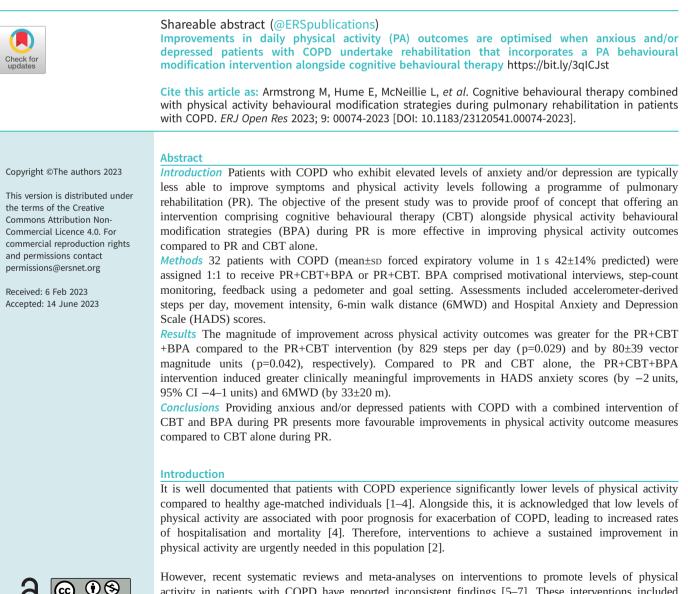


Cognitive behavioural therapy combined with physical activity behavioural modification strategies during pulmonary rehabilitation in patients with COPD

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activity in patients with COPD have reported inconsistent findings [5-7]. These interventions included pulmonary rehabilitation (PR), physical activity counselling as a standalone intervention or alongside PR, health monitoring and nocturnal noninvasive ventilation (NNIV) [5–7]. Of these interventions, physical activity counselling both as a standalone intervention and alongside PR provided the most effective intervention, especially when implemented alongside PR [5, 6]. In addition, recent research has shown that although PR is effective for improving functional capacity, it does not necessarily translate into improved physical activity outcomes unless physical activity behavioural modification strategies are incorporated [8]. Therefore, it is evident that a combination of both PR and behavioural interventions is necessary to see meaningful improvements in physical activity outcomes.

Interestingly, our earlier systematic review and meta-analysis found that patients with greater baseline physical activity levels (>4000 steps per day) showed greater improvements in steps per day compared to those with lower baseline physical activity levels (\leq 4000 steps per day), following a programme of physical activity counselling [5]. In patients with low baseline physical activity (\leq 4000 steps per day), studies that have implemented physical activity counselling alongside PR have reported an insignificant effect on steps per day [9–11]. Furthermore, patients with COPD who exhibit lower baseline exercise capacity prior to PR are less likely to achieve significant improvements in physical activity [12]. Indeed, patients demonstrating a 6-min walk distance (6MWD) >350 m exhibit clinically meaningful improvements in steps per day compared to patients with a 6WMD \leq 350 m [12]. The higher likelihood of physical activity improvement in this subgroup of patients is probably because they are more conditioned to cope with the physiological demands of the intervention and have greater self-efficacy, motivation and willingness to engage and improve their physical activity levels with the support of these interventions [12].

In addition, those reporting elevated levels of anxiety and/or depression are typically less able to manage symptoms and less likely to improve physical activity or 6MWD following PR [13, 14]. This may in part be due to anxiety and fear of exacerbating symptoms, leading many to avoid their day-to-day activities as a mechanism to manage their symptoms [15]. Meanwhile, with depression, people are typically led into a sedentary lifestyle through low mood and poor motivation, which in turn causes further physical inactivity and deconditioning [16].

Cognitive behavioural therapy (CBT) is an evidence-based intervention that can be implemented to help manage symptoms of anxiety and depression in patients with COPD [17]. Specifically, HESLOP-MARSHALL *et al.* [17] developed a brief COPD-specific CBT intervention that can be delivered by respiratory nurses and demonstrated that two to six fortnightly sessions either in clinic or at home were clinically effective in reducing symptoms of anxiety in patients with moderate to severe COPD. However, in a study implementing CBT alongside PR, similar improvements were shown in depression, anxiety and quality-of-life symptoms, when compared to PR alone [18]. It is therefore envisaged that adding physical activity behavioural modification strategies to CBT during PR may optimise improvements in physical activity outcomes through improved mood and better physical conditioning in this population [13, 14].

In our original study [8], we found that a combination of physical activity behavioural modification strategies and exercise training during PR was more effective than PR alone in improving physical activity levels in patients with COPD, characterised by variable levels of anxiety and/or depression. However, the original study did not specifically look at the effects of CBT added to physical activity behavioural modification strategies (BPA) during PR on anxious and/or depressed patients with COPD.

Accordingly, in this retrospective analysis of our original study [8], we evaluated the effect of combining CBT with BPA during PR on physical activity outcomes (steps per day, movement intensity) specifically in patients with COPD who exhibit elevated anxiety and/or depression levels at the outset of a PR programme. Considering the efficacy of PR+BPA in patients with COPD [8], and the known benefits of CBT to alleviate psychological difficulties in patients with COPD [17], we envisage that a combination of CBT+PR+BPA would be superior to the provision of CBT+PR alone in improving physical activity outcomes in patients with COPD.

Methods

Study design

This is a retrospective analysis of data from a randomised controlled trial [8] which was prospectively registered at www.clinicaltrials.gov (NCT03749655) and complied with National Institute for Health and Care Research Health Research Authority requirements (reference 18/YH/0376). In this retrospective analysis we focused solely on data from patients with COPD who exhibited high baseline levels of anxiety and/or depression (Hospital Anxiety and Depression Scale (HADS) \geq 8). The cut off (\geq 8) for the diagnosis of anxiety and/or depression using the HADS questionnaire was defined by NowAK *et al.* [19] and was reported to have high sensitivity (80%) and specificity (90%) in patients with COPD [20]. This was

achieved through the randomisation process in the original randomised controlled trial [8], with 1:1 allocation to either PR+BPA or PR, stratified for HADS (<8 or \geq 8) and 6MWD (<300 m or \geq 300 m; figure 1). Once randomised, those stratified for HADS \geq 8 received CBT as part of standard care provided by the Newcastle upon Tyne Hospitals NHS Foundation Trust (NuTH) on top of their randomised allocation to PR+BPA or PR (*i.e.* PR+BPA+CBT or PR+CBT).

Participants

Patients with COPD were recruited from NuTH chest clinic and PR services. Respiratory nurses and physiotherapists informed those eligible about the study and asked their willingness to participate in the study. Inclusion criteria included 1) COPD confirmed by obstructive spirometry (post-bronchodilator forced expiratory volume in 1 s (FEV₁) to forced vital capacity (FVC) ratio <0.70); 2) clinically stable male or female aged ≥ 40 years; 3) optimised medical therapy; 4) able to provide informed consent. Exclusion criteria included 1) orthopaedic, neurological or other concomitant disease that significantly impaired normal biomechanical movement patterns, as judged by the investigator; 2) moderate or severe COPD exacerbation within 4 weeks prior to study enrolment; 3) unstable ischaemic heart disease, including myocardial infarction within 6 weeks prior to study enrolment; 4) moderate or severe aortic

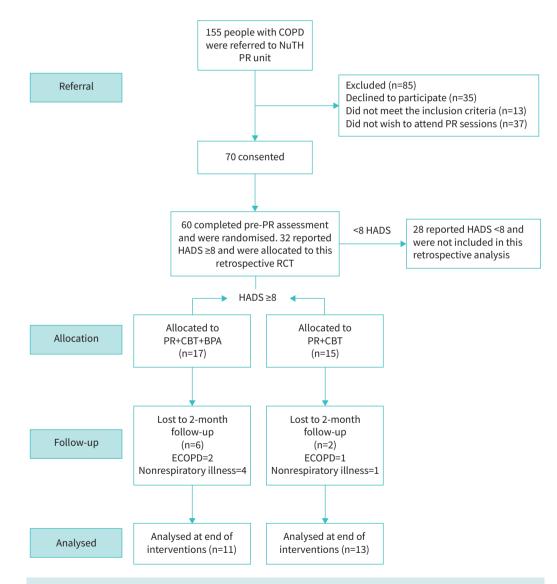


FIGURE 1 Consolidation Standards of Reporting Trials diagram of the study. NuTH: Newcastle upon Tyne Hospitals NHS Foundation Trust; PR: pulmonary rehabilitation; HADS: Hospital Anxiety and Depression Scale; RCT: randomised controlled trial; CBT: cognitive behavioural therapy; BPA: physical activity behavioural modification strategies; ECOPD: exacerbation of COPD.

stenosis or hypertrophic obstructive cardiomyopathy; 5) uncontrolled hypertension; or 6) another condition likely to limit life expectancy to <1 year (principally metastatic malignancy). Upon meeting the study entry criteria, those who agreed to participate were contacted by the research team. Detailed information regarding the study was provided and written informed consent was obtained prior to the study.

Pulmonary rehabilitation

The 8-week PR programme was delivered according to the British Thoracic Society guidelines on PR [21], with specific details regarding the PR programme available elsewhere [8]. Briefly, the 8-week PR programme comprised two 60-min sessions of exercise training per week, consisting of progressive, individualised aerobic exercise on a cycle ergometer/treadmill and resistance exercises using specific gym equipment/free weights. In addition, a 30-min session of education was delivered once per week by a multidisciplinary team comprising physiotherapists, dieticians, respiratory nurses and occupational therapists.

BPA

Specific details regarding the BPA are available elsewhere [8]. Patients received a one-to-one semi-structured motivational interview with the researcher at baseline lasting 15 min. During this time, motivational issues, favourite activities, facilitators and barriers to physical activity and strategies to become more physically active were discussed [22]. Following this, patients created three action plans, which were used throughout the programme to stimulate self-motivation to improve levels of physical activity.

BPA involved the provision of a pedometer (Fitbug, London, UK), an individualised daily step-count target (reviewed twice weekly for 8 weeks), and a step-count diary that was shared with a member of the research team at the end of every PR session. Each week, the researcher used the step count diary to calculate a new daily step-count target, by increasing the preceding week's average daily step-count by 10%. In addition, the researcher dedicated 10 min following the PR session to provide education on the importance of physical activity and advice on how to increase physical activity levels, referring to people's three action plans formed at baseline.

CBT

CBT lasting ~30 min was administered to patients with COPD with a HADS \geq 8, as part of standard PR service by respiratory nurses at either the chest clinic or at home. Specific details regarding delivery of CBT in this study can be found in a previously published randomised controlled trial [17]. Briefly, CBT focused on four key elements: behaviour, cognition/thoughts, feelings/emotions and physical sensations, with the aim of understanding how these experiences were interpreted [17]. Furthermore, to aid symptoms of anxiety and depression, several techniques including education on anxiety, depression and COPD, distraction techniques, breathing control, relaxation and rating achievement/pleasure of physical activity were used [17].

Outcome measures

All outcome measures were assessed 1 week prior to the onset of the PR programme and 1 week following completion of the PR programme. Objective measures of physical activity, including steps per day, intensity of physical activity (expressed as vector magnitude units (VMU)) and time spent in various domains of physical activity (sedentary, light and moderate vigorous physical activity) were assessed using the Actigraph wGT3X (Actigraph, Pensacola, FL, USA), validated for use in COPD [23, 24]. VMU, the sum of movements in three planes over each minute, were used to quantify the intensity of daily physical activity using the Actigraph triaxial accelerometer (Actigraph wGT3X) [25, 26]. VMU as an outcome measure is reported as the mean intensity of physical activity per minute over a specific period of accelerometer wear time, which is outlined later. Patients were instructed to wear an Actigraph accelerometer during waking hours for seven consecutive days, prior to the onset of PR and following completion of PR. Valid physical activity data from the Actigraph accelerometer were based on a minimum wear time of 8 h on \geq 4 weekdays within the 7-day period [27, 28]. Patients' experiences of physical activity were assessed using the Clinical Visit PROactive Physical Activity in COPD (C-PPAC) instrument, validated for use in COPD [29]. The instrument combines 12 questions regarding patients' subjective thoughts on amount and difficulty of physical activity alongside objective physical activity outcome data from the Actigraph accelerometer [27]. Based on these data, three categories were generated (amount of physical activity, difficulty of physical activity and total physical activity experience) ranging from 0 to 100, where higher numbers indicated a better score of physical activity experience. Other outcome measures included 1) the 6MWD [30]; 2) quadriceps muscle strength (one leg extension repetition maximum using a calibrated Myometer (MIE Medical Research, Leeds, UK), and endurance (30-s sit-to-stand repetitions) and 3) upper body strength (handgrip dynamometer) [31–33]; 4) health-related quality of life (QoL) (COPD assessment test (CAT)) [34]; 5) the Clinical COPD Questionnaire (CCQ) [35]; 6) anxiety and depression (HADS) [36]; and 7) Medical Research Council (MRC) dyspnoea scale [37].

Data analysis

Characteristics of patients with COPD and outcome data at baseline and following PR are reported as means \pm sD or 95% confidence intervals unless otherwise stated. All analyses were completed using SPSS version 28 (IBM, London, UK), with an intention-to-treat statistical analysis used for all outcome measures and missing data handled using a multiple imputation approach. Within- and between-group differences pre- to post-intervention are reported as mean (95% CI). Independent-samples t-tests were implemented to compare baseline group characteristics. A two-way repeated-measures ANOVA was implemented for all outcome variables, to identify differences between the two interventions. Statistical significance was set at p<0.05 for all analyses.

Results

Participants

In total, 32 patients with HADS \geq 8 (for anxiety and/or depression) provided consent for the study and were randomised to PR+CBT+BPA (n=17) and PR+CBT (n=15) (figure 1). Throughout the study, eight patients were lost due to nonrespiratory illness (n=5) and COPD exacerbations (n=3). Therefore, 24 patients completed the post-PR assessment visit, with 11 completing the PR+CBT+BPA and 13 completing the PR+CBT interventions (table 1). No significant differences in baseline lung function and physical activity characteristics were reported prior to both interventions (table 1).

Physical activity outcomes

Individual responses to Actigraph accelerometer-derived physical activity outcomes are shown in figures 2 and 3. Post-PR, the magnitude of improvement in steps per day was significantly greater (p=0.029) for the PR+CBT+BPA compared to PR+CBT interventions (by 829 steps per day, 95% CI 91–1564 steps per day) (table 2, figure 2). This difference was within the clinically important improvement margins (600–1100 steps per day) [38]. Following the completion of PR, the PR+CBT+BPA exhibited a significantly greater (p=0.042) improvement in the intensity of physical activity (by 80 VMU, 95% CI 3–162 VMU) (table 2, figure 3) compared to the PR+CBT intervention.

Experiences of physical activity

Individual responses to the C-PPAC instrument are shown in figure 2. Post-PR, the total score of the C-PPAC instrument was improved by a clinically important margin (>4 points) [27] in the PR+CBT+BPA intervention compared with the PR+CBT intervention, with a between-group difference of 10 points (95% CI 5–19 points, p=0.002 (table 2). For the difficulty score of the C-PPAC instrument, a clinically important

TABLE 1 Baseline characteristics							
	PR+CBT+BPA	PR+CBT	p-value				
Participants	17	15					
Age	66±4	71±5	0.306				
Male/female	4/13	7/8	NA				
BMI kg⋅m ⁻²	27±7	25±5	0.493				
FEV ₁ L	1.07±0.40	0.94±0.50	0.155				
FVC L	2.19±0.60	2.01±0.60	0.676				
FEV ₁ % pred	44±16	41±12	0.098				
FVC % pred	71±20	73±40	0.445				
FEV1/FVC %	49±13	46±14	0.228				
6MWD m	276±22	259±23	0.161				
Physical activity steps per day	2972±437	2962±466	0.217				
MVPA min	6±2	6±2	0.537				
CAT score	29±1	25±1	0.666				
HADS anxiety score	13±1	11±1	0.075				
HADS depression score	11±2	10±2	0.579				

Data are presented as n or mean±sEM, unless otherwise stated. PR: pulmonary rehabilitation; CBT: cognitive behavioural therapy; BPA: physical activity behavioural modification strategies; BMI: body mass index; FEV₁: forced expiratory volume in 1 s; FVC: forced vital capacity; 6MWD: 6-min walk distance; MVPA: moderate vigorous physical activity; CAT: COPD Assessment Test; HADS: Hospital Anxiety and Depression Scale; NA: not applicable.

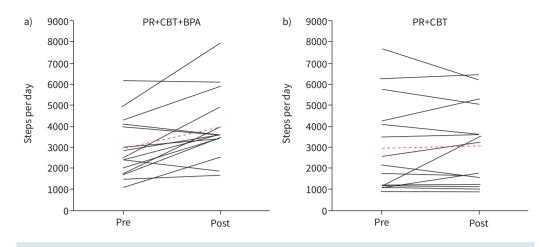


FIGURE 2 Individual responses to a) pulmonary rehabilitation (PR)+cognitive behavioural therapy (CBT)+physical activity behavioural modification strategies (BPA) and b) CBT+PR for changes in steps per day.

margin (>6 points) [27] of 13 points (95% CI 5–22 points, p=0.003 (table 2) was found. Finally, the amount score of the C-PPAC instrument improved by clinically important margin (>6 points) [27] of 13 points (95% CI 4–21 points, p=0.005 (table 2).

Anxiety and depression

Post-PR, a clinically important difference (<1.5 units) [39], in the magnitude of reduction in HADS anxiety score (by -2 unit, 95% CI -4-1 units; p=0.137) (table 3) was demonstrated in favour of the PR +CBT+BPA compared to the PR+CBT intervention. Following completion of PR, a clinically important difference in the magnitude of reduction (<1.5 units) in HADS depression score was demonstrated across both interventions, with a between-group difference of -1 unit (95% CI -3-1 units; p=0.125) (table 3).

Other outcomes

The 6MWD significantly improved following both PR+CBT+BPA and CBT+PR interventions, with no significant between group differences (table 3). However, the between-group difference (33±20 m) in 6MWD exceeded the clinically meaningful margins (30 m) (table 3) [40]. Both PR+CBT+BPA and CBT +PR demonstrated a clinically meaningful (>2 point) [39] worsening in CAT scores; however, the worsening was smaller in the PR+CBT+BPA group (table 3). Significant differences in upper body strength were demonstrated in favour of the PR+CBT+BPA group (table 3). Significant improvements in

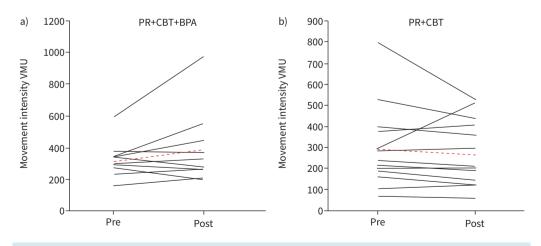


FIGURE 3 Individual responses to the a) pulmonary rehabilitation (PR)+cognitive behavioural therapy (CBT)+physical activity behavioural modification strategies (BPA) and b) CBT+PR for changes in movement intensity. VMU: vector magnitude units.

TABLE 2 Change in physical activity outcome measures in the pulmonary rehabilitation (PR)+cognitive behavioural therapy (CBT)+physical activity behavioural modification strategies (BPA) (n=17) and PR+CBT (n=15) interventions

	Baseline	Post-PR	Within-group mean difference (95% CI)	p-value	Between-group difference (95% CI)	p-value
Steps per day						
PR+CBT+BPA	2972±437	3920±422	948±247	0.001	829±360	0.029
PR+CBT	2962±466	3018±450	119±262	0.654		
Intensity of physical activity VMU						
PR+CBT+BPA	311±37	362±38	51±27	0.063	80±39	0.042
PR+CBT	298±39	266±41	-31±28	0.278		
Sedentary time min per day						
PR+CBT+BPA	513±18	461±21	-52±16	0.004	-18±24	0.458
PR+CBT	545±19	511±22	-34±18	0.065		
Light physical activity time min per day						
PR+CBT+BPA	158±12	148±10	-10±10	0.306	-3±14	0.805
PR+CBT	142±13	135±11	-7±10	0.529		
MVPA min per day						
PR+CBT+BPA	6±2	9±2	3±1	0.013	2±2	0.307
PR+CBT	6±2	7±2	1±2	0.305		
C-PPAC total score						
PR+CBT+BPA	47±3	61±9	14±2	0.001	10±3	0.002
PR+CBT	58±7	62±3	4±3	0.192		
C-PPAC difficulty score						
PR+CBT+BPA	43±4	63±4	20±3	0.001	13±4	0.003
PR+CBT	52±4	59±4	7±3	0.033		
C-PPAC amount score						
PR+CBT+BPA	44±4	63±4	19±3	0.001	13±4	0.005
PR+CBT	53±4	59±4	6±3	0.033		

Data are presented as mean±sD, unless otherwise stated. VMU: vector magnitude units; MVPA: moderate vigorous physical activity; C-PPAC: Clinical visit PROactive Physical Activity in COPD.

lower body strength were demonstrated in both the PR+CBT+BPA and PR+CBT interventions; however, no between-group differences in lower body strength were demonstrated (table 3).

Discussion

The novel findings of this study suggest that providing anxious and/or depressed patients with COPD with a combined intervention of CBT, physical activity behavioural modification strategies and PR, presents more favourable improvements in several physical activity outcome measures including steps per day, movement intensity and experiences of physical activity, compared to providing CBT alone during PR.

The magnitude of improvement in daily physical activity (by 828 steps per day) in the present study compares favourably with the majority of studies showing clinically important improvements across interventions of PR+physical activity [9, 10, 41, 42], telecoaching [43] and physical activity urban training [44] in patients with COPD. Such similarities in physical activity improvements are fundamental to patients with COPD with elevated anxiety/depression for several reasons. Firstly, previous literature has highlighted the negative impact that anxiety and/or depression may have on baseline levels of physical activity and the associated link between low levels of physical activity and increased risk of hospitalisation and mortality [4, 16]. This is particularly relevant in the current study, as the average baseline physical activity levels of our study population (2960 steps per day) were substantially lower than the average levels reported in previous studies implementing PR+physical activity in patients with COPD (~4200 steps per day) [9, 10, 41, 42, 45]. Linking closely to this, it has recently been evidenced that reduced baseline physical activity levels will probably constrain the effectiveness of physical activity interventions of this nature, highlighting the importance of our findings [5].

Accordingly, it's important to recognise the justification for these findings and gauge a better understanding of why the combined PR+CBT+BPA intervention may more effectively support physically inactive patients with COPD, compared to the CBT+PR intervention. Considering CBT, which is based upon a model of "talking therapy", patients can focus their negative thoughts and emotions on specific events that cause behavioural avoidance and explore the reasoning behind them [46]. Through the phrase

TABLE 3 Change in exercise capacity, muscular strength/endurance, quality of life and anxiety and depression outcome measures in the pulmonary rehabilitation (PR)+cognitive behavioural therapy (CBT)+physical activity behavioural modification strategies (BPA) (n=17) and PR+CBT (n=15) interventions

	Baseline	Post-PR	Within-group mean difference (95% CI)	p-value	Between-group mean difference (95% CI)	p-value
6MWD m						
PR+CBT+BPA	276±22	339±18	63±15	< 0.001	33±20	0.118
PR+CBT	259±23	289±19	30±15	0.044		
Handgrip kg						
PR+CBT+BPA	18.6±1.5	19.8±1.6	1.2±2.3	0.595	3.5±1.1	0.003
PR+CBT	23.4±1.7	21.1±1.8	-2.3±2.5	0.383		
QMVC kg						
PR+CBT+BPA	22.5±1.6	27.4±1.7	4.9±1.0	< 0.001	1.5±1.2	0.342
PR+CBT	21.2±1.7	24.6±1.8	3.4±1.1	0.004		
Sit-to-stand reps						
PR+CBT+BPA	9±1	12±1	3±1	< 0.001	1±1	0.023
PR+CBT	9±1	11±1	2±1	0.003		
CCQ (total)						
PR+CBT+BPA	3.1±0.3	2.6±0.3	-0.5±0.2	0.034	-0.8±0.3	0.020
PR+CBT	3.2±0.3	3.5±0.3	0.3±0.2	0.206		
CCQ (symptoms)						
PR+CBT+BPA	2.9±0.2	2.4±0.3	-0.5±0.2	0.023	-0.5±0.4	0.655
PR+CBT	3.4±0.2	3.4±0.3	0.0±0.3	0.973		
CCQ (functional)						
PR+CBT+BPA	2.9±0.3	2.6±0.3	-0.3±0.3	0.257	-0.2±0.4	0.564
PR+CBT	3.2±0.3	3.1±0.3	-0.1±0.3	0.778		
CCQ (mental)						
PR+CBT+BPA	2.7±0.4	1.6±0.4	-1.1±0.3	0.002	-1.1±0.5	0.034
PR+CBT	2.4±0.4	2.4±0.4	0.0±0.4	0.938		
CAT						
PR+CBT+BPA	29+1	31+1	2+1	0.519	-2+2	0.092
PR+CBT	25±1	29±1	4±2	0.020		
HADS anxiety						
PR+CBT+BPA	13±1	10±1	-3±1	0.002	-2±1	0.137
PR+CBT	10±1 11±1	10±1	-1±1	0.277		0.101
HADS depression		1011				
PR+CBT+BPA	11±1	8±1	-3±1	< 0.001	-1±1	0.125
PR+CBT	10±2	8±1	-2±1	0.001	<u><u><u></u></u><u></u><u></u><u></u><u></u></u>	0.120
MRC		511		0.000		
PR+CBT+BPA	3±1	3±1	0±1	0.302	0±1	0.205
PR+CBT	3±1	3±1 3±1	0±1	0.208	V±1	0.200

Data are presented as mean±sp, unless otherwise stated. 6MWD: 6-min walk distance; QMVC: quadriceps muscle voluntary capacity; CCQ: Clinical COPD Questionnaire; CAT: COPD Assessment Test; HADS: Hospital Anxiety and Depression Scale; MRC: Medical Research Council dyspnoea scale.

"it's not the event, but what individuals make of the event that is important", patients are able to identify unhelpful thoughts and behaviours, relieving anxious and depressive thoughts during stressful events such as exercise [46]. Subsequently, patients with COPD are less fearful during daily physical activities, preventing the vicious cycle of panic and avoidance during unexpected stages of breathlessness [15].

However, as reported in the CBT+PR group, simply alleviating anxious and/or depressive thoughts and emotions through CBT and improving exercise capacity through PR, does not automatically translate into improved physical activity levels. One reason for this links to the vicious cycle of inactivity model [47]. Within this, two vicious cycles, characterised by dyspnoea and/or deconditioning, lead to worsening of anxiety and/or breathing sensations during physical activity. As a result, patients begin to change their physical activity behaviour, choosing a sedentary lifestyle over greater levels of physical activity in response to the symptoms associated with their condition [48]. Therefore, the addition of BPA, including education on the benefits of physical activity and incorporation of behaviour change techniques such as goal setting, action planning and self-monitoring, alongside CBT and PR, may empower and motivate anxious/depressed patients with COPD to engage in more daily physical activities [8, 49].

Based on the novel findings of this study, it is plausible to suggest that through the mechanisms of behavioural modification strategies, PR, and CBT, as detailed earlier, deconditioned patients with COPD may improve physical activity by clinically meaningful margins. Furthermore, if these improvements can be replicated across future studies and clinical practice, patients may see additional improvements in key clinical outcomes including exacerbation risk, and hospital admissions, due to their close association with sustained improvements in physical activity [4, 50–53].

Another relevant finding from this study was the clinically important difference in anxiety following completion of the CBT+PR+BPA intervention, emphasising the potential beneficial effects of combining CBT with physical activity behavioural strategies on anxiety. One proposed reason for this may link to the previously detailed association between anxiousness and physical activity in patients with COPD, particularly with reference to fear avoidance [54]. Specifically, a recent study provided preliminary evidence that disease-specific anxiety and fear avoidance have a specific influence on predicted physical activity levels in patients with COPD who displayed lower relative physical capacity [54]. As a result, the smaller improvement in anxiety in those undertaking CBT+PR may have been associated with the poor baseline physical capacity and physical activity levels, leaving patients in continuous fear and avoidance of physical activity.

A clinically important difference in 6MWD following completion of both interventions was reported. confirming the effectiveness of PR for improving exercise capacity [55]. Interestingly, those who completed the CBT+PR+BPA intervention experienced superior improvements in 6MWD, which exceeded the clinically meaningful margin (30 m). Importantly, a significant improvement for the intensity of physical activity was found for the CBT+BPA+PR intervention, with a significant between-intervention magnitude of change in intensity of physical activity. It's plausible to suggest that the reduction in anxiety and fear as well as greater confidence in managing symptoms during daily activities would encourage patients to complete daily activities at a greater pace/intensity [15]. In turn, these greater intensities may have translated into improved experiences of physical activity, as reported through the significant improvement in C-PPAC instrument outcomes. The clinical relevance of this finding is important due to the severe reduction in walking pace typically reported in patients with severe COPD, that is known to be associated with poor prognosis [56]. Often, the intensity of physical activity presents an indication of a patient's ability to conduct physical activity, providing a breakdown of the time taken to complete a task. Unlike steps per day, which gives an overall accumulation of activity over a set period, the intensity of physical activity provides better understanding of how patients' symptoms impact physical activity and how they cope/engage with daily tasks [56]. Through observing improvements in both the duration (steps per day) and intensity (VMU) of physical activity, its plausible to suggest that patients in the current study were able to increase both the amount of walking activity and the pace at which walking activity was undertaken.

Based on the novel findings of this study, it's important to highlight that despite the equally poor physical activity levels (\leq 4000 steps per day) and 6MWD (\leq 350 m) across all patients with COPD in both the PR +CBT+BPA and PR+CBT groups, the combination of PR+CBT+BPA resulted in greater improvements for several outcomes including physical activity and 6MWD compared to PR+CBT. These findings provide vital evidence that combining the three elements of this intervention can challenge earlier claims that physical activity counselling interventions alongside PR are effective in improving physical activity levels only in patients with COPD exhibiting greater levels of physical activity (>4000 steps per day) and exercise capacity (>350 m) at baseline [9–12].

Study limitations

Several limitations should be considered from this study. Firstly, this retrospective analysis was unable to assess the true effect of CBT. This was due to CBT being part of standard care across NuTH, meaning we couldn't modify standard care for the purpose of this research. Secondly, eight out of 32 patients were lost during the study, meaning that only 24 patients with COPD completed both pre and post assessments. As a result, generalisability of the findings to clinical practice may be limited at this stage. Nevertheless, the 25% dropout rate across the study duration is in line with the average dropout rate reported by the UK National COPD Audit Programme for pulmonary rehabilitation [57]. Thirdly, CBT engages several techniques including education on anxiety and depression, thereby aiding the mitigation of the intensity of these symptoms. Nevertheless, despite our finding of a clinically important difference in anxiety following completion of the CBT+PR+BPA intervention, we cannot reliably confirm that the additional contact with the CBT specialist did not assist patients to improve their mental health, contributing to the observed benefit. Furthermore, all components of the intervention (CBT, PR and BPA) included factors that are difficult to keep separate while being delivered collectively (*e.g.* the PR+CBT group may receive elements of behaviour change which may contribute to BPA). Indeed, exercise training as part of PR may aid

improvements in daily physical activity, albeit to a smaller extent [5]. Finally, due to the intervention and all outcome measures being administered by a single researcher in a face-to-face manner, blinding of assessors was not possible. Our inability to blind assessors and those receiving the intervention may have reduced the overall quality of evidence and increased the risk of bias towards the intervention, however due to the study protocol this was unavoidable.

Conclusion

Incorporating CBT and physical activity behavioural modification interventions into a standard PR programme may provide patients with COPD who exhibit low baseline levels of physical activity and elevated anxiety and/or depression with clinically important improvements in physical activity levels and experiences of physical activity, alongside improvements in exercise capacity, health-related quality of life and anxiety. Further development and delivery of this combined intervention may support the management/treatment of patients with COPD who exhibit high levels of anxiety and/or depression, consequently lowering the risk of hospital admissions, morbidity and mortality, associated with physical inactivity in COPD.

Provenance: Submitted article, peer reviewed.

This study is registered at www.clinicaltrials.gov with identifier number NCT03749655. Individual participant data that underlie the results, reported in this article, will be shared after de-identification. Documents available will include the study protocol, statistical analysis plan and analytic code and will be available following publication to anyone who wishes to access the data.

Conflict of interest: M. Armstrong has no conflicts of interest to disclose for this study. E. Hume has no conflicts of interest to disclose for this study. L. McNeillie has no conflicts of interest to disclose for this study. F. Chambers has no conflicts of interest to disclose for this study. L. Wakenshaw has no conflicts of interest to disclose for this study. G. Burns has received a grant from NIHR for a CBT study. K. Heslop Marshall received a grant from NIHR for a CBT study and director of pivotal healthcare training for CBT skills. I. Vogiatzis is an associate editor of this journal.

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