

CONQUEST Quality Standards: For the Collaboration on Quality Improvement Initiative for Achieving Excellence in Standards of COPD Care

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Background: Chronic obstructive pulmonary disease (COPD) are managed predominantly in primary care. However, key opportunities to optimize treatment are often not realized due to unrecognized disease and delayed implementation of appropriate interventions for both diagnosed and undiagnosed individuals. The Collaboration on Quality improvement initiative for achieving Excellence in Standards of COPD care (CONQUEST) is the first-of-its-kind, collaborative, interventional COPD registry. It comprises an integrated quality improvement program focusing on patients (diagnosed and undiagnosed) at a modifiable and higher risk of COPD exacerbations. The first step in CONQUEST was the development of quality standards (QS). The QS will be imbedded in routine primary and secondary care, and are designed to drive patient-centered, targeted, risk-based assessment and management optimization. Our aim is to provide an overview of the CONQUEST QS, including how they were developed, as well as the rationale for, and evidence to support, their inclusion in healthcare systems.

Methods: The QS were developed (between November 2019 and December 2020) by the CONQUEST Global Steering Committee, including 11 internationally recognized experts with a specialty and research focus in COPD. The process included an extensive literature review, generation of QS draft wording, three iterative rounds of review, and consensus.

Results: Four QS were developed: 1) identification of COPD target population, 2) assessment of disease and quantification of future risk, 3) non-pharmacological and pharmacological intervention, and 4) appropriate follow-up. Each QS is followed by a rationale statement and a summary of current guidelines and research evidence relating to the standard and its components.

Conclusion: The CONQUEST QS represent an important step in our aim to improve care for patients with COPD in primary and secondary care. They will help to transform the patient journey, by encouraging early intervention to identify, assess, optimally manage and followup COPD patients with modifiable high risk of future exacerbations.

Keywords: identification, assessment, intervention, follow-up

Plain Language Summary

Under-diagnosis and under-treatment of COPD results in significantly higher risk of exacerbations, morbidity and death. Early identification and appropriate management should mitigate that risk. The Collaboration on Quality improvement initiative for achieving Excellence in Standards of COPD care (CONQUEST) aims to improve the management of patients at greater risk of future COPD exacerbations by developing and implementing

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quality standards (QS) into routine care and measuring implementation success. This is an area where important improvements in patient care and outcomes can be made and measured.

Four QS were developed: 1) identification of COPD target population, 2) assessment of disease and quantification of future risk, 3) appropriate non-pharmacological/pharmacological intervention, and 4) appropriate follow-up. The target population are patients with COPD (or potential COPD) with ≥ 2 moderate, or ≥ 1 severe exacerbations in the last 24 months (≥ 1 of these in the last 12 months) AND with scope for management optimization. Each QS is followed by a rationale statement and summary of current guidelines/evidence to support it and its components.

These QS represent a first step in empowering primary and secondary care providers to identify, assess, quantify risk, treat, and follow-up the millions of people living with COPD (both diagnosed and undiagnosed) at increased risk of exacerbations, morbidity and mortality. By incorporating these QS into routine care, CONQUEST aims to affect sustained change to COPD patient care. The QS will be used as a benchmark to monitor the impact of quality improvement on COPD outcomes in two planned CRTs in the UK and the USA.

Introduction

Despite being a largely preventable disease, the global burden of chronic obstructive pulmonary disease (COPD) is high and remains a public health priority in the 21st century.¹ COPD affects approximately 384 million people worldwide and is the third leading cause of death.^{2,3} It is frequently under-diagnosed, under-treated, and has high morbidity, including a greater risk of cardiovascular (CV) disease and death.^{3–5} COPD is projected to be responsible for a loss of 45 million quality-adjusted life years, 315 million exacerbations, and 9 million deaths cumulatively in the USA over the next 20 years, assuming current patterns of treatment and smoking rates.⁶ The socio-economic burden of COPD is also high, with projected 20-year (2019–2038) discounted direct and indirect costs estimated at \$801 billion and \$101 billion, respectively, in the USA.⁶ Similarly, the average total annual per patient cost of COPD management (excluding medications) in the United Kingdom has been estimated at £3396 for those experiencing ≥ 2 moderate-to-severe exacerbations. Primary care associated costs contribute most to overall expenditure.⁷

COPD is initially managed predominantly in primary care, but key opportunities to optimize management are often not realized due to unrecognized disease and delayed implementation of appropriate interventions for both diagnosed and undiagnosed individuals.^{8,9} A study carried out in the UK found that opportunities for diagnosis were missed in 85% of the patients in the 5 years immediately preceding COPD diagnosis.⁸ As a consequence, many patients receive no treatment or delayed initiation of maintenance therapy, even after diagnosis.⁹ This is important since interventions such as smoking cessation and pulmonary rehabilitation^{10,11} may improve prognosis and drugs such as inhaled corticosteroids (ICS) may improve current symptoms and exacerbations that are associated with rapid lung function loss.¹² Furthermore, patients who experience a greater exacerbation burden after initiation of maintenance therapy have worse lung function at diagnosis and a more rapid lung function decline thereafter, emphasizing the need for better treatment strategies.¹³ These findings highlight the importance of earlier intervention among patients with COPD, to more proactively diagnose and manage optimally. Current gaps in COPD management and the need for improved quality of care have been recognized both in the UK and the USA.^{14–18} In the USA, specific

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goals of the COPD national action plan¹⁴ are to raise public awareness of COPD, educate key stakeholders in diagnosis, prevention, and treatment, and encourage data sharing/dissemination. In the UK, the Department of Health has similar goals for COPD management.^{16,18}

The COLlaboratioN on QUality improvement initiative for achieving Excellence in STandards of COPD care (CONQUEST) is a new program providing a roadmap to improved quality of COPD care,¹⁹ and aligns with the aims of both the COPD National Action Plan and the UK Department of Health. CONQUEST is the first-of-its-kind collaborative, interventional, COPD registry with an integrated quality improvement program. Its vision is to drive change in the management of patients (both diagnosed and undiagnosed) at a greater risk of future COPD exacerbations and to measure the success of implementing this change in cluster randomized trials. This vision will be achieved by closing gaps in current COPD care provision, imbedding quality standards (QS) in routine primary care, integrating information from these patients, and presenting it to clinicians in a way that drives patient-centered, targeted, risk-based assessment and treatment optimization, thus empowering both patients and physicians to improve COPD care.¹⁹ The registry will be populated from local and global data sources and will hold anonymized data from over 200,000 primary and secondary care COPD patients (both diagnosed and undiagnosed) at modifiable high risk for COPD exacerbations. Patients will be identified through, and data collected from, electronic medical records (EMR). These data will be further supplemented through patient questionnaires and enhanced clinical assessments.¹⁹

The first step in the CONQUEST program is the development of QS. The aim of this article is to provide an overview of the CONQUEST QS, including how they were developed, as well as the rationale for, and evidence to support, their inclusion.

Methods

Quality Standard Design

These QS are core research and evidence-based statements that underpin the CONQUEST initiative and are intended to cover care for a targeted COPD population (see CONQUEST target patients below), a large proportion of whom may be positively impacted by quality COPD care. When developing these QS, consideration was given to 1) practicality and ease of use by all stakeholders (eg, clinicians, policy makers, advocacy groups), 2) future-proofing the QS, by advocating

general principles rather than a stringent and didactic approach, 3) feasibility of implementation in routine clinical practice, 4) the need to compare QS adherence and impact on disease outcomes across healthcare systems and 5) inter-country differences in COPD burden (morbidity, mortality, socio-economic), and patterns of treatment and access to care. Ethical approval from an Independent Review Board was not required as this article outlines QS informed by review of evidence in the public domain, no participants were recruited and data were neither collected nor analyzed.

The CONQUEST Target Patients

Two CONQUEST groups have been identified: those with a COPD diagnosis at baseline and those without a COPD diagnosis who fit the modifiable high-risk criteria. Please refer to [Table 1](#) for a summary of CONQUEST key terms. Modifiable high-risk patients are those with COPD (or potential COPD) who have had ≥ 2 moderate or ≥ 1 severe (hospitalized) exacerbation in the last 24 months (with at least one exacerbation occurring in the last 12 months) AND whose medical record data clearly indicates scope for management optimization. Management optimization focuses on patients who continue to have exacerbations while on their current therapy (including no therapy or short-acting β_2 -agonist only) and/or whose management may be optimized by correct diagnosis or additional non-pharmacological interventions. COPD exacerbations are defined as a significant worsening in respiratory symptoms in people with COPD (or an event analogous to a COPD exacerbation in people with suspected but undiagnosed COPD) and are categorized as moderate or severe. CONQUEST patients are required to be ≥ 40 years of age and, for those with potential COPD, to have a current or previous smoking history (ie, current or ex-smokers with ≥ 10 pack-year history or ≥ 10 years smoking duration).

CONQUEST Task Force

The CONQUEST QS task force comprises the Global Steering Committee, including 11 internationally recognized experts with a specialty and research focus in COPD from 8 countries across 4 continents, as well as 2 AstraZeneca members ([Table 2](#)). Membership of the task force was weighted towards the UK and the USA as it is in these countries where the QS will be initially implemented and trialed, before rolling out to other countries. Task force members are all experts in the field of COPD. Criteria for task force membership included 1) publication history; 2) leadership of national or international initiatives that have

Table I Lexicon of CONQUEST Key Terms

Term	Definition
CONQUEST Quality Improvement Program	Full name: Collaboration on a Quality Improvement Initiative to Achieve Excellence in Standards of COPD Care. The purpose of the CONQUEST program is to improve the prompt identification of patients with modifiable high-risk COPD, and to support the adoption of guideline-led clinical decision-making for all COPD patients, with a focus on assessment, therapy, and follow-up of patients with modifiable high-risk COPD. The components of the program form the intervention evaluated in PREVAIL.
PREVAIL Studies	
PREVAIL Cluster Randomized Controlled Trial	PRagmatic EVAluation of a quality Improvement program for people Living with modifiable high-risk COPD. The impact of the CONQUEST program on COPD outcomes will be evaluated in the subset of patients with modifiable high-risk COPD by a cluster-randomized controlled trial (CRT); self-contained primary care sites (PCS) or general practitioner (GP) practices will be the cluster units of randomization.
Baseline Period	The 24 months prior to randomization where primary care site (PCS) electronic medical record (EMR) data will be retrospectively searched to identify patients meeting modifiable high-risk criteria. The term “trial baseline” is used to describe the end of this 24-month-period, immediately prior to randomization.
Recruitment period	The period over which PCSs are recruited, electronic medical record (EMR) data are extracted, and sites are randomized to the CONQUEST intervention or control arm.
Implementation period	The intervention implementation period (start of the CONQUEST program) follows the recruitment period and runs for 6 months during which time the CONQUEST Quality Standards are implemented in intervention arm PCSs. This includes invitation of modifiable high-risk cohorts for assessment and clinical decision support prompted consultations to initiate optimization of COPD management. Control arm PCS continue with standard care.
Outcome Evaluation Period	Intervention arm site: The outcome evaluation period starts at the end of the 6-month CONQUEST implementation period in the intervention arm. During the outcome evaluation period control arm practices will continue to administer usual care for COPD. Control arm sites: The outcome evaluation period will start 6 months after recruitment of a PCS into the control arm. During the outcome evaluation period control arm practices will continue to administer usual care for COPD. The outcome evaluation period continues until the roll-out of CONQUEST to control arm PCSs.
Roll out	Roll-out of the CONQUEST intervention to the control arm practices commences at the start of the final year (year 5). Control arm roll-out mirrors the intervention arm implementation period; PCSs invite high-risk cohorts of patients for assessment and clinical decision support prompted consultations to initiate optimization of COPD management.
OPC affiliated sites	Those healthcare sites participating in the PREVAIL CRT or implementing the CONQUEST QI Program with the direct support of OPC.
Independent sites	Healthcare systems or sites wishing to implement the CONQUEST QI program (in accordance with the Global QI Operational Protocol) who are not supported by OPC directly. Independent sites will have access to resources and templates related to the program via the Global Operational Protocol and the CONQUEST website.

(Continued)

Table 1 (Continued).

Term	Definition
CONQUEST Target patients	
Modifiable “High-risk” patient	<p>Patients with COPD (or potential COPD) who have had 2 or more moderate, or 1 or more severe exacerbations in the last 24 months (with at least one exacerbation occurring in the last 12 months), AND whose medical record data indicates clearly that there is scope for management optimization.</p> <p>Patients with frequent exacerbations are at higher future risk of exacerbations. Frequent exacerbations are linked to accelerated lung function decline, greater risk of cardiovascular events and death. Guidelines state that frequent exacerbators should have their treatment optimized to reduce the risk of future exacerbations and potentially other adverse events. CONQUEST aims to address the management needs of such patients by promoting improved treatment and follow-up.</p>
Undiagnosed patient with potential modifiable high-risk COPD	<p>Patients without a COPD diagnosis who fit the modifiable high-risk criteria with respect to recent exacerbations of potential COPD, and who are smokers or former smokers ≥ 40 years of age.</p> <p>Exclusion: those with an active asthma diagnosis (ie, with an EMR diagnostic code for asthma and evidence of an asthma consultation in the 2 years before randomization (PREVAIL) or before identification (CONQUEST QI program)).</p> <p>N.B. 50 years of age will be used as the cut-off in the CRT in order to increase probability of identifying undiagnosed COPD patients.</p>
Newly diagnosed modifiable high-risk COPD patients	<p>The subset of the above undiagnosed patients who receive a COPD diagnosis following diagnostic assessment in the most recent CONQUEST QIP cycle.</p> <p>OR</p> <p>In the PREVAIL CRT, the subset of the above undiagnosed patients who receive a COPD diagnosis following assessment during the implementation and outcome evaluation period.</p>
Modifiable high-risk COPD diagnosed patients	<p>Patients with a COPD diagnosis at baseline who fit the modifiable high-risk criteria with respect to recent COPD exacerbations and are ≥ 40 years of age, and in whom there is an opportunity to optimize management.</p>
Patients with scope for management optimization in CONQUEST QI program	<p>The intervention will focus on patients who continue to have exacerbations while on their current therapy (including no therapy SABA, SAMA or SABA/SAMA only) and/or whose management may be optimized by correct diagnosis, or additional non-pharmacological interventions such as smoking cessation interventions or pulmonary rehabilitation.</p>
Exacerbations	
COPD exacerbation (EMR database definition)	<p>A significant worsening in respiratory symptoms in people with COPD.</p> <p>Either a moderate exacerbation defined as requiring an acute course of systemic corticosteroids and/or a course of antibiotics within 3 days of a lower respiratory consultation indicative of a probable COPD exacerbation, or an emergency room visit for a COPD-related respiratory cause*; or a severe exacerbation – ie an exacerbation resulting in a hospital admission or death from a COPD-related respiratory cause*.¹¹⁷</p> <p>Exacerbations occurring a minimum of 7 days after treatment of the initial exacerbation has ended will be considered as separate exacerbations. Prescribing instructions held in the EMR record will be used to identify the end of an exacerbation (ie, using recorded information on the duration of the treatment).</p>

(Continued)

Table 1 (Continued).

Term	Definition
Exacerbation of potential COPD (EMR database definition)	A significant worsening in respiratory symptoms analogous to a COPD exacerbation in people with suspected but undiagnosed COPD. Either a moderate potential exacerbation defined as requiring an acute course of systemic corticosteroids and/or a course of antibiotics within 3 days of a lower respiratory infection consultation, or an emergency room visit for a potentially COPD-related respiratory cause [†] ; or a severe exacerbation – ie an exacerbation resulting in a hospital admission or death for a potentially COPD-related respiratory cause [†] . Exacerbations occurring a minimum of 7 days after treatment of the initial exacerbation has ended will be considered as separate exacerbations. Prescribing instructions held in the EMR record will be used to identify the end of an exacerbation (ie, using recorded information on the duration of the treatment).
Complicated exacerbation	A severe COPD exacerbation OR A moderate COPD exacerbation involving any of the following: Additional acute course(s) of corticosteroids and/or respiratory antibiotic between 8 and 28 days after the initial event. Further record of moderate/severe COPD exacerbation between 8 and 28 days after the start of the initial event.
Other	
Systemic steroid burden	Total annual corticosteroid exposure from oral tablets, intravenous injection, or intramuscular injection measured as the average annual dose of prednisolone taken via any route, in milligrams. Records of systemic steroids other than prednisolone will be converted to prednisolone equivalent doses using USA and UK national formulary conversion tables. ^{118,119}
MACRE Major adverse cardiac or respiratory event	Occurrence of any of the below events: New diagnosis for heart failure, Hospitalization for heart failure, Revascularization, Myocardial Infarction, Stroke, All-cause mortality (further categorized as sudden death, cardiac deaths, respiratory death, none of these), Hospitalization or hospital admittance for respiratory event*, or Complicated exacerbations.
Collation of PRO/PRI questionnaire data	In patients receiving the QIP, PRO/PRI instruments will be completed at least annually, bringing together data received during the implementation of the intervention to be used in annual data analysis and feedback.
Opportunity Analysis	Retrospective study using EMR data to assess the management of patients with modifiable high-risk COPD over time in primary care and describe opportunities for management optimization in line with the Quality Standards. An opportunity analysis will be conducted in both the USA and the UK.

Notes: *Identified using validated code lists for moderate and severe COPD exacerbations. [†]Algorithm developed to define moderate and severe potential COPD exacerbations using appropriate respiratory diagnostic, symptom and investigation codes.

Abbreviations: CONQUEST, COllaboratioN on a QUality improvement initiative to achieve Excellence in STandards of COPD care; COPD, chronic obstructive pulmonary disease; EMR, electronic medical record; CRT, cluster randomized controlled trial; GOLD, Global initiative for chronic Obstructive Lung Disease; GP, general practitioner; ICS, Inhaled Corticosteroids; LABA, long-acting β_2 -agonist; LAMA, long-acting muscarinic antagonist; MACRE, Major Adverse Cardiac or Respiratory Event; OPC, Optimum Patient Care; PCS, primary care site; PREVAIL; PRagmatic EVAluation of a quality Improvement program for people Living with modifiable high-risk COPD; PRO/PRI, patient-reported outcome/patient-reported information; QIP, Quality Improvement Program; SABA, short-acting β_2 -agonist, SAMA, short-acting muscarinic antagonist.

Table 2 Members of the CONQUEST Quality Standard Task Force

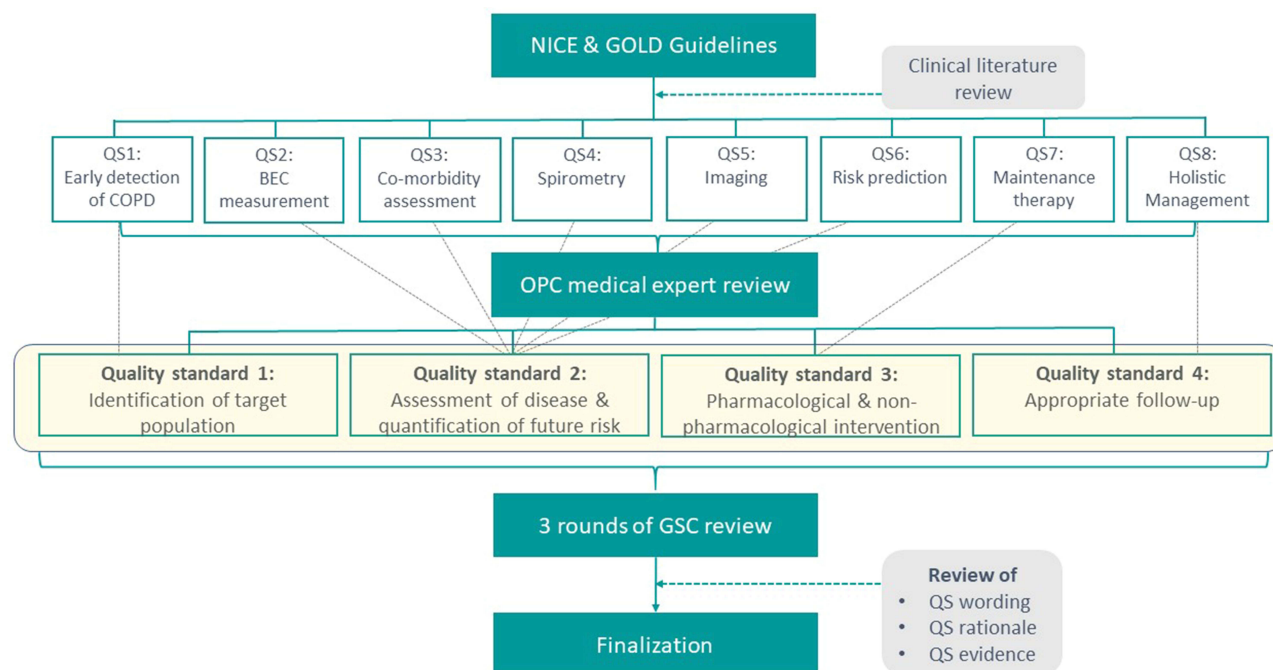
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Professor Rongchang Chen (China)	Director of Guangzhou Institute of Respiratory Diseases
Dr Mark Dransfield (USA)	Associate Professor of Medicine, Division of Pulmonary, Allergy and Critical Care Medicine
Professor John Hurst (UK)	Professor of Respiratory Medicine, University College London
Dr Fernando Martinez (USA)	Chief of Division of Pulmonary and Critical Care (GOLD committee)
Dr Marc Miravittles (Spain)	Pulmonologist, clinical research, Hospital Universitari Vall D'Hebron, Barcelona
Professor Shigeo Muro (Japan)	Professor of Respiratory Medicine, Nara Medical University, Japan
Professor David Price (Singapore)	Observational and Pragmatic Research Institute, Singapore
Dr Anita Sharma (Australia)	GP with cardiovascular interest at Platinum Medical Centre, Chair Scientific Programme Committee IPCRG
Professor Dave Singh (UK)	Professor of Clinical Pharmacology & Respiratory medicine, University of Manchester (GOLD committee)
Tonya Winders (USA)	President and Chief Executive Officer of Allergy & Asthma Network
Hana Müllerová	Medical Evidence Lead for COPD, AstraZeneca
Chris Blango	Employee of AstraZeneca at the time of the manuscript development

Abbreviations: IPCRG, International Primary Care Respiratory Group; GOLD, Global initiative for chronic Obstructive Lung Disease.

focused on high-risk COPD patients; or 3) experience as a primary care medical provider, patient representative or with cardiology expertise. Their role is to provide scientific advice on CONQUEST research projects and quality improvement program implementation, and to ensure that CONQUEST research is ethical, clinically appropriate, and continues to bring value to patients and physicians.

Quality Standard Development

The process included an extensive literature review as well as discussions with, and consensus of, experts of the CONQUEST Global Steering Committee and is summarized in Figure 1. Eight QS were initially identified, predominantly based on the National Institute for Health and Care Excellence (NICE) and Global initiative for chronic

**Figure 1** CONQUEST quality standards development process.

Abbreviations: BEC, blood eosinophil count; CONQUEST, COllaborationN on QUality improvement initiative for achieving Excellence in STandards of COPD care; COPD, chronic obstructive pulmonary disease; GOLD, Global initiative for chronic Obstructive Lung Disease; GSC, Global Steering Committee; NICE, National Institute for Health and Clinical Excellence; OPC, optimum patient care; QS, quality standard.

Obstructive Lung Disease (GOLD) recommendations, but also informed by the American Thoracic Society (ATS)/European Respiratory Society guidelines.^{3,20–22} Relevant references cited within these guidelines were also reviewed in addition to research identified through literature review.^{3,25–27} These eight QS were prioritized, consolidated (following several rounds of review by Optimum Patient Care medical experts) and sent to the CONQUEST QS Task Force for review and comment. The process consisted of three iterative rounds, and consensus was achieved following several meetings and via email correspondence. The process began in November 2019 and concluded in December 2020.

Global steering committee consensus was achieved for each of the four CONQUEST QS. Each QS is followed by a short statement explaining the rationale behind it and a summary of current guidelines and research evidence relating to the standard and its components.

Results

Quality Standard 1: Identification of Target Population

Identify individuals ≥ 40 years of age (with or without a pre-existing COPD diagnosis) with a history of smoking or relevant environmental exposure, at increased risk of exacerbations, morbidity, and mortality and with scope for COPD management optimization.

And, within this population, identify those with greater cardiovascular risk (Figure 2).

Rationale

The lives of COPD patients are significantly impacted by the disease. This informed the CONQUEST decision to focus our efforts on those individuals (both diagnosed and undiagnosed) at risk of exacerbating and to intervene earlier in the COPD journey in order to slow the speed of COPD progression (eg, accelerated lung function decline, increased exacerbation frequency, deterioration in co-morbidity control). A focus on patients at higher risk of exacerbating is an area in which large and important improvements in patient care and outcomes can be made and measured. These patients require more intense focus on identification and assessment in order to optimize management. The goal of optimized management is to reduce the number of exacerbations and consequent lung function decline and disease progression, thereby improving patient health-related quality of life (HRQoL) and decreasing CV risk associated with exacerbations.

Supporting Evidence

Evidence to support QS 1 is summarized in Table 3 and described in more detail here.

1. Diagnostic delay: COPD diagnosis is often delayed.^{8,23} Data from the Optimum Patient Care Research Database found that 85% of patients had consulted primary or secondary care for lower respiratory symptoms in the 5 years prior to their diagnosis, and 58% had consulted in the 10 years prior to diagnosis.⁸ Diagnosing COPD early (feasibly 5–10

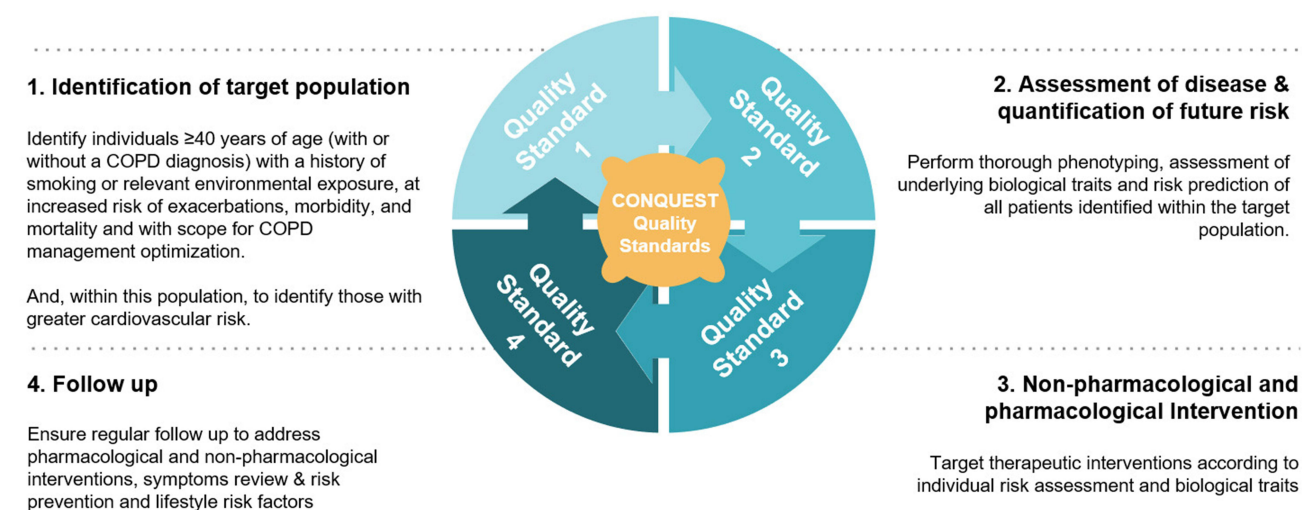


Figure 2 CONQUEST quality standards for the identification and management of patients with COPD at risk of exacerbations or acute respiratory events.

Abbreviations: CONQUEST, COLlaborationN on QUality improvement initiative for achieving Excellence in STandards of COPD care; COPD, chronic obstructive pulmonary disease.

years earlier based on UK data)⁸ in the disease progression is important, since undiagnosed patients have an increased risk of exacerbations and pneumonia, compared to their non-COPD counterparts.²⁴ Use of a targeted approach (focused case-finding methods) using characteristics such as age, smoking history, and respiratory symptoms may help to find these undiagnosed COPD patients.^{25,26} Physicians are more than seven times more likely to detect undiagnosed COPD in the targeted group (vs a routine care group), translating into one new case identified for every 21 targeted individuals assessed (after risk difference adjustment).²⁶ Undiagnosed patients who present to their health care provider (HCP) form part of the target population for CONQUEST, particularly those with a smoking history, who consult with lower respiratory symptoms, who received medication for lower respiratory illness or who were admitted to hospital or treated in an outpatient setting with respiratory symptoms.

2. Management optimization: COPD often remains undertreated, with a gap between guideline recommendations and real world practice.^{9,27,28} Medical and pharmacy claims data in the USA found that >65% of COPD patients were receiving no maintenance therapy,²⁷ whereas in the UK, three quarters of patients with a modified Medical Research Council (mMRC) dyspnea score <2 but ≥ 2 exacerbations/year were considered under-treated.⁹ Although patients hospitalized with severe exacerbations usually receive recommended care, once discharged, most patients receive inappropriate support and medications.²⁹
3. Exacerbation frequency: Many patients with COPD experience ≥ 2 exacerbations/year.³⁰ For example, 22% of patients with moderate COPD had frequent exacerbations in the ECLIPSE study.³¹ Furthermore, real-world data found that COPD patients who had ≥ 2 lower respiratory prescribing consultations in the 2 years preceding diagnosis were over three times more likely to have ≥ 2 exacerbations in the year after diagnosis.⁸ Earlier identification and treatment of patients who frequently exacerbate is likely to protect lung function, reduce CV risk and improve patients' HRQoL since 1) frequent exacerbations accelerate lung function decline,^{12,32} 2) exacerbations of COPD increase the risk of major adverse CV events,^{33–36} and 3)

frequency of exacerbation is associated with worse HRQoL.^{37,38} COPD patients with CV disease are four times more likely to have a subsequent CV event if they exacerbate, and 10 times more likely to have a subsequent CV disease (eg, cardiovascular death, myocardial infarction, stroke, unstable angina or transient ischemic attack) if the exacerbation required hospitalization.³⁶

Quality Standard 2: Assessment of Disease and Quantification of Future Risk

Perform thorough phenotyping, assessment of underlying biological traits and risk prediction of all patients identified within the target population (Figure 2).

Rationale

Different COPD phenotypes require different therapeutic interventions. Thorough assessment will enable accurate individual prediction of future risk of exacerbations, disease progression, mortality and CV risk, with the goal to guide therapy.

The assessment of individuals with COPD (both diagnosed and undiagnosed) with modifiable high risk of morbidity (including exacerbations) and mortality should include assessment of 1) symptoms and exacerbation history, 2) spirometry and other measures of lung function, 3) body mass index (BMI), 4) blood eosinophil count (BEC), 5) chest imaging, 6) cardiovascular risk assessment, 7) identification of co-morbidities, 8) use of relevant risk prediction tools and 9) smoking exposure and physical activity.

Supporting Evidence

Evidence to support QS 2 is summarized in Table 3 and described in more detail here.

1. Symptoms: NICE guidelines recommend the use of the MRC dyspnea scale for grading breathlessness at annual follow-up and when diagnosing COPD (in conjunction with other symptoms). The COPD Assessment Test is a method of assessing symptom burden in COPD that incorporates common symptoms.^{20,22} The results of symptom assessment measures can be used to guide pharmacological therapy in accordance with GOLD recommendations.³
2. Post-bronchodilator spirometry: Recommended in clinical guidelines to be used in conjunction with clinical assessment to confirm the diagnosis of COPD,^{3,22} and is incentivized as part of the

Quality Outcomes Framework in the UK. Longitudinal spirometry results can be used to measure lung function decline and response to therapy and can help guide therapy interventions.^{3,12}

3. Body mass index: BMI is associated with COPD prognosis,²⁰ and included in several risk prediction tools and models to determine exacerbation risk and/or mortality risk.^{39,40} Low BMI (ie, <21.75 kg/m²) has been associated with increased risk of pneumonia and mortality in COPD patients.^{41,42} Obesity, with BMI >30 kg/m², is a risk factor for the development of type II diabetes,⁴³ a relevant comorbidity for those who frequently exacerbate and require oral corticosteroids (OCS).
4. Blood eosinophil count: In stable COPD, BEC is recommended as a marker of steroid responsiveness, with higher BEC used as an indicator for ICS therapy.^{3,25,27} This recommendation is informed by the fact that those with higher BEC have less exacerbations when treated with ICS/long acting β_2 -agonist (LABA) versus LABA alone,^{44,45} or with ICS/LABA/long acting muscarinic agonist (LAMA) vs LAMA/LABA.^{46–48} Similarly, ICS are beneficial in reducing the number of exacerbations in those with elevated BEC, with exacerbation rate reductions increasing with increasing BEC.⁴⁹ Real-life data also showed that elevated BEC was associated with better outcomes for those on triple therapy (vs dual bronchodilator therapy),⁵⁰ and in frequent exacerbators is associated with more rapid lung function decline.¹² Higher BEC (ie, >200 cells/ μ L) may also be a predictor of OCS success in those experiencing exacerbations,⁵¹ and eosinophil guided therapy has been shown to reduce OCS exposure in patients admitted to hospital with acute exacerbation of COPD.⁵² Taken together, these results suggest the value of BEC to tailor treatments to specific COPD phenotypes and endotypes. However, BEC is commonly not available and/or underused in primary care and is not routinely considered in treatment decisions. More work on implementation in everyday clinical care is needed.
5. Imaging: Such as chest x-ray or CT scan is useful in suspected and/or confirmed COPD to exclude other respiratory pathologies.^{3,20} Chest CT scans are necessary when considering lung volume reduction,^{3,25} for the detection of bronchiectasis and for those requiring lung cancer assessment.³ CT chest scans (paired insp-exp) can aid early diagnosis of COPD by identifying air trapping, and standard CT scans can identify airway wall thickening, and can discriminate between emphysematous and non-emphysematous phenotypes.^{53,54} The former is a predictor for reduced survival as is pulmonary artery:aorta ratio.⁵⁵ However, it is recognized that CT scanning in all patients with COPD may not be feasible in the real-life setting.
6. CV risk assessment: CV diseases are perhaps the most important co-morbidities in COPD and carry an increased risk of death and hospitalization.^{5,56} COPD patients have a twofold higher risk of having CV disease (including ischemic heart disease, arrhythmias and heart failure) than their non-COPD counterparts.⁵⁷ Furthermore, cardiac mortality in patients with moderate COPD is higher than mortality due to respiratory failure,⁵⁸ and prognosis following an acute cardiac event is worse for those who also have a diagnosis of COPD.³⁵
7. Identification of co-morbidities: Co-morbidities besides CV disease (eg, anxiety/depression, gastroesophageal reflux disease, diabetes mellitus, and osteoporosis) are common in COPD and may be encountered at any level of disease severity.^{3,9,59} These co-morbidities are associated with a higher COPD symptom burden, higher likelihood of exacerbations and reduced therapy compliance.⁶⁰ Although the GOLD strategy document advises that, in general, the presence of co-morbidities should not change COPD treatment, these comorbidities can have a significant impact on disease course.³
8. Risk prediction tools: For example, the BODE index, which integrates BMI, airflow limitation (forced expiratory volume in one second (FEV₁)), dyspnea and 6-minute walk distance, are useful to predict COPD mortality.⁶¹ Another multicomponent index, DOSE (dyspnea, obstruction, smoking, exacerbations), has been shown to identify COPD patients with a greater risk of future worsening in health status and exacerbations.^{62,63} Others have found that risk of developing COPD in adulthood can be identified using lifetime lung function trajectory patterns and is 57% greater in those individuals with a parental history.^{64,65} These risk prediction tools may be used to guide therapy and highlight those most at risk of exacerbations, disease progression, and mortality within the target population.
9. Smoking (tobacco, e-cigarette/vaping), exposure and physical activity: The NICE guidelines suggest

a multifactorial approach to assess risk of exacerbations including ongoing smoking, exposure to pollutants, infection, insufficient physical activity, seasonal variation and frequency of exacerbations.²⁰ GOLD acknowledge that smoking cessation is key for COPD prevention and for maintenance therapy. However, the effectiveness of e-cigarettes as a smoking cessation aid is uncertain since e-cigarette use has been associated with severe acute lung injury, eosinophilic pneumonia, alveolar hemorrhage, respiratory bronchiolitis and other lung abnormalities.^{3,66,67}

Quality Standard 3: Non-Pharmacological and Pharmacological Intervention

Target therapeutic interventions according to individual risk assessment and biological traits (Figure 2).

Rationale

Both non-pharmacological and pharmacological interventions are essential for effective COPD management. These are used to reduce symptoms, reduce the frequency and severity of exacerbations, and improve exercise tolerance and health status.³ In accordance with published literature and expert opinion, the GOLD strategy document, and NICE and ATS guidelines, the CONQUEST program recommends the following^{3,22}:

Non-Pharmacological Interventions

1. Smoking cessation interventions as appropriate
2. Pulmonary rehabilitation referrals, where indicated
3. Long-term oxygen therapy, where indicated
4. Pneumococcal and annual influenza vaccination to all with COPD
5. Patient motivation and engagement enhanced by patient-reported outcome information and shared decision-making.

Pharmacological Interventions

1. Dual bronchodilators (LABA/LAMA) to all who are symptomatic on long-acting bronchodilator monotherapy.
2. Triple therapy (ICS/LABA/LAMA) for appropriate patients with exacerbations and elevated BEC, and potentially to those with greater CV disease risk and greater risk of mortality (ie, severe exacerbations).

a. In line with national guidelines for initial and follow-up therapy. Follow-up therapy should reflect both background therapy, symptom response and exacerbation risk based on actual exacerbations and COPD control.

3. Adequate and prompt pharmacological therapy of cardiac risk factors and disease according to latest guideline recommendations.
4. Regularly assess inhaler choice and technique.
5. Short courses of OCS and/or antibiotics during exacerbations as per GOLD or other national/international guideline recommendations.

Supporting Evidence

Evidence to support QS 3 is summarized in Table 3 and described in more detail here.

Non-Pharmacological Interventions

1. Smoking cessation: Has the greatest capacity to influence the natural history of COPD. The earlier smoking cessation is achieved the greater the impact on preventing lung function decline.⁶⁸ There is also an associated reduction in symptoms following smoking cessation.⁶⁹
2. Pulmonary rehabilitation: Improves dyspnea, health status and exercise tolerance in stable patients.^{3,10,11} Started soon after an exacerbation of COPD, pulmonary rehabilitation is safe, effective and associated with fewer subsequent hospital admissions for exacerbations.⁷⁰ It is recommended by NICE to all “who view themselves as functionally disabled by COPD (usually MRC of grade 3 or above)”,²⁵ and by GOLD as part of integrated COPD patient management.³
3. Long-term oxygen therapy: Has been shown to increase survival in patients with severe resting hypoxemia. GOLD recommend it for patients who have 1) $\text{PaO}_2 \leq 7.3$ kPa or $\text{SaO}_2 \leq 88\%$ with or without hypercapnia confirmed twice over a 3-week period or 2) for those with PaO_2 7.3–8.0 kPa or SaO_2 of 88% if there is evidence of pulmonary hypertension, peripheral edema suggesting congestive cardiac failure or polycythemia (hematocrit $>55\%$).³ Domiciliary non-invasive ventilation has proven utility in those with more severe disease.⁷¹
4. Vaccination: NICE and GOLD recommend offering pneumococcal and annual influenza vaccination to

all with COPD.^{3,20} More recently, people with more severe COPD have been prioritized for COVID-19 vaccine, and as these become available, local recommendations should be followed. Influenza vaccinations have been shown to reduce death and illness, such as lower respiratory tract infections, in COPD patients.^{72,73} An updated Cochrane review also concluded that inactivated influenza vaccine led to fewer COPD exacerbations, but noted that there were a limited number of randomized controlled trials available on the topic.⁷⁴ Pneumococcal vaccination also reduces the likelihood of COPD exacerbation.⁷⁵ While not currently part of any COPD guidelines, there has been some recent discussion regarding the role of pertussis booster vaccination in COPD patients, due to a resurgence of pertussis and evidence that COPD patients are at high risk of contracting the disease.⁷⁶

5. Patient motivation and engagement: GOLD recognizes the benefit of personalized, multi-component, education and training for patients with COPD (rather than simply providing information and advice).³ Shared decision-making and patient engagement have been shown to significantly improve perceived health status, COPD knowledge, adherence to pharmacological treatment, general functionality and healthy lifestyle measures for COPD patients hospitalized for acute exacerbations.⁷⁷

Pharmacological Interventions

The GOLD strategy document classifies COPD patients initiating therapy into ABCD groups based on health status as assessed by CAT, mMRC dyspnea score and exacerbation history, and recommends appropriate targeted pharmacological intervention for each group.³ The CONQUEST cohort are likely to fall within GOLD groups C or D. A dyspnea or exacerbation directed step up approach is advised if response to this initial treatment is not sufficient. Patients with dyspnea as their predominant symptom should be stepped up to dual LABA/LAMA (if on LABA or LAMA monotherapy) and consider switching inhaler device or investigation of other causes of dyspnea for those who remain symptomatic on dual bronchodilators.³ De-escalation of ICS should only be considered for those who become low risk over time or have reasons to be cautious of ICS use, including low eosinophil count and pneumonia.^{3,78} Exacerbating patients on LABA or LAMA mono-therapy should be stepped up to LABA/

LAMA or ICS/LABA (if $\text{BEC} \geq 300$ or $\text{BEC} \geq 100$ plus ≥ 2 exacerbations), and then to triple therapy if response is insufficient.³

1. Dual bronchodilators (to all who are symptomatic on long-acting bronchodilator monotherapy): Multiple studies have found improvements in lung function, symptoms and HRQoL with dual bronchodilators when compared to single therapy or placebo.⁷⁹
2. Triple therapy (ICS/LABA/LAMA) for appropriate patients with exacerbations and higher BEC, and potentially to those with greater CV disease risk and greater risk of mortality): Most beneficial to symptomatic COPD patients with an exacerbation history despite dual therapy. Also recommended for those patients with higher BEC (ie, ≥ 300 cells/ μL).^{80,81} COPD patients with a history of exacerbations treated with triple therapy experienced better HRQoL, better lung function and fewer moderate-to-severe exacerbations, compared to those treated with either LABA/LAMA or ICS/LABA; with 25% and 16% less exacerbations, respectively.⁸² ICS-containing regimens were associated with a lower rate of death from CV and respiratory causes and lower all-cause mortality.^{82,83} However, in keeping with other studies,^{84,85} an increased incidence of pneumonia was noted for those on ICS.⁸² For example, studies of salmeterol/fluticasone in COPD found that the number needed to treat to induce one case of pneumonia ranged from 16 to 20.⁸⁶ More recently, the ETHOS study also found that compared to those on dual therapy, patients on ICS triple therapy experienced a significantly lower annual exacerbation rate (budesonide/glycopyrronium/formoterol) vs LABA/LABA: (number needed to treat) (NNT)=3 and vs ICS/LABA: NNT=7],⁸⁷ with lower mortality rates (any cause) noted in the higher ICS group.⁴⁷ Pneumonia rates across triple and dual therapies were low, between 0.02 and 0.05 per patient-year and were lower for budesonide/glycopyrronium/formoterol than budesonide/formoterol (number needed to harm=58), suggesting a low pneumonia risk for budesonide/glycopyrronium/formoterol relative to its benefits on exacerbations.⁸⁷
3. Adequate and prompt pharmacological therapy of cardiac risk factors and disease: Although CV medications can cause concerns among prescribers over

possible side effects or interactions in COPD patients leading to under-treatment of CV disease,⁸⁸ the overall principle advocated by current COPD guidelines is that treatment of comorbidities should not be altered by the presence of COPD.³ The use of cardio-selective β_1 -blockers where indicated in ischemic heart disease and heart failure is considered safe with the benefits of treatment outweighing any potential risks.⁸⁹ The current GOLD report advises that heart failure should be treated no differently in COPD patients and use of cardio-selective β -blockers is recommended.³

4. Regularly assess inhaler choice and technique: While ensuring the right drug is prescribed to the patient at the correct dose is important for treatment success, method of delivery and inhaler training are often overlooked. Choice of inhaler should be tailored to the individual patient where possible and consideration given not only to what the inhaler contains (eg, drug class/particle size) but also inspiratory flow, ease of use, patient preference, age, cognition, manual dexterity and coordination.⁹⁰ Regular checking of inhaler technique is also important and has been associated with improved therapy adherence.⁹¹
5. Short course of OCS for exacerbations (when clinically indicated): GOLD strategy suggests consideration of a course of OCS of between 5 and 7 days for moderate or severe acute exacerbations.³ Use of OCS in these situations is associated with improved lung function, oxygenation and recovery time.⁹² Similarly for those with raised BEC, treatment with OCS during an acute exacerbation has been associated with a reduction in treatment failure and hospitalization.⁵¹ The NICE guidelines also recommend short-term OCS use (typically 5 days) in all people admitted to hospital with an exacerbation of COPD (if no contraindications) and to consider their use in community settings when an exacerbation is associated with “a significant increase in breathlessness that interferes with daily activities”.¹³ OCS can be associated with significant side effects in both the short and long term.⁹³

Quality Standard 4: Appropriate Follow-Up

Ensure regular follow-up to address non-pharmacological and pharmacological intervention by symptoms review and assessing risk prediction and lifestyle risk factors (Figure 2).

Rationale

Ongoing effort over time is required to change behavior and determine the appropriate course of therapy. Not all aspects of COPD care can be discussed and addressed in an initial consultation or risk assessment and therefore require a long-term patient treatment strategy.

Supporting Evidence

Evidence to support QS 4 is summarized in Table 3 and described in more detail here.

To improve quality of care for COPD patients, efforts to provide individualized treatment plans and interventions must be ongoing.³ This cycle of ongoing review and assessment involves pharmacological, clinical, and non-pharmacological components as outlined below:

1. Medication, inhaler device and symptom review: Current therapy should be recorded and reviewed with up-to-date biomarkers, spirometry, symptom severity scores and exacerbation rates. Pulse oximetry (at rest, during a 6-minute walk or overnight) can be used to estimate a patient’s arterial oxygen saturation and need for supplemental oxygen therapy, and is important in all patients with clinical signs suggestive of respiratory failure or right heart failure.³ Use of the COPD control tool may help to achieve appropriate and faster clinical review to evaluate response to therapy and escalate treatment where appropriate.^{94,95}
2. Inhaler technique: There is evidence that errors in inhaler technique are associated with worse disease outcomes in both asthma and COPD.⁹⁶ Conversely, significant improvement in inhaler technique has been associated with improved COPD control,⁹⁷ and checking of inhaler technique (within the last 2 years) is associated with better adherence.⁹¹
3. Smoking cessation: A Cochrane review in 2016 found evidence that smokers with COPD were more than twice as likely to quit if they received “high-intensity behavioral support” as well as medication, compared to those receiving only behavioral support.⁹⁸ There was no clear evidence that any one form of support or medication was better than another.
4. Nutrition: NICE COPD guidelines recommend calculation of BMI at the time of diagnosis and at follow-up, and referral for dietetic advice is recommended for those with low or high BMI.²⁰

5. Pulmonary rehabilitation: Patient needs and requirements can change over time, so the role of pulmonary rehabilitation should be part of regular patient follow-up. Patients should be referred if the therapy is likely to be beneficial. Considerations regarding patients suitable for referral include discussion of patient goals and their ability to perform the exercises, as well as noting the need to commit to a course of therapy lasting at least 6 to 8 weeks.^{3,99} The likely benefits of the program should be emphasized to the patient. The ability of clinicians to refer patients to a suitable pulmonary rehabilitation program may be limited by resources and availability.
6. Vaccination: Should be offered to COPD patients at review in accordance with local policies (please also see point 4 of QS 3 supporting evidence for non-pharmacological intervention).

Table 3 COPD Quality Standards and Rationale for Their Inclusion in CONQUEST

Quality Standard	Rationale to Include as a CONQUEST QS
1. Identification	<p>COPD diagnosis is often delayed in primary care.^{8,23}</p> <p>Use of a targeted approach on 'at risk' patients can help to find these undiagnosed COPD patients.^{25,26}</p> <p>COPD remains undertreated, with a gap between guideline recommendations and real world practice.^{9,27,28}</p> <p>COPD exacerbations are associated with accelerated lung function decline,^{12,32} and increased risk of major adverse CV events,^{33–36} and worse HRQoL.^{37,38}</p> <p>The socio-economic burden of COPD is high.^{6,7}</p>
2. Assessment of disease and risk quantification	<p>Symptom assessment using mMRC and or CAT score and exacerbation history to inform and guide therapy.³</p> <p>BEC is recommended as one marker of steroid responsiveness in stable COPD.^{3,25,27}</p> <p>CV disease in COPD patients carries an increased risk of death and hospitalization.^{5,56}</p> <p>Other co-morbidities (eg, anxiety/depression, GERD, DM, Osteoporosis) are associated with a higher COPD symptom burden, higher likelihood of exacerbations and reduced therapy compliance.⁶⁰</p>
3. Non-pharmacological and pharmacological interventions	<p>CONQUEST recommends:³</p> <p>Smoking cessation interventions as necessary.</p> <p>Pulmonary Rehabilitation referrals where indicated.</p> <p>Long-term oxygen therapy where indicated.</p> <p>Dual bronchodilators to all who are symptomatic despite being on single therapy.</p> <p>Triple therapy for appropriate patients with increased exacerbation risk, higher BEC, and potentially to those with greater CV disease risk.</p> <p>Adequate and prompt pharmacological therapy of cardiac risk factors and disease.</p> <p>Appropriate inhaler choice and checking of inhaler technique.</p> <p>Appropriate use of OCS and antibiotics to treat exacerbations.</p> <p>Pneumococcal and annual influenza vaccination to all with COPD and Covid-19 vaccination according to local recommendations.</p>
4. Follow-up	<p>GOLD recommends that current therapy should be recorded and reviewed with up-to-date biomarkers, post-bronchodilator spirometry, symptom severity scores and exacerbation rates.³</p> <p>Pulse oximetry used to evaluate the need for supplemental oxygen therapy.³</p> <p>Errors in inhaler technique are associated with worse disease outcomes.⁹⁶</p> <p>Smokers with COPD are > twice as likely to quit if they received 'high-intensity behavioral support'.⁹⁸</p> <p>NICE COPD guidelines recommend calculation of BMI at the time to diagnosis and at follow-up, and referral for dietetic advice is recommended for those with low or high BMI.²⁰</p> <p>Patient needs and requirements can change over time, so the role of PR should be part of regular patient follow-up.</p> <p>NICE and GOLD recommend pneumococcal and influenza vaccination to all with COPD.^{3,25}</p>

Abbreviations: BEC, blood eosinophil count; BMI, body mass index; CONQUEST, CollaborationN on Quality improvement initiative for achieving Excellence in Standards of COPD care; CAT, COPD Assessment Test; COPD, chronic obstructive pulmonary disease; CV, cardiovascular; DM, diabetes mellitus; GERD, gastroesophageal reflux disease; GOLD, Global initiative for chronic Obstructive Lung Disease; HRQoL, health-related quality of life; mMRC, modified Medical Research Council breathlessness score; NICE, National Institute for Health and Clinical Excellence; OCS, oral corticosteroid; PR, pulmonary rehabilitation.

7. Motivation and engagement: For quality improvement initiatives of this kind to be successful, there must be motivation to drive long-term behavioral changes, from both the physician/clinical team and the patient. In the past, therapeutic nihilism has been a feature of COPD care, but there is evidence that this feeling that little can be done for COPD patients is changing, with improving primary care physician knowledge and awareness of COPD guidelines.¹⁰⁰ Ensuring that motivation is sustained beyond the trial phase is a critical part of the CONQUEST initiative. Patients may not be motivated to seek medical help due to a lack of understanding of the disease itself, feelings of guilt and stigma associated with a COPD diagnosis,^{101,102} the feeling that it is “too late” to change or simply due to ignoring symptoms and/or attributing them to another cause.^{103,104} Greater public awareness and knowledge of the condition and the possibilities for its treatment are needed to counter these issues. Implementing behavior-targeted interventions with the aim of improving physical activity should also be considered as physical activity is often significantly reduced in COPD patients.¹⁰⁵ According to GOLD (2021),³ “This can lead to a downward spiral of inactivity, predisposing patients to reduced QoL, increased rates of hospitalization and mortality”.^{106,107}

Discussion

These CONQUEST QS represent a roadmap to improved quality of care for people with COPD (both diagnosed and undiagnosed). People with modifiable disease and higher risk of exacerbations, morbidity and mortality, are individuals for whom there is an opportunity to optimize management. The QS presented in the current article represent a first step in empowering primary and secondary care providers to identify, assess, quantify risk, treat, and follow-up the millions of people living with COPD who fall into this category, and differ from previous QS in that they are internationally agreed. They are important, as better identification of those at risk of exacerbating and those requiring management optimization is needed to reduce lung function decline and disease progression and improve patient HRQoL. Phenotyping disease using clinical, CV risk and biomarker variables should facilitate the identification of treatable traits, bringing us one step closer to the practice of precision medicine and selection of targeted phenotype-specific treatments to optimize COPD outcomes. Finally, regular follow-up of COPD patients is

critical to modify risks (with appropriate diagnosis and treatment), lock in behavior change(s), measure treatment successes, and encourage re-assessment of the patient (eg, identification of clinical, non-pharmacological and pharmacological interventions/issues, making adjustments to treatment when necessary). By engaging physicians and patients, and then incorporating and embedding a process of quality improvement into routine care, we aim to bring about sustained change to COPD patient care. The QS will be used as benchmark to monitor the impact of quality improvement in COPD management.

Although, many of the components of our initiative may be intuitive, data showing that it can be implemented successfully, in a way which can be scaled up globally, and improve disease outcomes is less apparent. Other initiatives to improve quality of care and guideline adherence in COPD management have shown some success.^{17,108–115} Specific quality improvement initiatives have included financial incentives,¹⁰⁸ national clinical audits,¹⁷ COPD diagnosis questionnaires,¹⁰⁹ educational programs for patients and physicians,^{111,114} introduction of care bundles to COPD management¹¹² and utilization of EMR for clinical decision-making support.¹¹⁰ For example, introduction of a COPD flow sheet into EMR at a tertiary care clinic resulted in significant improvement of clinical assessment and use of LABA.¹¹⁰ Use of a web-based disease management system to provide point-of-care support and information on primary care COPD management in the USA led to a significant reduction in exacerbations and improvement in COPD control medication prescription.¹¹¹ In the UK, a national clinical audit for hospitalized exacerbations of COPD found improvements in process indicators when continuous audit was combined with quality improvement support and, in particular, financial incentives.¹⁷ These studies contain elements that are similar to planned components of CONQUEST and suggest that providing information and guidance at the point-of-care can lead to closer adherence to clinical guidelines. However, overall, results have been mixed and a reduction in mortality has not yet been shown. This may have been because of the short-term nature of some of these initiatives, their tendency to focus on severe disease, and failure to fully integrate into health-care systems. Patient-reported outcomes and information were also under-utilized and quality improvement methodologies were of mixed quality and heterogeneous.¹¹⁶

CONQUEST overcomes many of these limitations. It promotes more proactive and hence earlier diagnosis, since undiagnosed COPD patients have an increased risk of

exacerbations and pneumonia, compared to those without COPD. Targeting undiagnosed patients at risk (≥ 40 years old and ever smokers) including the presence of previous respiratory symptoms and exacerbations, should increase diagnostic yield, improve disease outcomes, reduce health-care resource utilization, and help to inform national/international guidelines and policies to support earlier prevention and treatment intervention in COPD.

Limitations to CONQUEST QS implementation include the fact that global and local quality improvement protocols for implementation may need to be modified depending on the country adopting these standards. Any modifications required would be based on the various nuances of healthcare policy and differences in national recommendations and guidelines, which may impact a global delivery of the program. Moreover, variability in availability of technology and/or access to electronic record systems for patient identification (especially in lower income countries) may also impact ability to implement CONQUEST QS. As the QS will be implemented, and their impact assessed in the UK and the USA initially, we acknowledge that these countries may not be representative of the situation elsewhere. Challenges encountered in each country may be different (eg, data protection issues, completeness of EMRs and differentiation between primary and secondary care), particularly with respect to socio-economic burden of lower income countries, and also differences in access to, and availability of, pharmacological treatments. On the strengths side of the equation, CONQUEST QS have been informed by 11 key thought leaders from a variety of backgrounds, including representation from primary and specialist care, academics, patient advocacy groups and the pharmaceutical industry. The QS will help to address potential disconnects between local/national and global approaches, are consensus and evidence-based, will be reviewed and updated as new evidence emerges or whenever necessary, and will be embedded into primary and secondary care processes.

Next steps for CONQUEST include the translation of these QS into Quality Improvement Program operational protocols, which describe the core components of the program required to implement CONQUEST in a healthcare system or practice. The USA and the UK have been identified for the initial delivery of the intervention, as these countries have a well-documented burden of COPD, existing quality improvement programs, and EMR integration into healthcare systems. An Opportunity Analysis using retrospective analysis of EMRs to compare

the QS with current practice in each of these countries will be undertaken to describe the opportunities for management optimization. The impact of the CONQUEST intervention on COPD outcomes (eg, rate of exacerbations and major adverse cardiac or respiratory events, COPD control, lung function, systemic corticosteroid burden) will be evaluated by a cluster randomized controlled trial (CRT) in each country (PREVAIL study).

Conclusions

The CONQUEST QS are the first step on an international journey to improve care of the COPD patient in primary and secondary care. They aim to transform the patient pathway, by identifying patients with modifiable high risk of future exacerbations, optimizing management and follow-up of patients with diagnosed COPD to reduce symptoms, exacerbations, co-morbidity, and mortality. Moving forward CONQUEST will advocate for appropriate and early treatments to slow disease progression, optimize patient outcomes including improved HRQoL, encourage follow-up with patients and HCPs utilizing shared decision-making to lock in behavior change, and generate transformational evidence to measure treatment success.

Abbreviations

ATS, American Thoracic Society; BEC, blood eosinophil count; BMI, body mass index; BODE, Body-mass index, airflow Obstruction, Dyspnea, and Exercise; CONQUEST, CollaborationN on Quality improvement initiative for achieving Excellence STandards of COPD care; COPD, chronic obstructive pulmonary disease; CV, cardiovascular; DOSE, dyspnea, obstruction, smoking, exacerbations; ECLIPSE, Evaluation of COPD Longitudinally to Identify Predictive Surrogate Endpoints; EMR, electronic medical record; ETHOS, Efficacy and Safety of Triple Therapy in Obstructive Lung Disease; FEV₁, forced expiratory volume in one second; GOLD, Global initiative for chronic Obstructive Lung Disease; HCP, health care providers; HRQoL, health-related quality of life; ICS, inhaled corticosteroids; LABA, long-acting β_2 -agonist; LAMA, long-acting muscarinic antagonist; mMRC, modified Medical Research Council; NICE, National Institute for Health and Care Excellence; OCS, oral corticosteroids; QS, quality standards.

Acknowledgments

We thank Dr Seyi Soremekun, Jonathan Marshall, Jennie Medin and Irena Brookes-Smith for their valuable

contributions to the design of the study. We would also like to acknowledge Ms Andrea Teh Xin Yi (BSc, Hons) of the Observational and Pragmatic Research Institute (OPRI), Singapore, for editorial and formatting assistance which supported the development of this publication. Professor Dave Singh is supported by the National Institute for Health Research (NIHR) Manchester Biomedical Research Centre (BRC).

Funding

CONQUEST is conducted by Optimum Patient Care Global and Observational and Pragmatic Research Institute and is co-funded by Optimum Patient Care Global and AstraZeneca.

Disclosure

Rachel Pullen and Amy Couper are employees of the Observational and Pragmatic Research Institute, which is a research collaborator of the CONQUEST initiative with Optimum Patient Care and AstraZeneca. Marc Miravittles reports speaker fees from AstraZeneca, Boehringer Ingelheim, Chiesi, Cipla, Menarini, Rovi, Bial, Sandoz, Zambon, CSL Behring, Grifols and Novartis, consulting fees from AstraZeneca, Boehringer Ingelheim, Chiesi, GlaxoSmithKline, Bial, Gebro Pharma, Kamada, CSL Behring, Laboratorios Esteve, Ferrer, Mereo Biopharma, Verona Pharma, TEVA, Spin Therapeutics, pH Pharma, Novartis, Sanofi and Grifols and research grants from GlaxoSmithKline and Grifols. Anita Sharma is a practising Primary Care Physician and Senior Lecturer, School of Clinical Medicine-Primary Care Clinical Unit, University of Queensland. She supervises clinical training of primary care doctors and serves on advisory boards for Diabetes, Heart Failure and Osteoporosis for Novartis, Merck Sharp & Dohme and Boehringer Ingelheim, Eli Lilly and Amgen. Dave Singh has received personal fees from Aerogen, AstraZeneca, Boehringer Ingelheim, Chiesi, Cipla, CSL Behring, Epiendo, Glenmark, Gossamerbio, Kinaset, Menarini, Novartis, PulmatrixSanofi, Theravance and Verona. Fernando Martinez has received personal fees and non-financial support from the American College of Chest Physicians, AstraZeneca, Boehringer Ingelheim, ConCert, Genentech, GSK, Inova Fairfax Health System, Miller Communications, National Society for Continuing Education, Novartis, Pearl Pharmaceuticals, PeerView Communications, Prime Communications, Puerto Rico Respiratory Society, Chiesi, Sunovion, Theravance,

Potomac, University of Alabama Birmingham, Physicians Education Resource, Canadian Respiratory Network, Teva and Dartmouth; non-financial support from ProterrixBio, Gilead, Nitto and Zambon; and personal fees from Columbia University, Integritas, MD magazine, Methodist Hospital Brooklyn, New York University, UpToDate, WebMD/MedScape, Western Connecticut Health Network, Patara/Respivant, PlatformIQ, American Thoracic Society, Rockpointe, Rare Disease Healthcare Communications and France Foundation; grant support from NIH; and is a member of steering committees for Afferent/Merck, Biogen, Veracyte, Prometic, Bayer, Bridge Biotherapeutics and ProMedior. John Hurst has received personal payment and payment to his institution (UCL), including research grants, reimbursement for advisory work and educational activities, and support to attend meetings from pharmaceutical companies that make medicines to treat COPD, which includes AstraZeneca, Boehringer Ingelheim, Chiesi and Novartis. ; Luis Alves has served as an advisor or consultant for AstraZeneca, GlaxoSmithKline; served as a speaker or a member of a speakers bureau for Astra-Zeneca, BIAL, GlaxoSmithKline, and Novartis Pharmaceuticals Corporation. Luis Alves also reports personal fees from Optimum Patient Care Global Limited, during the conduct of the study. He is also a member of the Education Subcommittee of the International Primary Care Respiratory Group and a member of the of GRESP, the Portuguese Primary Care Study Group for Respiratory Diseases. Mark Dransfield has received grant support from the Department of Defense and NIH; personal fees from AstraZeneca, Boehringer Ingelheim, PneumRx/BTG, Genentech, BostonScientific, Quark Pharmaceuticals, Teva, and GSK; and contracted clinical trial support from Boehringer Ingelheim, Novartis, AstraZeneca, Yungjin, PneumRx/BTG, Pulmonx, Boston Scientific and GSK. Rongchang Chen has received grant support from AstraZeneca, Boehringer Ingelheim, and GSK; and reimbursement for advisory work and educational activities from AstraZeneca, Boehringer Ingelheim, GSK, Novartis. Shigeo Muro reported lecture and advisory fees from Novartis Pharma and AstraZeneca; lecture fees and grants from Boehringer Ingelheim and Fukuda Life Tech; advisory fees from GlaxoSmithKline; grants from Eisai Pharmaceutical, Otsuka Pharmaceutical and Fuji Film Medical; and lecture fees from Astellas Pharmaceutical, Kyorin Pharmaceutical and Meiji Seika Pharma. Tonya Winders has received personal fees from AstraZeneca,

GSK, Novartis, Genentech, and Sanofi/Regeneron alliance. Allergy & Asthma Network & Global Allergy & Airways Patient Platform has received funds for unbranded disease awareness and education from AZ, BI, GSK, Novartis, Genentech, Viatris and Sanofi/Regeneron. Christopher Blango was an employee of AstraZeneca and hold stock and/or stock options in the company. Currently employed at the Janssen Pharmaceutical Companies of J&J. Hana Müllerová, Tamsin Morris, Frank Trudo and Paul Dorinsky are employees of AstraZeneca and hold stock and/or stock options in the company. AstraZeneca is a co-funder of the CONQUEST initiative. Marianna Alacqua was an employee of AstraZeneca at the time of manuscript development. Currently employed at CSL Behring SpA. Victoria Carter is an employee of Optimum Patient Care, a co-funder of the CONQUEST initiative. Rupert Jones declares grants from Astra Zeneca, Glaxo Smith Kline, Novartis and Teva and personal fees for consultancy, speakers fees or travel support from Astra Zeneca, Boehringer Ingelheim, Glaxo Smith Kline, Novartis and OPRI. Konstantinos Kostikas was an employee and shareholder of Novartis Pharma AG until 31.10.2018. He has received honoraria for presentations and consultancy fees from AstraZeneca, Boehringer Ingelheim, Chiesi, ELPEN, GSK, Menarini, Novartis and Sanofi. His department has received funding and grants from AstraZeneca, Boehringer Ingelheim, Chiesi, Innovis, ELPEN, GSK, Menarini, Novartis and NuvoAir. He is a member of the GOLD Assembly. Ruth Murray reports personal fees from OPC, during the conduct of the study. David B Price has board membership with AstraZeneca, Boehringer Ingelheim, Chiesi, Mylan, Novartis, Regeneron Pharmaceuticals, Sanofi Genzyme, Thermofisher; consultancy agreements with Airway Vista Secretariat, AstraZeneca, Boehringer Ingelheim, Chiesi, EPG Communication Holdings Ltd, FIECON Ltd, Fieldwork International, GlaxoSmithKline, Mylan, Mundipharma, Novartis, OM Pharma SA, PeerVoice, Phadia AB, Spirosure Inc, Strategic North Limited, Synapse Research Management Partners S.L., Talos Health Solutions, Theravance and WebMD Global LLC; grants and unrestricted funding for investigator-initiated studies (conducted through Observational and Pragmatic Research Institute Pte Ltd) from AstraZeneca, Boehringer Ingelheim, Chiesi, Mylan, Novartis, Regeneron Pharmaceuticals, Respiratory Effectiveness Group, Sanofi Genzyme, Theravance and UK National Health Service; payment for lectures/speaking engagements from AstraZeneca, Boehringer Ingelheim,

Chiesi, Cipla, GlaxoSmithKline, Kyorin, Mylan, Mundipharma, Novartis, Regeneron Pharmaceuticals and Sanofi Genzyme; payment for travel/accommodation/meeting expenses from AstraZeneca, Boehringer Ingelheim, Mundipharma, Mylan, Novartis, Thermofisher; stock/stock options from AKL Research and Development Ltd which produces phytopharmaceuticals; owns 74% of the social enterprise Optimum Patient Care Ltd (Australia and UK) and 92.61% of Observational and Pragmatic Research Institute Pte Ltd (Singapore); 5% shareholding in Timestamp which develops adherence monitoring technology; is peer reviewer for grant committees of the UK Efficacy and Mechanism Evaluation programme, and Health Technology Assessment; and was an expert witness for GlaxoSmithKline. The authors report no other conflicts of interest in this work.

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