1	
2	British Thoracic Society Clinical Statement on Pulmonary Rehabilitation
3	Draft for consultation: 28 November 2022
4	
5	
6 7	Authors: William Man, Sally Singh, Emma Chaplin, Enya Daynes, Rachael Evans, Neil Greening, Claire Nolan, Matthew Pavitt, Nicola Roberts, Ioannis Vogiatzis
8	
9	
10	On behalf of the British Thoracic Society
11 12	
12	Available for public consultation from
14	28 November 2022 to 6 January 2023
15	
16	
17	
18	
19	
20	
21	
22	Contact:
23	British Thoracic Society,
24	17 Doughty St, London WC1N 2PL
25	miguel.souto@brit-thoracic.org.uk
26	
27	
28	
29	

30	
31	
32	
33	
34	Contents
35	Introduction
36	Section 1: Pulmonary rehabilitation: Access, Referrals and Uptake
37	Section 2: Assessment and Outcomes
38	Section 3: Extending the Scope of Pulmonary Rehabilitation
39	Section 4: Alternatives Models of Pulmonary Rehabilitation
40 41	Section 5: Adjuncts to and Maintenance of Pulmonary Rehabilitation
42	
43	
44	
45	
46	
47	
48	
49	
50	
51	
52	
53	
54	
55	

### 56 Introduction

- 57 The evidence-based British Thoracic Society (BTS) Guideline for pulmonary rehabilitation (PR) in adults was
- published in 2013.(1) There is a strong evidence base for the benefits of PR,(2) and it is one of the most cost-
- 59 effective interventions for adults with chronic obstructive pulmonary disease (COPD).(3) Furthermore, PR
- 60 improves exercise capacity and health related quality of life (HRQOL) in COPD to a much greater magnitude
- 61 than observed with bronchodilator therapy.(4)

62 Much of the guideline remains relevant today and does not need re-visiting. Since the guideline however, 63 there is deeper understanding of referral characteristics, outcome measures, patient selection, programme 64 delivery, potential adjuncts, and the role of maintenance following PR. The BTS Clinical Statement on PR will 65 provide a snapshot of current knowledge and best practice in topical areas by providing a series of clinical 66 practice points that are informed by evidence where this exists, or based on expert opinion and collective 67 clinical experience where evidence is limited. The intended audience are PR clinicians working within health 68 settings in the United Kingdom and beyond. The clinical statement will provide a framework to inform future 69 British Thoracic Society Quality Standards for PR. We have also highlighted areas of research priority, which 70 will be of interest to clinical researchers.

- 71 In this statement, we highlight the growing interest in alternative models of delivering PR (e.g. home-based,
- remote supervision, use of technology), accelerated by the restrictions placed on face-to-face PR delivery

73 during the global COVID-19 pandemic. Alternative PR models, typically delivered remotely, might potentially

increase provision of, and accessibility to PR. However research gaps remain and it is crucial these alternative

- 75 models are optimised and carefully evaluated before widespread adoption.(2).
- 76 A recent international workshop report, using a Delphi process, defined essential and desirable components
- of PR.(5) We have adapted this to define the core components of PR (Table 3), which will help health payers
- 78 decide if they are commissioning an intervention that is likely to produce good outcomes.
- 79

### 80 Methodology

The Clinical Statement Group (CSG) was chaired by Professor William Man and Professor Sally Singh. 81 82 Membership was drawn from an open application process. Members were selected for their experience 83 either in clinical delivery or academic evaluation (or both), and be representative of the multidisciplinary 84 team. The CSG identified key areas that reflect the scope approved by the BTS Standards of Care Committee 85 (SOCC). Following discussions of broad statement content, individual sections were drafted by CSG members 86 with the overall statement drafted by the chairs and reviewed by a patient representative. A final edited 87 draft was reviewed by the BTS SOCC before posting for public consultation and peer review on the BTS 88 website in (date to be confirmed). The revised document was re-approved by the BTS SOCC in TBC before 89 final publication.

- 90
- 91
- 92
- 93

### 94 Section 1: Pulmonary Rehabilitation: Access, Referrals, and Uptake

### 95 1.1 Access and Referrals

There is a large disparity between the number who are eligible and the number receiving PR.(6) Reasons for 96 97 this are complex, but barriers may exist at several points of the pathway. Referral from primary care appear to be influenced negatively by increasing age, gender (women less likely), deprivation, comorbidities, 98 99 respiratory disability and smoking status.(7) The PR outcomes from individuals with lower socioeconomic 100 status are not compromised, but they are less likely to be referred or to complete PR.(8) Over 10% of services 101 in England and Wales did not offer services to those with greatest respiratory disability (Medical Research 102 Council Dyspnoea Scale 5). Equity of access is rarely addressed within UK services, but modification of PR to 103 suit the needs of a diverse population has been proposed in other countries.(9) Health and digital literacy require attention, particularly with ever diversifying modes of PR delivery, including the use of 104 105 technology.(10, 11)

- Although there is a dearth of randomised controlled trial (RCT) data to support specific interventions designed to improve referral for PR,(12, 13) identified referrer barriers include a lack of referrer knowledge around eligibility criteria or how to refer for PR.(14) Several observational studies have provided indirect evidence that improving education can increase referral rates (summarised in Table 1).
- 110

### 111 Table 1: Effect of referrer education on pulmonary rehabilitation referrals

Action	Effect on referrals
Delivering education to primary care referrers (18-20)	3-5% increase
Patient education as part of a 'patient held score card' with advice to discuss referral at their next COPD review (21)	6% increase
Integrated approach to COPD care (22)	25% increase over three years
Delivering education to secondary care referrers (33, 34)	6% increase
	RR: 2.78 [2.65; 2.90]
Delivering COPD discharge bundles by pulmonary rehabilitation practitioners versus non-pulmonary rehabilitation practitioners (129)	OR: 14.46 [5.28 to 39.57]

<sup>112</sup> OR – adjusted odds ratio; RR – risk ratio; [] 95% confidence intervals

### 113

The most recent (pre-pandemic) national audit data identified that the median waiting time from receipt of referral to PR enrolment was 84 days, with only 54% receiving PR within 90 days of referral receipt.(15) A similar waiting time from prescription to receipt of an inhaler would be unacceptable, despite bronchodilators being a less cost-effective intervention to PR.(3) Commissioners need to ensure that accessibility to PR has at least the same priority as access to pharmacological therapy. This would require investment in workforce and training, with the BTS report "A workforce for the future" highlighting the substantial shortage of skilled health care professionals and support staff for PR.

121

122

### 123 1.2 Uptake and Completion

Barriers to uptake and completion of pulmonary rehabilitation are complex,(16, 17) but factors relating to the quality of a PR service, such as lack of patient-centeredness and coordination within PR team, inadequate professional competence of staff, lack of a holistic approach and limited accessibility, are relevant.(16)

There are few interventional studies targeting uptake and completion. Observational studies have explored interventions such as group opt-in sessions (which led to fewer patients attending assessment for PR (18)), patient-held manuals with research evidence summaries which improved attendance in the most socioeconomic disadvantaged patients,(19) and a nurse-general practitioner partnership care plan which increased attendance at PR by 21.5% compared to usual care.(20) In the acute setting, a patient co-designed education video did not improve post-hospitalisation PR uptake.(21) Other interventions currently being tested include the use of lay health workers to support patients.(22).

134

### 135 Clinical Practice Points

- PR provider leads should have designated sessional time to coordinate management and delivery of the service. This should include: regular education of potential referrers about PR and referral pathways; the expansion, training and skills maintenance of a specialist workforce to deliver PR; the collation of key organisational metrics.
- PR providers should demonstrate the offer of timely, accessible and high quality services by the regular monitoring and publication of key organisational metrics including waiting time from referral receipt to assessment and enrolment, percentage of referred patients who attend an assessment, percentage of patients who are assessed that attend at least one planned supervision, percentage of the number of attended to planned sessions, percentage of patients attending a discharge assessment.
- PR providers should work closely with relevant national professional societies and other stakeholders to
   develop consensus training programmes, competency documents and plans to develop and support a
   skilled workforce to deliver increased PR.
- 148

### 149 Research gaps

- Development of interventions to improve referrals to, uptake and completion of PR.
- The adaptations and evaluation of PR services to ensure programmes meet the needs of a diverse
   population, including equity of access.
- 153
- 154
- 155
- 156 157
- 158
- 159
- 160
- 161
- 101
- 162
- 163

### 164 Section 2: Assessment and Outcomes

PR assessments are documented in the previous BTS guideline and include measures of breathlessness, exercise capacity and HRQOL.(1) These remain core outcomes. However, this section will explore additional assessments and outcomes that are complementary and should be considered as part of a high-quality PR service.

169

### 170 2.1 Holistic Assessment

The PR pathway presents an opportunity to optimise holistic care. A thorough assessment for PR incorporates a multi-system approach. This should help identify individuals who might benefit from other cost-effective interventions such as vaccination and smoking cessation,(3) or those identified with treatable traits associated with poor prognosis that might prompt onward referral.

There is a significantly increased risk of several cardiovascular diseases in COPD (23) so unexplained 175 176 symptoms (such as chest pain or intermittent claudication), or identification of elevated blood pressure or 177 arrythmias should prompt referral for further evaluation. Long term oxygen therapy for severe hypoxaemia 178 remains one of the few interventions that influence prognosis in adults with COPD.(24, 25) Both low body 179 mass index (specifically unintentional weight loss) and extreme obesity are factors for poor prognosis.(26-180 28). Frailty, a multisystem syndrome characterised by reduced functional reserve and increased vulnerability 181 following minor stressor events, is associated with adverse prognosis in adults with COPD,(29-31) and 182 increases likelihood of PR non-completion.(32) Mental health issues are common in patients referred for PR (15, 33) and are associated with reduced adherence to interventions, increased dyspnoea, and lower levels 183 184 of patient activation.(34-37)

Education is a key component of PR; yet assessing the effects of this component is challenging, with limited availability of validated questionnaires, particularly for non-COPD conditions.(38) Although validated COPD knowledge questionnaires have been used in PR settings,(39, 40) further research is needed to determine the impact of the educational component beyond knowledge acquisition. A list of suggested educational topics were published in the previous BTS guidelines.(1)

190

### 191 2.2 Home-based or remote assessment of core outcomes

Since the COVID-19 pandemic, there has been increasing interest in home-based or remote assessment options. Many non-exercise outcomes, such as HRQOL, are assessed through questionnaires. The COPD Assessment Test, Saint Georges Respiratory Questionnaire and Hospital Anxiety and Depression Scale have comparable validity and reliability when delivered over the phone compared to face-to-face delivery.(41, 42)

196 However, evidence is lacking to support remote delivery of functional or field walking tests as a reliable 197 alternative to face-to-face testing. Although sit to stand, step, and timed up and go tests are feasible in the 198 home-setting, they do not accurately reflect oxygen desaturation with walking or allow exercise 199 prescription.(43) Six-minute walk tests (6MWT) supported by mobile phone application algorithms offers a 200 potentially attractive approach but has not been validated in chronic respiratory disease populations.(44) 201 There are some data to suggest that there is no significant difference in six-minute walk distance when 202 performed indoors or outdoors,(45) although further corroboration is required in variable environmental 203 conditions. Current assessment of patient safety for exercise-training and exercise capacity to facilitate 204 exercise prescription should be conducted in-person, irrespective of the PR delivery model (see Section 4).

205

### 206 2.3 Functional assessments

Simple functional assessments are attractive as they do not require as much space as field walking tests (46) and can be performed in most healthcare settings including the home. These include four metre gait speed (47-49), sit to stand tests (five repetition, 30 seconds, one minute) (50-53), step tests (54-57), timed up and go (58), and composite measures combining several functional tests. These have been reviewed in detail elsewhere (43, 59-61). These functional tests are safe and feasible in the home setting, have a moderate relationship with field walking test performance or muscle strength and are responsive to exercise-based interventions or PR.

214 However, there are several caveats. Most validation studies have taken place in clinical settings where the 215 tests were directly supervised and therefore the safety and validity of remotely supervised functional tests 216 in patients with chronic respiratory disease have not been established. Some functional tests have floor or 217 ceiling effects that might limit their application in PR. For example, 15% of those referred for PR were not 218 able to complete the five repetition sit to stand, (50) whilst the four metre gait speed is less responsive to PR 219 in higher functioning individuals with COPD.(47) Functional tests are also typically submaximal, and therefore 220 not able to support individualised exercise prescription.(47) Others have used functional tests as surrogate markers of muscle strength. However the relationship between five repetition sit-to-stand test and 221 222 quadriceps strength is only moderate.(50)

223 224

### 225 2.4 Physical activity

Reduced physical activity (PA) is associated with poor prognosis in COPD.(62) Although PA can be measured subjectively using questionnaires, there are limitations to this method including recall bias.(63) There is growing literature on measuring PA using wearable devices, including pedometers and accelerometers, but considerable variability has been reported in clinical trials.(64) An International Taskforce on Physical Activity has recommended implementation of standard operating procedures for PA data collection and reporting.(62) Although PA has been identified as an important outcome that may be potentially amenable to PR, further research is required before adoption into routine clinical practice.

233

### 234 Clinical Practice Points

As well as establishing safety and suitability for exercise training and facilitation of exercise prescription,
 a high quality PR assessment should encompass a holistic approach incorporating documentation of
 vaccination and smoking/vaping status; resting oxygen saturations, heart rate and blood pressure
 measurements; nutritional assessment; frailty; presence of anxiety and depression; and disease
 knowledge. This information should be communicated to other relevant healthcare professionals
 involved in the individual's management so that required action can be coordinated.

Assessment of patient safety for exercise-training and exercise capacity to facilitate exercise prescription
 should be conducted in-person using a validated field walking test (incremental shuttle walk, 6MWT) or
 laboratory cardiopulmonary exercise test.

• There is no evidence to support the safety or validity of field walking tests or simple functional tests that are supervised remotely.

- When routine face-to-face assessments are restricted, hybrid assessments can be considered with
   questionnaire-based assessments conducted over the telephone and a directly supervised, face-to-face
   assessment of exercise capacity.
- Functional tests are complementary to, but not a replacement for, validated exercise walking tests. There
   is no evidence to support aerobic or strength exercise prescription from simple functional tests.

### 252 Research gaps

- Development of outcomes that assess the effectiveness of the education component of the PR
   programme
- Studies to assess the safety and validity of remotely supervised exercise and functional outcomes through
   video-conferencing or mobile applications.
- Alternative strategies to prescribe exercise and deliver effective PR in the absence of a directly supervised
   validated exercise test.
- Clarify the value of measuring PA and other physiological data obtainable from wearables as part of
   routine clinical practice in PR.

### 278 Section 3: Extending the Scope of Pulmonary Rehabilitation

### 279 3.1 Chronic Respiratory Disease other than COPD

There is a growing evidence-base and real-world experience of delivering PR to people with asthma, bronchiectasis and interstitial lung disease (ILD). Systematic reviews have demonstrated that exercise training, compared with control interventions, significantly improves exercise capacity and HRQOL.(65-68) Furthermore, real-world data suggest that these improvements are of similar magnitude to those observed in matched patients with COPD.(33, 69, 70)

Considerations and potential adaptations needed to deliver PR to people with non-COPD chronic respiratory 285 286 disease are outlined in Table 2. For asthma, to minimise risk of adverse events, patients should be medically 287 optimised prior to referral for PR.(71) Similarly, as bronchiectasis is characterised by excessive sputum 288 production, a review and optimisation of airway clearance technique should be considered prior to starting 289 PR.(72) There are no data to support increased risk of cross-infection of multi-resistant organisms,(73) but 290 local infection control policies should be followed. Compared with COPD, profound exercise-induced oxygen 291 desaturation is more common in IPF and some subtypes of ILD; this needs to be considered as part of the 292 safe assessment and delivery of PR in these patients. Although most standard PR education is relevant to 293 people with non-COPD respiratory disease, some adaptations are needed (eg. medications) or particular 294 components prioritised (eg. airway clearance in bronchiectasis).

295	
296	
297	
298	C
299	
300	
301	XO
302	
303	
304	
305	
306	
307	
308	
309	
310	
311	
312	
313	

### 314 Table 2. Disease-specific considerations for pulmonary rehabilitation

Asthma	<ul> <li>To minimise risk of adverse events, patients should be medically optimised prior to PR referral.(71)</li> </ul>
Bronchiectasis	<ul> <li>Bronchiectasis is characterised by excessive sputum production, therefore a review and optimisation of airway clearance technique is recommended prior to starting PR.(72)</li> <li>There are no data on risk of cross-infection of multi-resistant organisms during PR, but local infection control policies should be followed.</li> </ul>
Interstitial lung disease	<ul> <li>Compared with COPD, profound exercise-induced desaturation is more common in idiopathic pulmonary fibrosis and some sub-types of interstitial lung disease; this needs to be considered as part of the safe assessment and delivery of PR in these patients.</li> </ul>
Post-COVID-19	<ul> <li>Check for contraindications and beware unexplained chets pain. Unidentified (and therefore untreated) pulmonary thromboembolic disease (84) and myocarditis (172) have been reported in the post- COVID-19 syndrome, which are relative contraindications to PR.</li> <li>Assessment: A proportion of patients will have post-intensive care syndrome with multi-systemic symptoms. The following symptoms should be assessed to enable the exercise and education components to be individualised: fatigue, muscle weakness, breathing pattern disorder, post-traumatic stress, swallow/speech difficulties, and peripheral neuropathy.</li> <li>Monitoring: Post-exertional symptom exacerbation (PESE) is a widely reported symptom in post-Covid 19 syndrome.(173) Given the potential for deterioration in function following overexertion, fatigue and PESE</li> </ul>
lung cancer	<ul> <li>should be closely monitored during PR.</li> <li>Due to time sensitivity for curative surgery conventional PR programmes</li> </ul>
	would require adaptation to be suitable for prehabilitation.
Chronic heart failure	<ul> <li>Programme adaptations/considerations might include:(99)</li> <li>Exercise assessment with an exercise ECG.</li> <li>Provision of disease-specific education and non-exercise interventions to address breathless and psychological needs of patients and carers.</li> <li>Workforce training to understand signs of an episode of decompensated heart failure.</li> <li>Inclusion of a heart failure nurse in the multi-disciplinary team.</li> </ul>
Pulmonary hypertension	<ul> <li>To be eligible for PR, people with pulmonary hypertension should:(101, 103)</li> <li>Have stable disease (&gt;3 months).</li> <li>Be prescribed drug therapy with no change in previous two months.</li> <li>Have no recent syncope.</li> <li>International guidelines recommend that exercise is supervised by specialist exercise professionals (103)</li> </ul>

<sup>316</sup> Abbreviations: ECG: Electrocardiogram; PR: Pulmonary Rehabilitation.

### 320 3.2 Post-COVID-19

321 Previous guidance from the BTS regarding the role of adapted PR to meet the recovery needs in post-COVID-322 19 syndrome has been previously published. (74) Several observational studies have demonstrated that PR 323 following hospitalised COVID-19 is associated with significant improvements in exercise capacity, 324 breathlessness, and HRQOL.(75-80) Without a control group, natural recovery cannot be dismissed as the 325 main driver of improvements.(56) However, symptom burden, reduced exercise tolerance and sequelae of 326 hospitalisation for COVID-19 remain substantial at five months post-discharge,(81) with negligible 327 improvement one year after discharge. (82) Initial trial data suggest a role for PR in the recovery of individuals 328 with post-COVID-19 syndrome, (83) and the results of further trials are awaited

329 Several factors need to be considered when providing PR to individuals with post-COVID-19 syndrome (Table 330 2). A proportion will have post-intensive care syndrome with multi-organ impairment, and there should be a 331 wider assessment for symptoms such as fatigue, muscle weakness, breathing pattern disorder, post-332 traumatic stress, swallow/speech difficulties, and peripheral neuropathy. These should also be considered 333 with regards to individualising the exercise and education components of the programme. Unidentified (and 334 therefore untreated) pulmonary thromboembolic disease (84) and myocarditis (85) have been reported in the post-COVID-19 syndrome, which are relative contraindications to PR. Furthermore, post-exertional 335 336 symptom exacerbation (PESE) is a widely reported symptom in post-Covid 19 syndrome.(86) Given the 337 potential for deterioration in function following over-exertion, fatigue and PESE should be closely monitored 338 during PR.

339

### 340 3.3 Lung Cancer

Prehabilitation is the focus on modifiable risk factors in individuals preparing for lung cancer treatment, typically commencing at the point of diagnosis and is multimodal in approach.(87) A systematic review suggested that exercise pre-surgery improves physical and pulmonary function, although the interventions were very heterogeneous in nature and duration.(88) Whilst PR addresses some modifiable factors, the timesensitivity of lung cancer resection means that the traditional outpatient PR model would need significant adaptations to be suitable for prehabilitation (Table 2).

A Cochrane review identified eight RCTs of exercise-training following surgical resection of non-small cell lung cancer (89). Compared with usual care, improvement in exercise capacity was greater in the intervention group, but trial populations were small and there was lower certainty for other outcomes. Due to the significant heterogeneity of the interventions, the optimal timing, setting, nature or duration of exercisetraining for post-lung cancer surgery patients remains unclear. Few patients are currently referred for PR after lung cancer surgery.(90) Little data exists on rehabilitation interventions that combine pre- and postlung cancer surgery exercise-training.

354

### 355 3.4 Lung Volume Reduction Surgery

Lung volume reduction surgery (LVRS) is recommended by the National Institute for Health and Care Excellence (NICE) for the treatment of selected individuals with emphysema and hyperinflation.(91) As part of the work-up for LVRS, all individuals should receive PR, a prerequisite to randomisation in landmark trials of LVRS.(92) Furthermore, it plays an important role in selecting individuals for LVRS with up to 20% improving their exercise tolerance to such an extent that they change LVRS risk stratification groups.(93) In the UK, only a small minority of eligible patients undergo LVRS due to the absence of standardised referral pathways.(94) However, PR practitioners may have a role in identifying potential candidates as the post-PR assessment represents the point at which the patient's functional capacity and management of breathlessness should be optimised. Recent analysis of data from the National Asthma and COPD Audit suggested that up to 18.1% of PR completers met the NICE criteria for a LVRS-focused respiratory review (Non-smoker, MRC≥3, 6MWT > 140m or ISWT >80m).(95)

367

### 368 3.5 Lung Transplantation

369 Before referral for lung transplantation, individuals with advanced lung disease should have been optimised, 370 including completion of PR. Unlike for lung cancer, waiting time for lung transplantation is unpredictable, and 371 there is little guidance on the ideal content or duration of a prehabilitation programme for lung 372 transplantation, and consequently few published data.

Exercise-training following lung transplantation has been studied in more detail. A Cochrane review to determine the benefits and safety of exercise training in adult lung transplant recipients included eight RCTs involving 438 participants.(96) However, results could not be aggregated due to the small number of underpowered trials and the heterogeneity of the interventions. The authors concluded that the effects of exercise-based rehabilitation following lung transplantation were uncertain due to imprecise estimates of effects and high risk of bias.(96)

379

### 380 3.6 Cardiac Disease and Pulmonary Hypertension

Cardiac comorbidity is highly prevalent in patients attending PR.(15) There is no convincing data to suggest that stable cardiac comorbidity is associated with worse outcomes to PR.(97) Exercise-based cardiac rehabilitation is safe in individuals with chronic heart failure (CHF) and improves exercise capacity and HRQOL.(98) Integrating individuals with CHF and those with chronic respiratory disease into breathlessness rehabilitation programmes is feasible with minor adaptations (Table 2).(99) These improve exercise capacity in CHF, with a magnitude similar to that observed in COPD.(100) Only 18% of PR services in the UK currently accept patients with CHF.(15)

388 In a systematic review of seven trials in patients with primarily pulmonary arterial hypertension (PAH) 389 (including some with chronic thromboembolic pulmonary hypertension: CTEPH), exercise-based 390 rehabilitation improved 6MWT distance and peak oxygen consumption compared with usual care.(101) 391 However, the quality of evidence was low and the rehabilitation interventions were inpatient-based and 392 atypical of PR practice in the NHS. Collective experience is that exercise-training is safe and effective in 393 PAH,(102) and in those with pulmonary hypertension secondary to chronic lung disease. However, expert 394 consensus is that patient selection is key (stable disease with no recent change in drug therapy or recent 395 history of syncope).(101, 103). In PAH and CTEPH, exercise-based rehabilitation should be directly supervised 396 in person by specialist exercise health care professionals.(103)

397

### **398 3.7 Pulmonary Rehabilitation around the time of a hospitalised exacerbation of COPD**

Extrapulmonary manifestations of hospitalised exacerbations include reduced walking performance (104, 105), HRQOL (106, 107), low physical activity levels (108) and muscle dysfunction (109) – all of which are associated with poor prognosis, (30, 104, 110) but also potentially responsive to PR.

The BTS Guideline on PR recommended that individuals hospitalised for acute exacerbation of COPD should be offered PR at hospital discharge to commence within one month of discharge.(1) The Cochrane systematic review included 20 trials and 1477 participants and demonstrated moderate to large effects of rehabilitation on HRQOL and exercise capacity in patients with COPD after an exacerbation.(111) Additionally there is evidence that PR after hospitalised exacerbation may reduce the risk of readmission (112, 113) and improve survival with a dose-response effect.(114) However, the content, setting and duration of rehabilitation interventions were heterogeneous.

- 409 In the UK setting, inpatient rehabilitation may not be feasible given the short duration of hospital stays. Two 410 trials conducted in the NHS setting evaluated PR initiated during the inpatient stay and progressing to a more 411 "light touch" approach to post-discharge outpatient treatment with the aim of addressing both the initial 412 insult of the hospitalisation as well promoting recovery.(115, 116); however benefits were less impressive 413 than observed in post-exacerbation outpatient PR trials.(112, 115-117) Rehabilitation started one month 414 after hospitalisation yielded better overall results than rehabilitation started during the hospital 415 admission.(118) A systematic review, including 30 studies, identified that longer programmes, starting after 416 hospital discharge and including an educational component (as well as exercise), were most effective at 417 reducing hospital readmissions.(119)
- Implementation of PR following an exacerbation remains a challenge. Real-world data suggests that uptake is between 1.5% and 9%.(114, 120) Strategies to improve referral, uptake and completion have been limited.(21) "Delayed" PR following a hospital admission is still associated with benefits (121) and therefore it is important to re-offer PR to people who initially decline in the immediate post-hospitalisation period.

422

### 423 Clinical Practice Points

- PR should be offered to symptomatic individuals with asthma, bronchiectasis and ILD.
- PR may be helpful in the recovery of subgroups of patients with post-Covid-19 syndrome.
- The assessment, exercise and education components of PR should be adapted for relevant
   cardiorespiratory diseases, taking into account disease-specific issues.
- The workforce should receive training and be competent to deliver high-quality PR for relevant
   cardiorespiratory diseases.
- PR practitioners should have the skill set to support prehabilitation interventions for patients awaiting
   lung cancer and lung transplant surgery, but the current delivery model of PR needs to be adapted in
   order to be appropriately time sensitive.
- PR practitioners have a role in identifying potential candidates for LVRS.
- Patients with stable CHF, PAH or CTEPH can be incorporated safely within directly supervised outpatient
   PR programmes.
- Outpatient supervised PR, incorporating both exercise-training and education should be offered to all
   appropriate patients discharged from hospital after exacerbation of COPD.
- Members of the integrated care team should re-offer "delayed" PR in individuals who decline an initial
   offer of post-hospitalisation PR.

440		
441	Research gaps	
442	<ul> <li>Trials to understand the role of PR in the recovery of post-Covid-19</li> </ul>	9 syndrome.
443 444	<ul> <li>Trials to determine the optimal timing, setting, nature or duratic cancer and post-lung transplant surgery.</li> </ul>	ion of exercise-training for post-lung
445 446	<ul> <li>Trials to evaluate the effects of PR in hospitalised exacerbations of COPD.</li> </ul>	chronic respiratory disease other than
447	<ul> <li>Interventional trials designed to increase referral to and uptake of</li> </ul>	post-exacerbation PR.
448	• The role of alternative remote PR models in the post-exacerbation	setting.
449		
450		
451		$\mathcal{O}$
452		<b>)</b>
453		
454		
455		
456		
457		
458		
459		
460		
461		
462		
463		

#### 465 Section 4: Alternatives Models of Pulmonary Rehabilitation

466

### Barriers to traditional hospital-based PR have been well documented.(17, 122) This has highlighted the need 467

for alternative modes of delivering PR, as these may potentially increase uptake and accessibility. 468

469 National audit data show that non-medical, community-based settings are increasingly used to deliver

470 supervised PR in the UK.(6) PR delivered in a community setting has similar efficacy to that produced in a

471 hospital-based setting.(123) Supervised PR using minimal resources have similar efficacy to programmes

using specialist exercise equipment.(124) 472

473 Home-based rehabilitation spans a range of delivery options ranging from standardised manuals, web-based 474 applications, tele-rehabilitation and face-to-face supervision. Across all these modes, the level and frequency 475 of supervision and contact with a health care professional may vary dramatically. Commissioners need to 476 consider carefully whether alternative models delivered by providers include core components detailed in 477 Table 3. Although some PR models might involve remote supervision, published trials have all incorporated 478 a directly supervised face-to-face, validated exercise test prior to the intervention to evaluate safety and 479 facilitate exercise prescription. A further consideration is digital literacy (10, 11) and avoiding the exclusion 480 of individuals uncomfortable with technology.

# Table 3: Core Components of a Pulmonary rehabilitation programme

- An initial face-to-face assessment by a suitably trained health care professional; ٠
- Initial assessment must include a validated exercise test from which an individualised exercise prescription can be obtained;
- Endurance and resistance training, which is individually prescribed and progressed with regular supervision from suitably trained health care professionals;
- A structured education programme;
- Delivered by a dedicated team of health care professionals trained in exercise assessment, prescription and progression and delivery of education on chronic respiratory disease management;
- The programme model, including assessment and delivery components, must have been previously tested in a clinical trial and shown to be safe and effective;
- Measurement of core outcomes before and after PR, including a validated exercise test;
- Participation in regular audit of organisational and clinical outcomes; for example engagement with a recognised national audit programme where available.
- Regular external peer review, for example engagement with a recognised accreditation programme where available.

481

#### Home-based, non-digital 482 4.1

483 In this model, individual patients are provided with a manual, exercise diary or written material which provides structured exercise and educational components (Table 4). These are usually supported by remote 484

485 supervision from skilled PR health care professionals. Previous data suggest that this model does improve 486 HRQOL and exercise capacity compared with usual care, although differences are modest.(125) Trials that 487 have compared home-based models supported by manual and telephone support with outpatient, centrebased PR have produced short-term clinical outcomes that are similar to centre-based PR.(126-128) 488 489 However, an interesting observation is that "gold-standard" centre-based rehabilitation did not produce the 490 expected improvements in exercise capacity. In a real-world study, a home-based, manual-structured 491 programme with weekly telephone supervision produced similar improvements in HRQOL, but smaller 492 changes in exercise capacity, compared to a propensity-matched cohort undergoing twice-weekly centre-493 based supervised programme.(129)

Although home-based programmes typically involve less frequent staff contact than centre-based approaches, that contact is conducted one-to-one, and therefore data are required to evaluate the cost effectiveness of such an approach. Other home-based therapies include the use of neuromuscular electrical stimulation which improves muscle weakness in those with advanced disease.(130, 131) However, the effect on exercise capacity is unclear.(130, 132).

# Table 4: Comparison of Home-based, non-technology versus centre based PR or usual care: summary of selective studies

Study	Population	Intervention / Control	Outcomes
Maltais 2018 (128)	252 with COPD	Home based (including one home visit and weekly telephone calls) versus Outpatient centre based rehabilitation supervised PR for eight weeks. Both groups received four weeks of in-person centre- based education	Similar changes in dyspnoea, health status and exercise capacity at 3 months and 12 months
Holland 2017 (126)	166 with COPD	Home based (including one home visit and weekly telephone calls) programme versus Outpatient centre based supervised PR for eight weeks	Short term clinical outcomes equivalent to centre based PR but neither effective maintenance at 12 months
Horton 2018 (127)	287 with COPD	Structured unsupervised home based programme including a manual and telephone support for seven weeks versus Centre based supervised PR for seven weeks	Evidence of significant gains in CRQ-D at 7 weeks in both groups. Inconclusive that homebased PR was non- inferior to PR in dyspnoea favouring the centre group at 7 weeks
Nolan 2019 (129)	154 with COPD	Home based structured exercise programme with weekly telephone calls versus Centre-based supervised PR for eight weeks	Significant improvements in both groups in exercise capacity but home-based group demonstrated smaller improvements; clinically and statistically significant improvements in QoL within each group. Completion rates were low in both groups

Mitchell 2014 (125)	184 with COPD	Structured unsupervised home based programme including a manual and telephone support for 6-weeks versus usual care	Significant differences between groups in QoL, exercise performance, anxiety, and disease knowledge at 6 weeks; Intervention did not improve dyspnoea over and above usual care at 6 months
			1

501 PR – Pulmonary rehabilitation; CRQ-D – Chronic respiratory questionnaire – dyspnoea domain; QoL – Quality
 502 of life

# 503 4.2 Home-based Web Platform

These are similar to home-based models described in 4.1, except that the programme is supported by a web-504 505 based platform or app (Table 5). A home-based, online platform, "MyPR", was compared with face-to-face 506 PR delivered in an outpatient setting, and demonstrated that "MyPR" was safe and well tolerated, and non-507 inferior to the control arm in terms of effects on exercise capacity and symptom scores.(133) However, the 508 trial population was selective (exclusion criteria included exercise-induced oxygen desaturation, functional 509 limitation, comorbidities, poor digital literacy), and the control arm was not a conventional supervised PR 510 programme, but comprised exercise stations matched to those provided by the online platform.(133) 511 Completers of both a home-based interactive web platform "SPACE for COPD" and a standard care outpatient PR programme showed similar improvements in endurance shuttle walk and dyspnoea.(134) However 512 engagement with digital technology was challenging; only 103 of 2646 invited patients were randomised, 513 514 whilst 57% of the web platform arm dropped out.(134) Both platforms provided an introductory face to face 515 session, with either contact details provided for further queries (133) or weekly contact via email or 516 telephone using a standardised proforma.(134)

Study Population		Population	Intervention / Control	Outcomes	
	Chaplin 2017 (134)	103 with COPD	Web-based programme (SPACE for COPD) of exercise and education versus centre based supervised PR, twice weekly, 2 hourly sessions for 7 weeks (4 weeks supervised; 3 weeks unsupervised)	Interactive web-based PR programme is feasible and acceptable when compared with centre based PR; statistically significant improvements within groups for exercise capacity and dyspnoea but not between groups. Dropout rates were higher in the web-based programme	
	Bourne 2017 (133)	90 with COPD	6-week Online PR via log in and access to 'myPR' versus a supervised PR programme group sessions in a local rehabilitation facility	Online supported PR was non-inferior to a conventional model delivered in face-to-face sessions in terms of effects on 6MWT distance and symptom scores. Online PR was safe and well tolerated.	

### 517 Table 5: Comparison of Home-based, web platform versus centre based PR: summary of selective studies

518 SPACE for COPD – Self management programme of activity, coping and education for Chronic obstructive

519 pulmonary disease; PR – Pulmonary Rehabilitation; 6MWT – six minute walk test

520

### 521 4.3 Video Tele-rehabilitation

- 522 Video tele-rehabilitation encompasses synchronous real-time PR supported by video-conferencing. A small 523 trial showed that video tele-rehabilitation improved endurance exercise capacity and self-efficacy in patients 524 with COPD when compared with usual care.(135) Two studies have compared video tele-rehabilitation with
- face-to-face centre-based PR, and shown similar effects on exercise capacity and HRQOL.(136, 137) However
   the improvements in exercise capacity were modest in both intervention and standard care arms (Table 6).
- 527 Furthermore, participants were provided with video technology and specialist exercise equipment to use in
- 528 the home for free, which may not be generalisable to the NHS setting.
- 529 Outside of the home-setting, video-conferencing has also been utilised to support satellite tele-rehabilitation
- 530 centres ("hub and spoke" model).(138, 139) Trials are needed to test the effects of such models on patient
- 531 throughput, staffing ratios and travelling for patients.(139)

## 532 Table 6: Comparison of video tele-rehabilitation versus usual care without exercise or standard care:

### 533 summary of selective studies

Study	Population	Intervention / Control	Outcomes
Tsai 2017 (135)	37 with COPD	Supervised home-based real-time video tele-rehabilitation (exercise three times/week for eight weeks) versus usual care without exercise training	Statistical, and clinically significant, improvement in endurance shuttle walk time in video tele-rehabilitation group, but underpowered to demonstrate improvements in incremental shuttle walk or six-minute walk
Hansen 2020 (137)	134 with COPD	10-week video tele-rehabilitation programme versus supervised face- to-face rehabilitation	Similar changes in exercise capacity, breathlessness and HRQOL, but changes in both groups very modest and probably not clinically significant
Cox 2021 (136)	142 with chronic respiratory disease (100 with COPD)	Video tele-rehabilitation programme versus supervised centre-based PR, both interventions 8 weeks with 16 sessions	Video-telerehabilitation appeared safe and provided clinically meaningful improvements in dyspnoea and HRQOL, but equivalence to traditional PR not shown

### 534 HRQOL – health-related quality of life; PR – pulmonary rehabilitation

### 535 4.4 Virtual reality

536 Virtual reality is an emerging technology that might provide an interactive and visually stimulating approach 537 to providing PR in the home setting.(140) To date, there are few published data, of which most have 538 limitations in the reporting quality.(141) Acceptability is also unknown in a patient population that 539 traditionally have digital hesitancy.(10) 540

### 541 4.5 Active Mind-Body Movement Therapies

542 Three systematic reviews have examined the deployment of active mind-body movement therapies as an 543 alternative to pulmonary rehabilitation.(142-144) Two reviews compared Tia' Chi or yoga against non-544 exercise control groups and identified statistically significant improvements in both exercise capacity and 545 HRQOL, concluding that tai chi or yoga may be a useful adjunct to rehabilitation (142, 143). A later review 546 compared active mind-body movement therapies (largely tai chi and/or qigong) as an adjunct to or in 547 comparison with pulmonary rehabilitation.(144) Overall, the data was of poor quality, the impact on both 548 exercise capacity and HRQOL remained inconclusive, and none conducted in NHS settings. (53) A recent trial 549 directly compared PR (three sessions a week) to Tai Chi (five sessions a week) for 12 weeks.(145) While there 550 were important changes in HRQOL in both groups, neither group reached the minimal clinically important 551 difference for the 6MWT distance.(145) The population was atypical of those usually referred for PR with a 552 pre-PR 6MWT distance of over 500 metres. Standardised reporting is crucial to our understanding and development of these modes of delivery, which is important to attract a more diverse population.(146) 553

554 Overall, the outcomes of alternative models of PR have been heterogeneous and studies need to be 555 interpreted with caution. Although systematic reviews have suggested that alternative models of PR achieve 556 outcomes similar to those seen in traditional centre-based PR,(147) the certainty of evidence is limited by 557 the small number of studies with relatively few participants, varying models of care, and whether models are 558 generalisable to the NHS setting. Almost all published data are restricted to COPD.

559 Notably, a near universal observation is the lower-than-expected benefits associated with the "gold-560 standard" centre-based arm in equivalence or non-inferiority trials. This may reflect selective trial 561 populations lacking equipoise. Furthermore, systematic reviews of telerehabilitation studies have shown that 562 the mean change in six minute walk distance with telerehabilitation are lower than the established minimum 563 clinically important difference, (147) and lower than that observed with centre-based PR (Figure 1). (2) Real-564 world observational data have shown that home-based, remotely supervised PR are associated with a smaller 565 magnitude of change in exercise-capacity, about half of that seen in directly supervised, centre-based PR.(129) There is no published data on hybrid models (which combine limited centre-based with home-based 566 567 PR).

- 568
- 569
- 570
- 571
- 572
- 573

574

- 575
- 576
- 577

### 578 Figure 1

- 579 Data taken from two Cochrane reviews a) Cox N et al (147) and McCarthy B et al (2) comparing the 6MWT
- response data to home-based telerehabilitation and directly supervised PR based interventions. Data for
- the 4 trials on the left is data from 4 RCT's comparing tele-rehabilitation and centre based rehabilitation,
- and the data point on the right is combined data from 38 trials included in the Cochrane review of
- 583 supervised pulmonary rehabilitation which used six minute walk as an outcome.



584 585

# 586 Clinical Practice Points

- Every individual referred for PR should have the opportunity to access directly supervised, centre-based
   PR in a timely way as this model is supported by a convincing evidence base.
- In patients who decline or drop out from supervised centre-based PR, providers should offer an alternative model of delivery. Any alternative model should have a supporting evidence base (published trials, ideally in the NHS setting), and incorporate a directly supervised, validated exercise test from which individualised exercise can be prescribed.
- Both staff and patients require training to support alternative PR models, particularly those involving
   digital technology.
- Commissioners and providers should ensure that the delivery of alternative PR models do not promote
   digital exclusion.
- 597

### 598 Research gaps

Further trials are required to evaluate the efficacy and clinical effectiveness of alternative models of PR,
 including hybrid models, particularly in the NHS setting.

601	•	An agreed framework for the reporting of technology-based interventions, including core datasets and
602		outcomes.

603	Alternative models of PR delive	v should be evaluated in chronic	respiratory diseases other than COPD.
000			respiratory discuses officer than oor br

604	
605	
606	
607	
608	
609	
610	
611	
612	
613	
614	
615	
616	
617	
618	
619	
620	
621	
622	
623	

### 624 Section 5: Adjuncts to and Maintenance of Pulmonary Rehabilitation

625

626 Since the BTS guideline,(1) several trials have informed on the potential utility of adjunctive strategies to 627 improve PR outcomes.

628

### 629 5.1 Oxygen supplementation

630 Oxygen supplementation in the experimental setting acutely enhances endurance exercise performance in 631 individuals with COPD.(148-150) However, this has not translated to augmented outcomes in PR. In a 632 multicentre trial, 111 participants with COPD and exercise-induced oxygen desaturation were randomised to 633 receive either supplemental oxygen or room air during an eight-week exercise-training programme.(151) 634 Exercise capacity and HRQOL improved in both groups, with no additional benefit from training with supplemental oxygen.(151) The majority of participants had only modest exercise induced oxygen 635 636 desaturation, and the acute physiological response to oxygen was not tested prior to the training programme.(152) Limited data exist regarding the role of supplemental oxygen during PR in conditions other 637 638 than COPD.

639

### 640 5.2 Non-invasive ventilation (NIV)

Systematic reviews and meta-analyses of studies using NIV during supervised exercise training provide 641 642 conflicting evidence of the benefits. One meta-analysis showed improvements in endurance exercise capacity 643 with the addition of NIV (153), whilst another meta-analysis found similar responses to exercise training 644 between NIV supported and sham arms (154). In hospitalised exacerbations of cystic fibrosis and 645 bronchiectasis, Dyer and colleagues demonstrated that application of NIV could acutely improve endurance 646 cycling time (155), but there were concerns about patient acceptability. Practical considerations include the 647 additional equipment needed and time required to supervise patients on NIV during PR; this is less 648 problematic in those already established on domiciliary NIV.(156)

649

### 650 5.3 Inspiratory muscle training (IMT)

51 Since the guideline, three large RCTs have investigated the value of IMT as an adjunct to PR. Although IMT 52 improved inspiratory muscle strength, particularly in those with inspiratory muscle weakness,(157) 53 significant additive benefits of IMT to PR in core outcomes such as exercise capacity or HRQOL are less 54 convincing (157-159). Limited and conflicting data exist in respiratory disease other than COPD.(160, 161)

655

### 656 5.4 Physical activity (PA) counselling

Physical inactivity is associated with poor prognosis in COPD.(62) The effects of PR alone on physical activity levels are relatively modest (162). A systematic review demonstrated that PA promotion with pedometers as an adjunct to PR improves step counts/day,(163) although studies were small and results heterogeneous. A trial conducted in the NHS setting randomised 152 participants with COPD to an eight-week PR programme either with or without pedometer-directed step targets reviewed weekly.(164) No significant differences in change in time spent in moderate intensity activity, exercise capacity or HRQOL were seen between groups.(164) Studies exploring behavioural counselling as an adjunct to PR, typically using motivational

- interviewing, have produced mixed results.(165-167) As discussed in Section 2.4, PA data collection and
- 665 reporting should conform to international consensus recommendations.(62)
- 666

### 667 5.5 Maintenance of pulmonary rehabilitation

The beneficial effects of PR decline over one year.(168) The previous BTS guidelines recommended that PR
 graduates should be encouraged to continue exercise. However the format and delivery of maintenance
 programmes reported in the literature vary significantly.(169)

The evidence for maintenance programmes after PR are inconsistent (Table 7). A Cochrane review of supervised maintenance programmes showed clinically important improvements in HRQOL with maintenance intervention but no significant differences in exercise capacity.(170) In contrast, the long-term efficacy of PR with home-based or low frequency maintenance programmes showed improved maintenance of exercise capacity but no differences in HRQOL.(171)

- 676 Further studies are needed to explore the optimal frequency and duration of supervised and unsupervised
- 677 maintenance programmes, and the cost-effectiveness of such programmes compared with alternative
- 678 approaches (e.g. repeated PR offers).

Study	Number	Review question	Results
	of trials		
Malaguti, 2021 (170)	21 RCTs	Supervised maintenance programmes following pulmonary rehabilitation compared to usual care for COPD.	Supervised maintenance programmes not associated with increased adverse events, may improve health-related quality of life, and could improve exercise capacity at 6-12 months. Strength of evidence was limited (high risk of bias and small sample size).
lmamura, 2020 (171)	7 RCTs	Long-term efficacy of pulmonary rehabilitation with home-based or low frequent maintenance programs in COPD patients compared to those who had no maintenance programme.	PR with maintenance significantly improved 6MWD, but not HRQOL was observed.
Jenkins, 2018 (174)	8 RCTs	Efficacy of supervised maintenance exercise programmes following pulmonary rehabilitation compared to usual care on health care use.	Supervised maintenance exercise led to clinically important reduction in the rate of respiratory-cause hospital, overall risk of an exacerbation and mortality).
Busby, 2014 (175)	8 RCTs	Review of existing maintenance interventions following pulmonary rehabilitation	Most studies showed initial positive intervention effects, which declined to non- significance within 3-12 months after completion of maintenance.

### 679 Table 7: Systematic Reviews of Maintenance PR: summary of selective studies

680

### 681 Clinical Practice Points

- Oxygen supplementation should not be routinely used as an adjunct to PR except in individuals already
   established on long-term or ambulatory oxygen therapy.
- NIV should not be routinely used as an adjunct to PR in those naïve to domiciliary NIV, but could be
   offered to those already established on domiciliary NIV.
- IMT, as an adjunct to PR, is associated with improvements in muscle function, but this has not translated
   to improvements in core outcomes.
- PA counselling should be a core component of the PR educational component. The use of pedometers
   or/and additional PA counselling as adjuncts to PR require further evaluation.
- PR programmes should deliver self-management education and advice around the importance of regular
   exercise after the PR programme has been completed. There is insufficient evidence to support the
   routine formal delivery of maintenance programmes
- 693

### 694 Research gaps

- The role of oxygen supplementation during PR in specific subgroups: severe exercise induced oxygen
   desaturation (e.g. below 80%), those who demonstrate acute physiological response to oxygen.
- 697 Understanding the role of behavioural change on physical activity promotion and maintenance of the
   698 benefits of PR.
- Optimising the frequency, duration and content of supervised and unsupervised maintenance
   programmes with concomitant assessment of cost-effectiveness.
- 701 Trials comparing maintenance interventions with repeated PR.
- 702
- 703 704

### References

1. Bolton CE, Bevan-Smith EF, Blakey JD, Crowe P, Elkin SL, Garrod R, et al. British Thoracic Society guideline on pulmonary rehabilitation in adults. Thorax. 2013;68 Suppl 2:ii1-30.

2. McCarthy B, Casey D, Devane D, Murphy K, Murphy E, Lacasse Y. Pulmonary rehabilitation for chronic obstructive pulmonary disease. Cochrane Database Syst Rev. 2015;2(2):CD003793.

3. Hopkinson NS, Molyneux A, Pink J, Harrisingh MC. Chronic obstructive pulmonary disease: diagnosis and management: summary of updated NICE guidance. Bmj. 2019;366:I4486.

4. Casaburi R. Pulmonary Rehabilitation: Where We've Succeeded and Where We've Failed. Copd. 2018;15(3):219-22.

5. Holland AE, Cox NS, Houchen-Wolloff L, Rochester CL, Garvey C, ZuWallack R, et al. Defining Modern Pulmonary Rehabilitation. An Official American Thoracic Society Workshop Report. Ann Am Thorac Soc. 2021;18(5):e12-e29.

6. National COPD Audit Programme: Pulmonary rehabilitation workstream. Pulmonary Rehabilitation: Time to breathe better. National Chronic Obstructive Pulmonary Disease (COPD) Audit programme: Resources and organisation of Pulmonary Rehabilitation services in England and Wales 2015. 2015.

7. Stone PW, Hickman K, Steiner MC, Roberts CM, Quint JK, Singh SJ. Predictors of Referral to Pulmonary Rehabilitation from UK Primary Care. Int J Chron Obstruct Pulmon Dis. 2020;15:2941-52.

8. Steiner MC, Lowe D, Beckford K, Blakey J, Bolton CE, Elkin S, et al. Socioeconomic deprivation and the outcome of pulmonary rehabilitation in England and Wales. Thorax. 2017;72(6):530-7.

9. Perera ACH, Jayamaha AR, Jones AV, Yusuf ZK, Wijayasiri K, Amarasekara T, et al. Developing Appropriate Pulmonary Rehabilitation Services in Sri Lanka: Assessment of People Living with COPD and Healthcare Providers in Urban and Semi Urban Areas in Sri Lanka. Int J Chron Obstruct Pulmon Dis. 2022;17:631-41.

10. Polgar O, Aljishi M, Barker RE, Patel S, Walsh JA, Kon SS, et al. Digital habits of PR service-users: Implications for home-based interventions during the COVID-19 pandemic. Chron Respir Dis. 2020;17:1479973120936685.

11. Polgar O, Patel S, Walsh JA, Barker RE, Ingram KA, Kon SS, et al. Digital habits of pulmonary rehabilitation service-users following the COVID-19 pandemic. Chron Respir Dis. 2022;19:14799731221075647.

12. Early F, Wellwood I, Kuhn I, Deaton C, Fuld J. Interventions to increase referral and uptake to pulmonary rehabilitation in people with COPD: a systematic review. Int J Chron Obstruct Pulmon Dis. 2018;13:3571-86.

13. Jones AW, Taylor A, Gowler H, O'Kelly N, Ghosh S, Bridle C. Systematic review of interventions to improve patient uptake and completion of pulmonary rehabilitation in COPD. ERJ Open Res. 2017;3(1).

14. Cox NS, Oliveira CC, Lahham A, Holland AE. Pulmonary rehabilitation referral and participation are commonly influenced by environment, knowledge, and beliefs about consequences: a systematic review using the Theoretical Domains Framework. J Physiother. 2017;63(2):84-93.

15. National Asthma and COPD Audit Programme. Pulmonary rehabilitation clinical and organisational audits 2019. 2020 [Available from: <u>https://www.nacap.org.uk/nacap/welcome.nsf/reportsPR.html</u>.

16. Sami R, Salehi K, Hashemi M, Atashi V. Exploring the barriers to pulmonary rehabilitation for patients with chronic obstructive pulmonary disease: a qualitative study. BMC Health Serv Res. 2021;21(1):828.

17. Keating A, Lee A, Holland AE. What prevents people with chronic obstructive pulmonary disease from attending pulmonary rehabilitation? A systematic review. Chron Respir Dis. 2011;8(2):89-99.

18. Graves J, Sandrey V, Graves T, Smith DL. Effectiveness of a group opt-in session on uptake and graduation rates for pulmonary rehabilitation. Chron Respir Dis. 2010;7(3):159-64.

19. Harris M, Smith BJ, Veale AJ, Esterman A, Frith PA, Selim P. Providing reviews of evidence to COPD patients: controlled prospective 12-month trial. Chron Respir Dis. 2009;6(3):165-73.

20. Zwar NA, Hermiz O, Comino E, Middleton S, Vagholkar S, Xuan W, et al. Care of patients with a diagnosis of chronic obstructive pulmonary disease: a cluster randomised controlled trial. Med J Aust. 2012;197(7):394-8.

21. Barker RE, Jones SE, Banya W, Fleming S, Kon SSC, Clarke SF, et al. The Effects of a Video Intervention on Posthospitalization Pulmonary Rehabilitation Uptake. A Randomized Controlled Trial. Am J Respir Crit Care Med. 2020;201(12):1517-24.

22. White P, Gilworth G, Lewin S, Hogg L, Tuffnell R, Taylor SJC, et al. Improving uptake and completion of pulmonary rehabilitation in COPD with lay health workers: feasibility of a clinical trial. Int J Chron Obstruct Pulmon Dis. 2019;14:631-43.

23. Morgan AD, Rothnie KJ, Bhaskaran K, Smeeth L, Quint JK. Chronic obstructive pulmonary disease and the risk of 12 cardiovascular diseases: a population-based study using UK primary care data. Thorax. 2018;73(9):877-9.

24. Continuous or nocturnal oxygen therapy in hypoxemic chronic obstructive lung disease: a clinical trial. Nocturnal Oxygen Therapy Trial Group. Ann Intern Med. 1980;93(3):391-8.

25. Long term domiciliary oxygen therapy in chronic hypoxic cor pulmonale complicating chronic bronchitis and emphysema. Report of the Medical Research Council Working Party. Lancet. 1981;1(8222):681-6.

26. Kwan HY, Maddocks M, Nolan CM, Jones SE, Patel S, Barker RE, et al. The prognostic significance of weight loss in chronic obstructive pulmonary disease-related cachexia: a prospective cohort study. J Cachexia Sarcopenia Muscle. 2019;10(6):1330-8.

27. Celli BR, Cote CG, Marin JM, Casanova C, Montes de Oca M, Mendez RA, et al. The body-mass index, airflow obstruction, dyspnea, and exercise capacity index in chronic obstructive pulmonary disease. N Engl J Med. 2004;350(10):1005-12.

28. Brigham EP, Anderson JA, Brook RD, Calverley PMA, Celli BR, Cowans NJ, et al. Challenging the obesity paradox: extreme obesity and COPD mortality in the SUMMIT trial. ERJ Open Res. 2021;7(3).

29. Hanlon P, Lewsey J, Quint JK, Jani BD, Nicholl BI, McAllister DA, et al. Frailty in COPD: an analysis of prevalence and clinical impact using UK Biobank. BMJ Open Respir Res. 2022;9(1).

30. Walsh JA, Barker RE, Kon SSC, Jones SE, Banya W, Nolan CM, et al. Gait speed and adverse outcomes following hospitalised exacerbation of COPD. Eur Respir J. 2021;58(5).

31. Singer JP, Diamond JM, Gries CJ, McDonnough J, Blanc PD, Shah R, et al. Frailty Phenotypes, Disability, and Outcomes in Adult Candidates for Lung Transplantation. Am J Respir Crit Care Med. 2015;192(11):1325-34.

32. Maddocks M, Kon SS, Canavan JL, Jones SE, Nolan CM, Labey A, et al. Physical frailty and pulmonary rehabilitation in COPD: a prospective cohort study. Thorax. 2016;71(11):988-95.

33. Wynne SC, Patel S, Barker RE, Jones SE, Walsh JA, Kon SS, et al. Anxiety and depression in bronchiectasis: Response to pulmonary rehabilitation and minimal clinically important difference of the Hospital Anxiety and Depression Scale. Chron Respir Dis. 2020;17:1479973120933292.

34. Gordon CS, Waller JW, Cook RM, Cavalera SL, Lim WT, Osadnik CR. Effect of Pulmonary Rehabilitation on Symptoms of Anxiety and Depression in COPD: A Systematic Review and Meta-Analysis. Chest. 2019;156(1):80-91.

35. Leander M, Lampa E, Rask-Andersen A, Franklin K, Gislason T, Oudin A, et al. Impact of anxiety and depression on respiratory symptoms. Respir Med. 2014;108(11):1594-600.

36. McNamara RJ, Kearns R, Dennis SM, M FH, Gardner K, McDonald J. Knowledge, Skill, and Confidence in People Attending Pulmonary Rehabilitation: A Cross-Sectional Analysis of the Effects and Determinants of Patient Activation. J Patient Exp. 2019;6(2):117-25.

37. Volpato E, Toniolo S, Pagnini F, Banfi P. The Relationship Between Anxiety, Depression and Treatment Adherence in Chronic Obstructive Pulmonary Disease: A Systematic Review. Int J Chron Obstruct Pulmon Dis. 2021;16:2001-21.

38. Roberts NJ, Kidd L, Kirkwood K, Cross J, Partridge MR. A systematic review of the content and delivery of education in pulmonary rehabilitation programmes. Respir Med. 2018;145:161-81.

39. Jones RC, Wang X, Harding S, Bott J, Hyland M. Educational impact of pulmonary rehabilitation: Lung Information Needs Questionnaire. Respir Med. 2008;102(10):1439-45.

40. White R, Walker P, Roberts S, Kalisky S, White P. Bristol COPD Knowledge Questionnaire (BCKQ): testing what we teach patients about COPD. Chron Respir Dis. 2006;3(3):123-31.

41. Hedman E, Ljótsson B, Blom K, El Alaoui S, Kraepelien M, Rück C, et al. Telephone versus internet administration of self-report measures of social anxiety, depressive symptoms, and insomnia: psychometric evaluation of a method to reduce the impact of missing data. J Med Internet Res. 2013;15(10):e229.

42. Rocha V, Jácome C, Martins V, Marques A. Are in Person and Telephone Interviews Equivalent Modes of Administrating the CAT, the FACIT-FS and the SGRQ in People With COPD? Frontiers in Rehabilitation Sciences. 2021;2.

43. Holland AE, Malaguti C, Hoffman M, Lahham A, Burge AT, Dowman L, et al. Home-based or remote exercise testing in chronic respiratory disease, during the COVID-19 pandemic and beyond: A rapid review. Chron Respir Dis. 2020;17:1479973120952418.

44. Salvi D, Poffley E, Orchard E, Tarassenko L. The Mobile-Based 6-Minute Walk Test: Usability Study and Algorithm Development and Validation. JMIR Mhealth Uhealth. 2020;8(1):e13756.

45. Brooks D, Solway S, Weinacht K, Wang D, Thomas S. Comparison between an indoor and an outdoor 6-minute walk test among individuals with chronic obstructive pulmonary disease. Arch Phys Med Rehabil. 2003;84(6):873-6.

46. Holland AE, Spruit MA, Troosters T, Puhan MA, Pepin V, Saey D, et al. An official European Respiratory Society/American Thoracic Society technical standard: field walking tests in chronic respiratory disease. Eur Respir J. 2014;44(6):1428-46.

47. Kon SS, Canavan JL, Nolan CM, Clark AL, Jones SE, Cullinan P, et al. The 4-metre gait speed in COPD: responsiveness and minimal clinically important difference. Eur Respir J. 2014;43(5):1298-305.

48. Kon SS, Patel MS, Canavan JL, Clark AL, Jones SE, Nolan CM, et al. Reliability and validity of 4-metre gait speed in COPD. Eur Respir J. 2013;42(2):333-40.

49. Nolan CM, Maddocks M, Maher TM, Canavan JL, Jones SE, Barker RE, et al. Phenotypic characteristics associated with slow gait speed in idiopathic pulmonary fibrosis. Respirology. 2018;23(5):498-506.

50. Jones SE, Kon SS, Canavan JL, Patel MS, Clark AL, Nolan CM, et al. The five-repetition sit-to-stand test as a functional outcome measure in COPD. Thorax. 2013;68(11):1015-20.

51. Zhang Q, Li YX, Li XL, Yin Y, Li RL, Qiao X, et al. A comparative study of the five-repetition sit-tostand test and the 30-second sit-to-stand test to assess exercise tolerance in COPD patients. Int J Chron Obstruct Pulmon Dis. 2018;13:2833-9.

52. Hansen H, Beyer N, Frølich A, Godtfredsen N, Bieler T. Intra- and inter-rater reproducibility of the 6minute walk test and the 30-second sit-to-stand test in patients with severe and very severe COPD. Int J Chron Obstruct Pulmon Dis. 2018;13:3447-57.

53. Crook S, Buesching G, Schultz K, Lehbert N, Jelusic D, Keusch S, et al. A multicentre validation of the 1-minute sit-to-stand test in patients with COPD. European Respiratory Journal. 2017.

54. da Costa JN, Arcuri JF, Gonçalves IL, Davi SF, Pessoa BV, Jamami M, et al. Reproducibility of cadence-free 6-minute step test in subjects with COPD. Respir Care. 2014;59(4):538-42.

55. Pessoa BV, Arcuri JF, Labadessa IG, Costa JN, Sentanin AC, Di Lorenzo VA. Validity of the six-minute step test of free cadence in patients with chronic obstructive pulmonary disease. Braz J Phys Ther. 2014;18(3):228-36.

56. Finney LJ, Doughty R, Lovage S, Spurr L, Mehta B, Kemp SV, et al. Lung Function Deficits and Symptom Burden in Survivors of COVID-19 Requiring Mechanical Ventilation. Ann Am Thorac Soc. 2021;18(10):1740-3.

57. Patel S, Jones SE, Walsh JA, Barker RE, Polgar O, Maddocks M, et al. The Six-Minute Step Test as an Exercise Outcome in COPD. Ann Am Thorac Soc. 2022.

58. Mesquita R, Wilke S, Smid DE, Janssen DJ, Franssen FM, Probst VS, et al. Measurement properties of the Timed Up & Go test in patients with COPD. Chron Respir Dis. 2016.

59. Houchen-Wolloff L, Daynes E, Watt A, Chaplin E, Gardiner N, Singh S. Which functional outcome measures can we use as a surrogate for exercise capacity during remote cardiopulmonary rehabilitation assessments? A rapid narrative review. ERJ Open Res. 2020;6(4).

60. Bisca GW, Morita AA, Hernandes NA, Probst VS, Pitta F. Simple Lower Limb Functional Tests in Patients With Chronic Obstructive Pulmonary Disease: A Systematic Review. Arch Phys Med Rehabil. 2015;96(12):2221-30.

61. Bui KL, Nyberg A, Maltais F, Saey D. Functional Tests in Chronic Obstructive Pulmonary Disease, Part 2: Measurement Properties. Ann Am Thorac Soc. 2017;14(5):785-94.

62. Demeyer H, Mohan D, Burtin C, Vaes AW, Heasley M, Bowler RP, et al. Objectively Measured Physical Activity in Patients with COPD: Recommendations from an International Task Force on Physical Activity. Chronic Obstr Pulm Dis. 2021;8(4):528-50.

63. Garfield BE, Canavan JL, Smith CJ, Ingram KA, Fowler RP, Clark AL, et al. Stanford Seven-Day Physical Activity Recall questionnaire in COPD. Eur Respir J. 2012;40(2):356-62.

64. Burtin C, Mohan D, Troosters T, Watz H, Hopkinson NS, Garcia-Aymerich J, et al. Objectively Measured Physical Activity as a COPD Clinical Trial Outcome. Chest. 2021;160(6):2080-100.

65. Feng Z, Wang J, Xie Y, Li J. Effects of exercise-based pulmonary rehabilitation on adults with asthma: a systematic review and meta-analysis. Respir Res. 2021;22(1):33.

66. Dowman L, Hill CJ, May A, Holland AE. Pulmonary rehabilitation for interstitial lung disease. Cochrane Database Syst Rev. 2021;2(2):Cd006322.

67. Lee AL, Hill CJ, McDonald CF, Holland AE. Pulmonary Rehabilitation in Individuals With Non-Cystic Fibrosis Bronchiectasis: A Systematic Review. Arch Phys Med Rehabil. 2017;98(4):774-82 e1.

68. Osadnik CR, Gleeson C, McDonald VM, Holland AE. Pulmonary rehabilitation versus usual care for adults with asthma. Cochrane Database Syst Rev. 2022;8(8):Cd013485.

69. Nolan CM, Polgar O, Schofield SJ, Patel S, Barker RE, Walsh JA, et al. Pulmonary Rehabilitation in Idiopathic Pulmonary Fibrosis and COPD: A Propensity-Matched Real-World Study. Chest. 2022;161(3):728-37.

70. Patel S, Cole AD, Nolan CM, Barker RE, Jones SE, Kon S, et al. Pulmonary rehabilitation in bronchiectasis: a propensity-matched study. Eur Respir J. 2019;53(1).

71. Eichenberger PA, Diener SN, Kofmehl R, Spengler CM. Effects of exercise training on airway hyperreactivity in asthma: a systematic review and meta-analysis. Sports Med. 2013;43(11):1157-70.

72. Hill AT, Sullivan AL, Chalmers JD, De Soyza A, Elborn SJ, Floto AR, et al. British Thoracic Society Guideline for bronchiectasis in adults. Thorax. 2019;74(Suppl 1):1-69.

73. Sobala R, Brooks K, Davison J, Lumb J, De Soyza A. Family case studies: absence of Pseudomonas aeruginosa transmission in bronchiectasis. ERJ Open Res. 2022;8(4).

74. Delivering rehabilitation to patients surviving COVID-19 using an adapted pulmonary rehabilitation approach – BTS guidance 2020 [Available from: <u>https://www.brit-thoracic.org.uk/document-library/quality-improvement/covid-19/pulmonary-rehabilitation-for-covid-19-patients/</u>.

75. Al Chikhanie Y, Veale D, Schoeffler M, Pépin JL, Verges S, Hérengt F. Effectiveness of pulmonary rehabilitation in COVID-19 respiratory failure patients post-ICU. Respir Physiol Neurobiol. 2021;287:103639.

76. Daynes E, Gerlis C, Chaplin E, Gardiner N, Singh SJ. Early experiences of rehabilitation for individuals post-COVID to improve fatigue, breathlessness exercise capacity and cognition - A cohort study. Chron Respir Dis. 2021;18:14799731211015691.

77. Nopp S, Moik F, Klok FA, Gattinger D, Petrovic M, Vonbank K, et al. Outpatient Pulmonary Rehabilitation in Patients with Long COVID Improves Exercise Capacity, Functional Status, Dyspnea, Fatigue, and Quality of Life. Respiration. 2022;101(6):593-601.

78. Gloeckl R, Leitl D, Jarosch I, Schneeberger T, Nell C, Stenzel N, et al. Benefits of pulmonary rehabilitation in COVID-19: a prospective observational cohort study. ERJ Open Res. 2021;7(2).

79. Spielmanns M, Pekacka-Egli AM, Schoendorf S, Windisch W, Hermann M. Effects of a Comprehensive Pulmonary Rehabilitation in Severe Post-COVID-19 Patients. Int J Environ Res Public Health. 2021;18(5).

80. Zampogna E, Paneroni M, Belli S, Aliani M, Gandolfo A, Visca D, et al. Pulmonary Rehabilitation in Patients Recovering from COVID-19. Respiration. 2021;100(5):416-22.

81. Evans RA, McAuley H, Harrison EM, Shikotra A, Singapuri A, Sereno M, et al. Physical, cognitive, and mental health impacts of COVID-19 after hospitalisation (PHOSP-COVID): a UK multicentre, prospective cohort study. Lancet Respir Med. 2021;9(11):1275-87.

82. Group P-CC. Clinical characteristics with inflammation profiling of long COVID and association with 1-year recovery following hospitalisation in the UK: a prospective observational study. Lancet Respir Med. 2022.

83. Li J, Xia W, Zhan C, Liu S, Yin Z, Wang J, et al. A telerehabilitation programme in post-discharge COVID-19 patients (TERECO): a randomised controlled trial. Thorax. 2022;77(7):697-706.

84. Remy-Jardin M, Duthoit L, Perez T, Felloni P, Faivre JB, Fry S, et al. Assessment of pulmonary arterial circulation 3 months after hospitalization for SARS-CoV-2 pneumonia: Dual-energy CT (DECT) angiographic study in 55 patients. EClinicalMedicine. 2021;34:100778.

85. Puntmann VO, Martin S, Shchendrygina A, Hoffmann J, Ka MM, Giokoglu E, et al. Long-term cardiac pathology in individuals with mild initial COVID-19 illness. Nat Med. 2022.

86. Wright J, Astill SL, Sivan M. The Relationship between Physical Activity and Long COVID: A Cross-Sectional Study. Int J Environ Res Public Health. 2022;19(9).

87. Davis JF, van Rooijen SJ, Grimmett C, West MA, Campbell AM, Awasthi R, et al. From Theory to Practice: An International Approach to Establishing Prehabilitation Programmes. Curr Anesthesiol Rep. 2022;12(1):129-37.

88. Himbert C, Klossner N, Coletta AM, Barnes CA, Wiskemann J, LaStayo PC, et al. Exercise and lung cancer surgery: A systematic review of randomized-controlled trials. Crit Rev Oncol Hematol. 2020;156:103086.

89. Cavalheri V, Burtin C, Formico VR, Nonoyama ML, Jenkins S, Spruit MA, et al. Exercise training undertaken by people within 12 months of lung resection for non-small cell lung cancer. Cochrane Database Syst Rev. 2019;6(6):Cd009955.

90. Cavalheri V, Jenkins S, Hill K. Physiotherapy practice patterns for patients undergoing surgery for lung cancer: a survey of hospitals in Australia and New Zealand. Intern Med J. 2013;43(4):394-401.

91. NICE Evidence Reviews Collection. Referral criteria for lung volume reduction procedures, bullectomy or lung transplantation: Chronic obstructive pulmonary disease in over 16s: diagnosis and management: Evidence review G. London: National Institute for Health and Care Excellence (NICE)

### Copyright © NICE 2018.; 2018.

92. Fishman A, Martinez F, Naunheim K, Piantadosi S, Wise R, Ries A, et al. A randomized trial comparing lung-volume-reduction surgery with medical therapy for severe emphysema. N Engl J Med. 2003;348(21):2059-73.

93. Ries AL, Make BJ, Lee SM, Krasna MJ, Bartels M, Crouch R, et al. The effects of pulmonary rehabilitation in the national emphysema treatment trial. Chest. 2005;128(6):3799-809.

94. McNulty W, Jordan S, Hopkinson NS. Attitudes and access to lung volume reduction surgery for COPD: a survey by the British Thoracic Society. BMJ Open Respir Res. 2014;1(1):e000023.

95. Buttery SC, Lewis A, Kemp SV, Banya W, Quint JK, Steiner MC, et al. Lung volume reduction eligibility in patients with COPD completing pulmonary rehabilitation: results from the UK National Asthma and COPD Audit Programme. BMJ Open. 2020;10(11):e040942.

96. Gutierrez-Arias R, Martinez-Zapata MJ, Gaete-Mahn MC, Osorio D, Bustos L, Melo Tanner J, et al. Exercise training for adult lung transplant recipients. Cochrane Database Syst Rev. 2021;7(7):CD012307.

97. Mesquita R, Vanfleteren LE, Franssen FM, Sarv J, Taib Z, Groenen MT, et al. Objectively identified comorbidities in COPD: impact on pulmonary rehabilitation outcomes. Eur Respir J. 2015;46(2):545-8.
98. Taylor RS, Walker S, Ciani O, Warren F, Smart NA, Piepoli M, et al. Exercise-based cardiac

rehabilitation for chronic heart failure: the EXTRAMATCH II individual participant data meta-analysis. Health Technol Assess. 2019;23(25):1-98.

99. Chaplin E, Ward S, Daynes E, Bourne C, Stenson A, Watt A, et al. Integrating patients with chronic respiratory disease and heart failure into a combined breathlessness rehabilitation programme: a service redesign and pilot evaluation. BMJ Open Respir Res. 2021;8(1).

100. Evans RA, Singh SJ, Collier R, Loke I, Steiner MC, Morgan MD. Generic, symptom based, exercise rehabilitation; integrating patients with COPD and heart failure. Respir Med. 2010;104(10):1473-81.

101. Yan L, Shi W, Liu Z, Zhao Z, Luo Q, Zhao Q, et al. The benefit of exercise-based rehabilitation programs in patients with pulmonary hypertension: a systematic review and meta-analysis of randomized controlled trials. Pulm Circ. 2021;11(2):20458940211007810.

102. Morris NR, Kermeen FD, Holland AE. Exercise-based rehabilitation programmes for pulmonary hypertension. Cochrane Database Syst Rev. 2017;1:CD011285.

103. Grünig E, Eichstaedt C, Barberà JA, Benjamin N, Blanco I, Bossone E, et al. ERS statement on exercise training and rehabilitation in patients with severe chronic pulmonary hypertension. Eur Respir J. 2019;53(2).

104. Kon SS, Jones SE, Schofield SJ, Banya W, Dickson MJ, Canavan JL, et al. Gait speed and readmission following hospitalisation for acute exacerbations of COPD: a prospective study. Thorax. 2015;70(12):1131-7.

105. Man WD, Kon SS, Maddocks M. Rehabilitation after an exacerbation of chronic respiratory disease. BMJ. 2014;349:g4370.

106. Kon SS, Canavan JL, Jones SE, Nolan CM, Clark AL, Dickson MJ, et al. Minimum clinically important difference for the COPD Assessment Test: a prospective analysis. Lancet Respir Med. 2014;2(3):195-203.

107. Steer J, Gibson GJ, Bourke SC. Longitudinal change in quality of life following hospitalisation for acute exacerbations of COPD. BMJ Open Respir Res. 2015;2(1):e000069.

108. Pitta F, Troosters T, Probst VS, Spruit MA, Decramer M, Gosselink R. Physical activity and hospitalization for exacerbation of COPD. Chest. 2006;129(3):536-44.

109. McAuley HJC, Harvey-Dunstan TC, Craner M, Richardson M, Singh SJ, Steiner MC, et al. Longitudinal changes to quadriceps thickness demonstrate acute sarcopenia following admission to hospital for an exacerbation of chronic respiratory disease. Thorax. 2021;76(7):726-8.

110. Steer J, Gibson GJ, Bourke SC. Predicting outcomes following hospitalization for acute exacerbations of COPD. Qjm. 2010;103(11):817-29.

Puhan MA, Gimeno-Santos E, Cates CJ, Troosters T. Pulmonary rehabilitation following exacerbations of chronic obstructive pulmonary disease. Cochrane Database Syst Rev. 2016;12:CD005305.
Seymour JM, Moore L, Jolley CJ, Ward K, Creasey J, Steier JS, et al. Outpatient pulmonary rehabilitation following acute exacerbations of COPD. Thorax. 2010;65(5):423-8.

113. Ko FW, Cheung NK, Rainer TH, Lum C, Wong I, Hui DS. Comprehensive care programme for patients with chronic obstructive pulmonary disease: a randomised controlled trial. Thorax. 2017;72(2):122-8.

114. Lindenauer PK, Stefan MS, Pekow PS, Mazor KM, Priya A, Spitzer KA, et al. Association Between Initiation of Pulmonary Rehabilitation After Hospitalization for COPD and 1-Year Survival Among Medicare Beneficiaries. JAMA. 2020;323(18):1813-23.

115. Greening NJ, Williams JE, Hussain SF, Harvey-Dunstan TC, Bankart MJ, Chaplin EJ, et al. An early rehabilitation intervention to enhance recovery during hospital admission for an exacerbation of chronic respiratory disease: randomised controlled trial. BMJ. 2014;349:g4315.

116. Cox M, O'Connor C, Biggs K, Hind D, Bortolami O, Franklin M, et al. The feasibility of early pulmonary rehabilitation and activity after COPD exacerbations: external pilot randomised controlled trial, qualitative case study and exploratory economic evaluation. Health Technol Assess. 2018;22(11):1-204.

117. Man WD, Polkey MI, Donaldson N, Gray BJ, Moxham J. Community pulmonary rehabilitation after hospitalisation for acute exacerbations of chronic obstructive pulmonary disease: randomised controlled study. BMJ. 2004;329(7476):1209.

118. Güell-Rous MR, Morante-Vélez F, Flotats-Farré G, Paz-Del Río LD, Closa-Rusinés C, Ouchi-Vernet D, et al. Timing of Pulmonary Rehabilitation in Readmitted Patients with Severe Chronic Obstructive Pulmonary Disease: A Randomized Clinical Trial. Copd. 2021;18(1):26-34.

119. Wageck B, Cox NS, Lee JYT, Romero L, Holland AE. Characteristics of Pulmonary Rehabilitation Programs Following an Exacerbation of Chronic Obstructive Pulmonary Disease: A SYSTEMATIC REVIEW. J Cardiopulm Rehabil Prev. 2021;41(2):78-87.

120. Jones SE, Green SA, Clark AL, Dickson MJ, Nolan AM, Moloney C, et al. Pulmonary rehabilitation following hospitalisation for acute exacerbation of COPD: referrals, uptake and adherence. Thorax. 2014;69(2):181-2.

121. Kjærgaard JL, Juhl CB, Lange P, Wilcke JT. Early pulmonary rehabilitation after acute exacerbation of COPD: a randomised controlled trial. ERJ Open Res. 2020;6(1).

122. Rochester CL, Vogiatzis I, Holland AE, Lareau SC, Marciniuk DD, Puhan MA, et al. An Official American Thoracic Society/European Respiratory Society Policy Statement: Enhancing Implementation, Use, and Delivery of Pulmonary Rehabilitation. Am J Respir Crit Care Med. 2015;192(11):1373-86.

123. Waterhouse JC, Walters SJ, Oluboyede Y, Lawson RA. A randomised 2 x 2 trial of community versus hospital pulmonary rehabilitation, followed by telephone or conventional follow-up. Health Technol Assess. 2010;14(6):i-v, vii-xi, 1-140.

124. Patel S, Palmer MD, Nolan CM, Barker RE, Walsh JA, Wynne SC, et al. Supervised pulmonary rehabilitation using minimal or specialist exercise equipment in COPD: a propensity-matched analysis. Thorax. 2021;76(3):264-71.

Mitchell KE, Johnson-Warrington V, Apps LD, Bankart J, Sewell L, Williams JE, et al. A self-management programme for COPD: a randomised controlled trial. Eur Respir J. 2014;44(6):1538-47.
Holland AE, Mahal A, Hill CJ, Lee AL, Burge AT, Cox NS, et al. Home-based rehabilitation for COPD using minimal resources: a randomised, controlled equivalence trial. Thorax. 2017;72(1):57-65.

127. Horton EJ, Mitchell KE, Johnson-Warrington V, Apps LD, Sewell L, Morgan M, et al. Comparison of a structured home-based rehabilitation programme with conventional supervised pulmonary rehabilitation: a randomised non-inferiority trial. Thorax. 2018;73(1):29-36.

128. Maltais F, Bourbeau J, Shapiro S, Lacasse Y, Perrault H, Baltzan M, et al. Effects of home-based pulmonary rehabilitation in patients with chronic obstructive pulmonary disease: a randomized trial. Ann Intern Med. 2008;149(12):869-78.

Nolan CM, Kaliaraju D, Jones SE, Patel S, Barker R, Walsh JA, et al. Home versus outpatient pulmonary rehabilitation in COPD: a propensity-matched cohort study. Thorax. 2019;74(10):996-8.
Jones S, Man WD, Gao W, Higginson IJ, Wilcock A, Maddocks M. Neuromuscular electrical stimulation for muscle weakness in adults with advanced disease. Cochrane Database Syst Rev. 2016;10:CD009419.

131. Maddocks M, Nolan CM, Man WD, Polkey MI, Hart N, Gao W, et al. Neuromuscular electrical stimulation to improve exercise capacity in patients with severe COPD: a randomised double-blind, placebo-controlled trial. Lancet Respir Med. 2016;4(1):27-36.

132. Nolan CM, Patel S, Barker RE, Walsh JA, Polgar O, Maddocks M, et al. Muscle stimulation in advanced idiopathic pulmonary fibrosis: a randomised placebo-controlled feasibility study. BMJ Open. 2021;11(6):e048808.

133. Bourne S, DeVos R, North M, Chauhan A, Green B, Brown T, et al. Online versus face-to-face pulmonary rehabilitation for patients with chronic obstructive pulmonary disease: randomised controlled trial. BMJ Open. 2017;7(7):e014580.

134. Chaplin E, Hewitt S, Apps L, Bankart J, Pulikottil-Jacob R, Boyce S, et al. Interactive web-based pulmonary rehabilitation programme: a randomised controlled feasibility trial. BMJ Open. 2017;7(3):e013682.

135. Tsai LL, McNamara RJ, Moddel C, Alison JA, McKenzie DK, McKeough ZJ. Home-based telerehabilitation via real-time videoconferencing improves endurance exercise capacity in patients with COPD: The randomized controlled TeleR Study. Respirology. 2017;22(4):699-707.

136. Cox NS, McDonald CF, Mahal A, Alison JA, Wootton R, Hill CJ, et al. Telerehabilitation for chronic respiratory disease: a randomised controlled equivalence trial. Thorax. 2022;77(7):643-51.

137. Hansen H, Bieler T, Beyer N, Kallemose T, Wilcke JT, Ostergaard LM, et al. Supervised pulmonary tele-rehabilitation versus pulmonary rehabilitation in severe COPD: a randomised multicentre trial. Thorax. 2020;75(5):413-21.

138. Alwakeel AJ, Sicondolfo A, Robitaille C, Bourbeau J, Saad N. The Accessibility, Feasibility, and Safety of a Standardized Community-based Tele-Pulmonary Rehab Program for Chronic Obstructive Pulmonary Disease: A 3-Year Real-World Prospective Study. Ann Am Thorac Soc. 2022;19(1):39-47.

139. Knox L, Dunning M, Davies CA, Mills-Bennet R, Sion TW, Phipps K, et al. Safety, feasibility, and effectiveness of virtual pulmonary rehabilitation in the real world. Int J Chron Obstruct Pulmon Dis. 2019;14:775-80.

140. Jung T, Moorhouse N, Shi X, Amin MF. A Virtual Reality-Supported Intervention for Pulmonary Rehabilitation of Patients With Chronic Obstructive Pulmonary Disease: Mixed Methods Study. J Med Internet Res. 2020;22(7):e14178.

141. Condon C, Lam WT, Mosley C, Gough S. A systematic review and meta-analysis of the effectiveness of virtual reality as an exercise intervention for individuals with a respiratory condition. Adv Simul (Lond). 2020;5(1):33.

142. Liu XC, Pan L, Hu Q, Dong WP, Yan JH, Dong L. Effects of yoga training in patients with chronic obstructive pulmonary disease: a systematic review and meta-analysis. J Thorac Dis. 2014;6(6):795-802.

143. Wu W, Liu X, Wang L, Wang Z, Hu J, Yan J. Effects of Tai Chi on exercise capacity and health-related quality of life in patients with chronic obstructive pulmonary disease: a systematic review and metaanalysis. Int J Chron Obstruct Pulmon Dis. 2014;9:1253-63.

144. Gendron LM, Nyberg A, Saey D, Maltais F, Lacasse Y. Active mind-body movement therapies as an adjunct to or in comparison with pulmonary rehabilitation for people with chronic obstructive pulmonary disease. Cochrane Database Syst Rev. 2018;10(10):Cd012290.

145. Polkey MI, Qiu ZH, Zhou L, Zhu MD, Wu YX, Chen YY, et al. Tai Chi and Pulmonary Rehabilitation Compared for Treatment-Naive Patients With COPD: A Randomized Controlled Trial. Chest. 2018;153(5):1116-24. 146. Sahasrabudhe SD, Orme MW, Jones AV, Tillu G, Salvi SS, Singh SJ. Potential for integrating yoga within pulmonary rehabilitation and recommendations of reporting framework. BMJ Open Respir Res. 2021;8(1).

147. Cox NS, Dal Corso S, Hansen H, McDonald CF, Hill CJ, Zanaboni P, et al. Telerehabilitation for chronic respiratory disease. Cochrane Database Syst Rev. 2021;1(1):CD013040.

148. Somfay A, Porszasz J, Lee SM, Casaburi R. Dose-response effect of oxygen on hyperinflation and exercise endurance in nonhypoxaemic COPD patients. Eur Respir J. 2001;18(1):77-84.

149. Louvaris Z, Vogiatzis I, Aliverti A, Habazettl H, Wagner H, Wagner P, et al. Blood flow does not redistribute from respiratory to leg muscles during exercise breathing heliox or oxygen in COPD. J Appl Physiol (1985). 2014;117(3):267-76.

150. O'Donnell DE, D'Arsigny C, Webb KA. Effects of hyperoxia on ventilatory limitation during exercise in advanced chronic obstructive pulmonary disease. Am J Respir Crit Care Med. 2001;163(4):892-8.

151. Alison JA, McKeough ZJ, Leung RWM, Holland AE, Hill K, Morris NR, et al. Oxygen compared to air during exercise training in COPD with exercise-induced desaturation. Eur Respir J. 2019;53(5).

152. Walsh JA, Maddocks M, Man WD. Supplemental oxygen during exercise training in COPD: full of hot air? Eur Respir J. 2019;53(5).

153. Menadue C, Piper AJ, van 't Hul AJ, Wong KK. Non-invasive ventilation during exercise training for people with chronic obstructive pulmonary disease. Cochrane Database Syst Rev. 2014(5):Cd007714.

154. Ricci C, Terzoni S, Gaeta M, Sorgente A, Destrebecq A, Gigliotti F. Physical training and noninvasive ventilation in COPD patients: a meta-analysis. Respir Care. 2014;59(5):709-17.

155. Dyer F, Flude L, Bazari F, Jolley C, Englebretsen C, Lai D, et al. Non-invasive ventilation (NIV) as an aid to rehabilitation in acute respiratory disease. BMC Pulm Med. 2011;11:58.

156. Vitacca M, Kaymaz D, Lanini B, Vagheggini G, Ergun P, Gigliotti F, et al. Non-invasive ventilation during cycle exercise training in patients with chronic respiratory failure on long-term ventilatory support: A randomized controlled trial. Respirology. 2018;23(2):182-9.

157. Charususin N, Gosselink R, Decramer M, Demeyer H, McConnell A, Saey D, et al. Randomised controlled trial of adjunctive inspiratory muscle training for patients with COPD. Thorax. 2018;73(10):942-50.

158. Schultz K, Jelusic D, Wittmann M, Kramer B, Huber V, Fuchs S, et al. Inspiratory muscle training does not improve clinical outcomes in 3-week COPD rehabilitation: results from a randomised controlled trial. Eur Respir J. 2018;51(1).

159. Beaumont M, Mialon P, Le Ber C, Le Mevel P, Peran L, Meurisse O, et al. Effects of inspiratory muscle training on dyspnoea in severe COPD patients during pulmonary rehabilitation: controlled randomised trial. Eur Respir J. 2018;51(1).

160. Zaki S, Moiz JA, Mujaddadi A, Ali MS, Talwar D. Does inspiratory muscle training provide additional benefits during pulmonary rehabilitation in people with interstitial lung disease? A randomized control trial. Physiother Theory Pract. 2022:1-11.

161. Newall C, Stockley RA, Hill SL. Exercise training and inspiratory muscle training in patients with bronchiectasis. Thorax. 2005;60(11):943-8.

162. Cindy Ng LW, Mackney J, Jenkins S, Hill K. Does exercise training change physical activity in people with COPD? A systematic review and meta-analysis. Chron Respir Dis. 2012;9(1):17-26.

163. Armstrong M, Winnard A, Chynkiamis N, Boyle S, Burtin C, Vogiatzis I. Use of pedometers as a tool to promote daily physical activity levels in patients with COPD: a systematic review and meta-analysis. Eur Respir Rev. 2019;28(154).

164. Nolan CM, Maddocks M, Canavan JL, Jones SE, Delogu V, Kaliaraju D, et al. Pedometer Step Count Targets during Pulmonary Rehabilitation in Chronic Obstructive Pulmonary Disease. A Randomized Controlled Trial. Am J Respir Crit Care Med. 2017;195(10):1344-52.

165. Cruz J, Brooks D, Marques A. Walk2Bactive: A randomised controlled trial of a physical activityfocused behavioural intervention beyond pulmonary rehabilitation in chronic obstructive pulmonary disease. Chron Respir Dis. 2016;13(1):57-66.

166. Rausch Osthoff AK, Beyer S, Gisi D, Rezek S, Schwank A, Meichtry A, et al. Effect of counselling during pulmonary rehabilitation on self-determined motivation to be physically active for people with chronic obstructive pulmonary disease: a pragmatic RCT. BMC Pulm Med. 2021;21(1):317.

167. Armstrong M, Hume E, McNeillie L, Chambers F, Wakenshaw L, Burns G, et al. Behavioural modification interventions alongside pulmonary rehabilitation improve COPD patients' experiences of physical activity. Respir Med. 2021;180:106353.

168. Griffiths TL, Burr ML, Campbell IA, Lewis-Jenkins V, Mullins J, Shiels K, et al. Results at 1 year of outpatient multidisciplinary pulmonary rehabilitation: a randomised controlled trial. Lancet. 2000;355(9201):362-8.

169. Alison JA, McKeough ZJ, Johnston K, McNamara RJ, Spencer LM, Jenkins SC, et al. Australian and New Zealand Pulmonary Rehabilitation Guidelines. Respirology. 2017;22(4):800-19.

170. Malaguti C, Dal Corso S, Janjua S, Holland AE. Supervised maintenance programmes following pulmonary rehabilitation compared to usual care for chronic obstructive pulmonary disease. Cochrane Database Syst Rev. 2021;8(8):Cd013569.

171. Imamura S, Inagaki T, Terada J, Nagashima K, Katsura H, Tatsumi K. Long-term efficacy of pulmonary rehabilitation with home-based or low frequent maintenance programs in patients with chronic obstructive pulmonary disease: a meta-analysis. Ann Palliat Med. 2020;9(5):2606-15.

172. Kornowski R, Witberg G. Acute myocarditis caused by COVID-19 disease and following COVID-19 vaccination. Open Heart. 2022;9(1).

173. Twomey R, DeMars J, Franklin K, Culos-Reed SN, Weatherald J, Wrightson JG. Chronic Fatigue and Postexertional Malaise in People Living With Long COVID: An Observational Study. Phys Ther. 2022;102(4).
174. Jenkins AR, Gowler H, Curtis F, Holden NS, Bridle C, Jones AW. Efficacy of supervised maintenance exercise following pulmonary rehabilitation on health care use: a systematic review and meta-analysis. Int J Chron Obstruct Pulmon Dis. 2018;13:257-73.

175. Busby AK, Reese RL, Simon SR. Pulmonary rehabilitation maintenance interventions: a systematic review. Am J Health Behav. 2014;38(3):321-30.