

# Effectiveness of perioperative pulmonary rehabilitation in thoracic surgery

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**Background:** Functional condition is crucial for operability of patients with lung cancer and/or chronic respiratory diseases. The aim of the study was to measure changes of functional and quality of life parameters in terms of the effectiveness of perioperative pulmonary rehabilitation (PR).

**Methods:** A total of 208 COPD patients (age: 63±9 years, man/woman: 114/94, FEV1: 62±14 %pred) participated in a perioperative PR program. The indication was primary lung cancer in 72% of the patients. The 68 patients participated in preoperative (PRE) rehabilitation, 72 in a pre- and postoperative rehabilitation (PPO) and 68 patients only in postoperative rehabilitation (POS). PR program included respiratory training techniques, individualized training and smoking cessation. Lung function tests, 6 minutes walking distance (6MWD) were measured before and after the rehabilitation. Quality of life tests [COPD Assessment Test (CAT) and Modified Medical Research Council Dyspnoea Scale (mMRC)] were evaluated as well.

**Results:** There was a significant improvement in FEV1 (PRE: 64±16 vs. 67±16 %pred; PPO: 60±13 vs. 66±13 %pred before the operation, 48±13 vs. 52±13 %pred after the operation; POS: 56±16 vs. 61±14 %pred, P<0.05) and 6MWD (PRE: 403±87 vs. 452±86 m; PPO: 388±86 vs. 439±83 m before, 337±111 vs. 397±105 m after the operation; POS: 362±89 vs. 434±94 m, P<0.0001). Significant improvement was detected in FVC, grip strength, mMRC and CAT questionnaires as an effectiveness of PR, also. Average intensive care duration was 3.8±5.2 days with vs. 3.1±3.6 without preoperative PR.

**Conclusions:** Improvements in exercise capacity and quality of life were seen following PR both before and after thoracic surgery.

**Keywords:** Perioperative pulmonary rehabilitation (perioperative PR); thoracic surgery; lung cancer; chest physiotherapy; endurance training; chronic obstructive pulmonary disease (COPD)

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## 1 Introduction

2 Severe chronic obstructive pulmonary disease (COPD),  
 3 severely impaired functional capacity, low physical  
 4 activity, obesity, smoking and comorbidities are significant  
 5 factors for risk stratification before thoracic operation  
 6 (1-3). Preoperative rehabilitation can improve functional  
 7 condition. Improvement of cardiovascular function,  
 8 metabolism, muscle-function and lung mechanics can be  
 9 achieved by pulmonary rehabilitation (4).

10 More interest is focusing the non-pharmacological  
 11 interventions, such as exercise and improving functional  
 12 capacity during, before and after thoracic surgery and  
 13 cancer treatment. Exercise is proved to be successful  
 14 intervention that makes physical and psychological health  
 15 better in different cancer states, including lung cancer (5,6).  
 16 Cardiopulmonary rehabilitation is part of the management  
 17 of COPD. It can reduce symptoms and minimize the  
 18 exacerbation rate of the disease (5,6). Smoking cessation  
 19 is important in terms of reduction of postoperative  
 20 complications as well (7).

21 It has been shown, that pulmonary rehabilitation (PR)  
 22 can improve exercise capacity and Health Related Quality  
 23 of Life (HRQOL), and can reduce the main symptoms such  
 24 as dyspnoea, fatigue and depression (8). There is limited  
 25 availability of specific exercise intervention for patients with  
 26 lung cancer (8). It is important to define the optimal design  
 27 of exercise intervention that will be feasible, acceptable and  
 28 positively affect under perioperative conditions.

29 Our aim was to investigate the effectiveness of  
 30 perioperative rehabilitation by monitoring the parameters  
 31 of lung function, lung mechanics, chest kinematics, exercise  
 32 capacity and quality of life.

## 34 Methods

### 36 Study subjects

37 A total of 208 COPD patients participated in the  
 38 perioperative pulmonary rehabilitation program in  
 39 connection with thoracic operations between 2012 and  
 40 2015 at Department of Thoracic Surgery in National  
 41 Koranyi Institute for Pulmonology, Hungary. Indication of  
 42 the operation was primary lung cancer in 150, pulmonary  
 43 metastasis in 11, benign disease in 10, infection in 16, other  
 44 cause in 21 cases. The patient's characteristic is presented  
 45 in *Table 1*. All of the patients gave consent for the study  
 46 in the Department of Pulmonary Rehabilitation. It was  
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an observational study using the general management of  
 the patient, it was a non-interventional study. There was  
 no significant difference between the groups in terms of  
 patient's characteristics (*Table 1*).

Our patients participated in three groups. The  
 68 patients performed only preoperative pulmonary  
 rehabilitation (PRE). In PPO group, 72 patients performed  
 pre- and postoperative rehabilitation also, and 68 patients  
 had postoperative rehabilitation only (POS) (*Figure 1*).

### *Pulmonary function*

According to ATS/ERS guidelines all patients underwent  
 post-bronchodilator pulmonary function testing ( $V_{\max}$  229  
 and Autobox 6200, Sensormedics) including spirometry  
 measurements (9). COPD patients inhaled 400  $\mu$ g  
 salbutamol 20 minutes before testing.

### *Functional follow-up and quality of life questionnaire*

Functional follow-up included measurement of lung  
 functions, 6 minutes walking distance (6MWD) test (10)  
 and quality of life tests such as COPD Assessment Test  
 (CAT) (11) and Modified Medical Research Council  
 Dyspnoea Scale (mMRC) (12).

### *Personalized training programs*

Our pulmonary rehabilitation program includes  
 30 minutes of respiratory training in the morning, chest wall  
 mobilization, learning the controlled breathing techniques,  
 inhalation, expectoration, improving the psychological  
 condition, smoking cessation and a personalized training.  
 Patients participate an individualized continuous or interval  
 type of cycle- and/or treadmill training for 10–30 minutes,  
 2–3 times a day at a level of 60–80% of maximal intensity.  
 The duration of the rehabilitation program is 3 weeks (13).  
 The intensity of the training was progressive from 60–80%  
 of peak work rate, the intensity was increased based on  
 maintaining Borg dyspnea scale breathlessness and leg  
 fatigue both on grade No. 7.

### *Smoking cessation*

Smoking cessation is an important part of the perioperative  
 rehabilitation program. Our institute has a special smoking  
 cessation program for the patients once per week for 45

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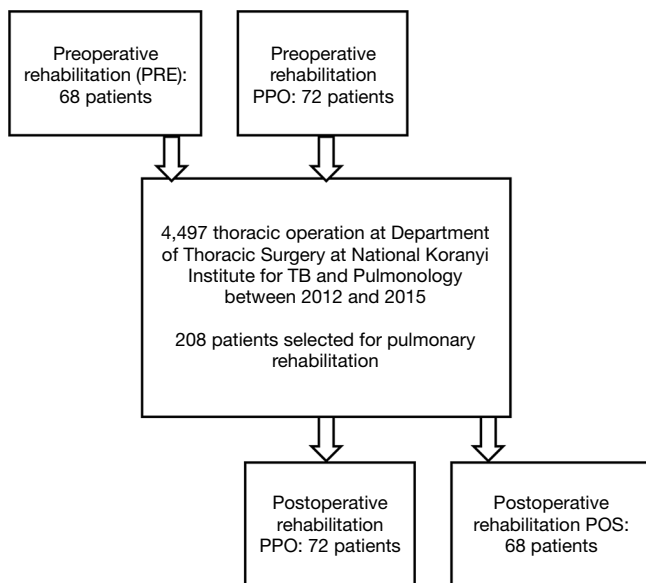
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**Table 1** Patient's characteristics

Characteristics, N=208	Group PRE, n=68	Group POS, n=68	Group PPO, n=72	Significance
Age (years)	65±6	60±11	65±7	NS
Male:female	45:23 (66%:34%)	35:33 (51%:49%)	34:38 (47%:53%)	NS
BMI (kg/m <sup>2</sup> )	27±5	25±5	26±6	NS
FEV <sub>1</sub> (%pred)	64±16	55±16	60±13	NS
Hypertension	46 (68%)	43 (68%)	46 (72%)	NS
Diabetes	25 (37%)	25 (37%)	23 (32%)	NS
Atherosclerosis	22 (32%)	23 (34%)	24 (33%)	NS
Pulmonary hypertension	10 (15%)	9 (13%)	8 (11%)	NS
Quitting rate of smoking cessation	52 (76%)	51 (75%)	54 (75%)	NS

**Figure 1** Flow of participants in the perioperative rehabilitation program.

94 minutes, with help of psychologists (7).

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### 96 **Statistical analysis**

97 Subject's characteristics, lung function and exercise  
 98 physiologic variables were compared by paired *t*-test and  
 99 non-parametric sign test and Wilcoxon test. Significance  
 100 was accepted at the  $P < 0.05$  level. The distribution around  
 101 the mean was expressed as  $\pm$  SD, in tables and the figures  
 102 also. Distributions were tested for normality by the  
 103 Kolmogorov-Smirnov test.

## Results

Clinically significant improvement was detected in FEV<sub>1</sub>, FVC, 6MWD, grip strength, in PRE, in PPO before and after the operation and in POS group (*Tables 2-4*,  $P < 0.05$ ). The level of dyspnoea (measured by the mMRC Dyspnoea Scale) and quality of life (measured by CAT) improved significantly in PRE group, in the PPO group before and after the operation, and in POS group, also (*Tables 2-4*;  $P < 0.001$ ). There was no significant difference between the result of cancer and non-cancer patients in functional parameters. Average intensive care duration was  $3.8 \pm 5.2$  days with *vs.*  $3.1 \pm 3.6$  without preoperative pulmonary rehabilitation. The quitting rate was high, and not significantly different between groups (*Table 1*) (*Figures 2,3*).

## Discussion

Perioperative pulmonary rehabilitation was performed before and/or after thoracic surgery in patients with COPD. Pre- and postoperative pulmonary rehabilitation resulted significant improvement in FEV<sub>1</sub>, FVC, 6MWD and quality of life questionnaires (CAT, mMRC). There was no significant difference between intensive care duration with or without preoperative pulmonary rehabilitation.

Exercise tolerance is a crucial part of risk stratification before surgical resection. Exercise interventions can lead to improve exercise capacity parallel with reduction of risk in cardiopulmonary function (14). Reduction in postoperative complications and length of hospital stay can be detected as a result of pre-surgical interventions, which

**Table 2** Changes in functional parameters as the effectiveness of preoperative pulmonary rehabilitation (PRE group)

Parameter	PRE (preoperative pulmonary rehabilitation), n=68		
	Before rehab.	After rehab.	Change, significance
FEV1	1.75±0.58 L (63.7±16.0 %pred)	1.90±0.58 L (67.4±16.3 %pred)	+5.89% (P=0.0025)
FVC	2.92±0.90 L (85.8±17.6 %pred)	3.12±0.89 L (90.6±16.8 %pred)	+5.66% (P=0.0109)
6MWD (m)	403±87	452±86	+12.07% (P<0.0001)
mMRC	1.0±0.7	0.7±0.6	-31.71% (P=0.0004)
Grip strength (kg)	19.9±14.4	21.8±15.5	+10.02% (P=0.0002)
CAT	8.4±5.3	5.4±4.7	-35.47% (P<0.0001)

6MWD, 6 minutes walking distance; mMRC, Modified Medical Research Council Dyspnoea Scale; CAT, COPD Assessment Test; COPD, chronic obstructive pulmonary disease.

**Table 3** Changes of functional parameters as the effectiveness of postoperative pulmonary rehabilitation (POS group)

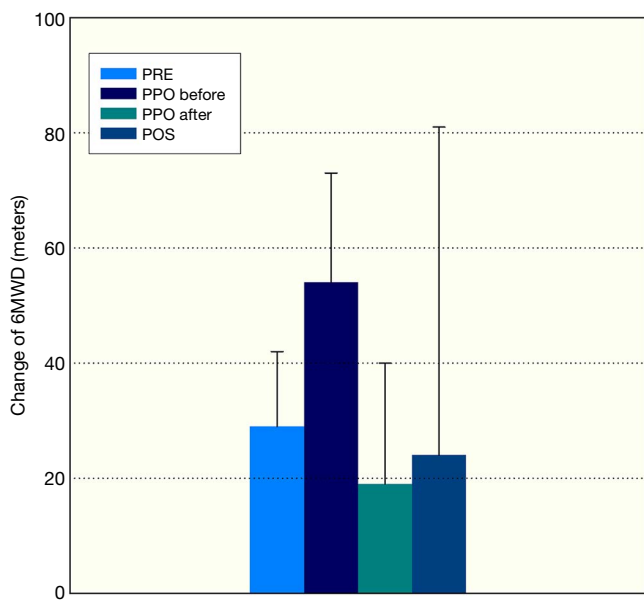
Parameter	POS (postoperative pulmonara rehabilitation), n=68		
	Before rehab.	After rehab.	Change, significance
FEV1	1.50±0.49 L (55.6±16.2 %pred)	1.75±0.61 L (60.8±14.2 %pred)	+9.34% (P=0.0044)
FVC	2.19±0.74 L (66.8±21.3 %pred)	2.54±0.83 L (74.9±19.6 %pred)	+12.09% (P=0.0001)
6MWD (m)	362±89	434±94	+19.88% (P<0.0001)
mMRC	1.5±1.0	1.0±0.8	-32.31% (P<0.0001)
Grip strength	19.2±12.3	21.2±13.2	+10.14% (P=0.0008)
CAT	17.6±9.0	12.8±8.8	-27.00% (P<0.0001)

6MWD, 6 minutes walking distance; mMRC, Modified Medical Research Council Dyspnoea Scale; CAT, COPD Assessment Test; COPD, chronic obstructive pulmonary disease.

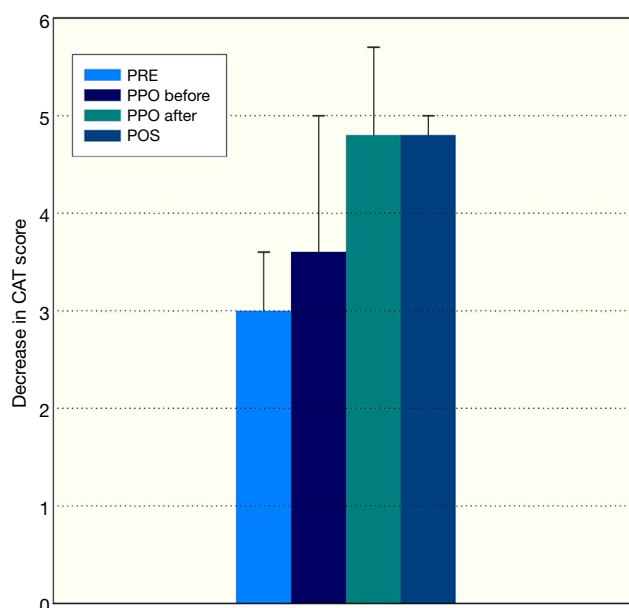
**Table 4** Changes of functional parameters as the effectiveness of pre- and postoperative pulmonary rehabilitation (PPO group)

Parameter	PPO (pre- and postoperative pulmonary rehabilitation), n=72					
	Preoperative			Postoperative		
	Before rehab.	After rehab.	Change, significance	Before rehab.	After rehab.	Change, significance
FEV1	1.49±0.53 L (60.1±12.8 %pred)	1.68±0.53 L (66.3±12.9 %pred)	+10.39% (P<0.0001)	1.21±0.43 L (48.4±12.7 %pred)	1.30±0.40 L (51.8±13,0 %pred)	+7.14% (P=0.0247)
FVC	2.57±0.81 L (88.7±14.7 %pred)	2.86±0.82 L (97.9±14.7 %pred)	+10.41% (P=0.0001)	2,00±0.72 L (63.6±16.1 %pred)	2.13±0.68 L (67.7±17.7 %pred)	+6.57% (P=0.1126)
6MWD (m)	388±86	439±83	+13.06% (P<0.0001)	337±111	397±105	+17.74% (P<0.0001)
mMRC	1.2±1.0	0.8±0.8	-35.30% (P=0.0002)	1.8±0.9	1.4±0.8	-18.47% (P=0.0017)
Grip strength (kg)	22.0±12.0	23.3±12.8	+5.69% (P=0.0057)	20.2±14.1	21.1±14.5	+4.23% (P=0.3658)
CAT	12.0±6.9	8.4±5.5	-30.08% (P<0.0001)	16.0±6.2	11.4±5.3	-28.75% (P<0.0001)

6MWD, 6 minutes walking distance; mMRC, Modified Medical Research Council Dyspnoea Scale; CAT, COPD Assessment Test; COPD, chronic obstructive pulmonary disease.



**Figure 2** Improvement of maximal exercise capacity measured by 6-minute walking test, PRE-preoperative rehabilitation group, PPO-perioperative rehabilitation group, PPO-before: pulmonary rehabilitation before thoracic surgery, PPO-after: pulmonary rehabilitation after thoracic surgery, POS-postoperative rehabilitation group, mean ± standard deviation were presented in the columns.



**Figure 3** Improvement of quality of life measured by COPD Assessment Test (CAT), PRE-preoperative rehabilitation group, PPO-perioperative rehabilitation group, PPO-before: pulmonary rehabilitation before thoracic surgery, PPO-after: pulmonary rehabilitation after thoracic surgery, POS-postoperative rehabilitation group, mean ± standard deviation were presented in the columns. COPD, chronic obstructive pulmonary disease.

135 may be in connection with increased exercise capacity (14).  
 136 We detected significant improvement in exercise capacity  
 137 as a potential predictor of reduction of postoperative  
 138 complication, but we did not collect the complication rate  
 139 in this study.

DEMO Pulmonary rehabilitation can improve lung mechanics,  
 140 exercise capacity and quality of life using controlled  
 141 breathing techniques, chest wall mobilization and specific  
 142 training modalities. Specialized training modalities are  
 143 favourable for respiratory and peripheral muscles as well  
 144 (15). The effectiveness of our rehabilitation program  
 145 was supported by improvement of lung mechanics, exercise  
 146 capacity and quality of life.

147 Clinical data underline that exercise intervention  
 148 compared with usual care both pre and post-surgery is a  
 149 safe, feasible and acceptable method, which can increase  
 150 exercise capacity, reduce postoperative complications and  
 151 hospital stay (16). Exercise can increase muscle strength  
 152 and reduce fatigue, as well (16). In the other hand, the  
 153 improvement in pulmonary function, quality of life,

and blood gas analysis is not consistent. Our exercise  
 interventions were safe, feasible and effective for the  
 patients.

As a result of rehabilitation, increased muscle strength  
 and reduced fatigue are important findings. Resistance  
 training can improve muscle strength, highlighting the  
 importance of interventions (17). Exercise intervention  
 suggests that physical exercise can help to reduce fatigue  
 both during and after treatment for cancer including lung  
 cancer (18,19). However, there is a question about the  
 component of rehabilitation against fatigue.

Different types of exercise, aerobic exercise in  
 comparison with resistance exercise alone, may have  
 superior effect for improving exercise capacity and  
 oxygen uptake (VO<sub>2</sub> max), but there are not so many data  
 about (17) resistance exercise. Resistance training can  
 result (17) significant improvement in muscle strength.  
 The combination of resistance and aerobic exercise  
 training may result the optimal training program for this  
 population. Our rehabilitation program included strength

DEMO and endurance part, also.

173 Endurance training is an important component of  
 174 pulmonary rehabilitation. Supervised high intensity interval  
 175 training can be more effective compared to continuous  
 176 training in healthy subjects. In patients with COPD, these  
 177 two methods have no significant difference in results  
 178 (13,20). Both methods have advantages according to self-  
 179 paced training (13,20). It is important to consider the  
 180 patient's functional capacities, significant co-morbidities  
 181 (like impaired pulmonary hemodynamics) and desaturation  
 182 index during exercise to find the optimal, personalized  
 183 training program (20). In our program, patients performed  
 184 continuous and interval training based on functional  
 185 capacity and comorbidities.

186 Improvements in exercise capacity, together with no  
 187 change in pulmonary function may be an unexpected  
 188 findings in this type of patients like other pulmonary  
 189 diseases, for example, COPD and restrictive disorders  
 190 (21,22). Overall there is a question about the evidence that  
 191 exercise may or may not improve pulmonary function in  
 192 this population. We found significant improvement in lung  
 193 function using these rehabilitation techniques.

194 Quality of life has a non-significant change in one part  
 195 of the studies. Preoperative rehabilitation period (23) may  
 196 result no change in QOL due to a possible ceiling effect.  
 197 Randomized controlled trials (24-26) for post-operative  
 198 interventions resulted no significant differences in QOL  
 199 between groups, however there were certain limitations  
 200 of the studies. In the other hand, in the study by Arbane  
 201 *et al.* (27), there was no data about the adherence to home-  
 202 based exercise and how much exercise was undertaken,  
 203 in another clinical study (18) the intervention group was  
 204 significantly fitter at baseline, and finally, Gao *et al.* (28)  
 205 did not use a disease specific tool to measure QOL.  
 206 Differences in tools, design of intervention and extent of  
 207 surgery made comparisons and conclusions concerning  
 208 QOL difficult. There is a need for future studies to use  
 209 equal measurement instruments for which reliability and  
 210 validity (27,28). We detected significant improvement  
 211 in quality of life and dyspnoea measured by CAT and  
 212 mMRC.

213 Our perioperative pulmonary rehabilitation program  
 214 was shorter compared to general duration of pulmonary  
 215 rehabilitation program based on ATS, ERS, BTS guidelines.  
 216 These types of patients performed the pulmonary  
 217 rehabilitation before thoracic surgery because of lung  
 218 cancer or other severe thoracic diseases. We needed to  
 219 reduce the time before thoracic surgery and there are some

studies about a shorter duration of pulmonary rehabilitation DEMO  
 effective for improvement of cardiovascular response, 220  
 metabolism, muscle function and lung mechanics. 221

222 Our study had limitations as well. This was a cohort  
 223 study and there was no control group. We hadn't collected  
 224 postoperative complications in this study. We can choose  
 225 a wide range of other quality of life scores to evaluate  
 226 symptoms, like Saint George Respiratory Questionnaire,  
 227 Chronic Clinical Questionnaire (CCQ). We may focus the  
 228 change of depression and anxiety in the future as a result of  
 229 perioperative pulmonary rehabilitation, as well.

230 Pulmonary rehabilitation had positive effects before  
 231 lung surgery by improving exercise capacity and functional  
 232 reserves. Thoracic surgical functional operability  
 233 and postoperative management was promoted by the  
 234 conditioning effect of pulmonary rehabilitation. Positive  
 235 effects of pulmonary rehabilitation appeared objectively  
 236 in all three groups. There was significant improvement  
 237 in lung function, chest kinematics, exercise capacity and  
 238 quality of life, also. We consider this rehabilitation program  
 239 as an effective management of pulmonary rehabilitation in  
 240 perioperative condition. 241

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 249 psychologist and Erika Pataki for coordinating the smoking  
 250 cessation program. 251

## Footnote 252

253 *Conflicts of Interest:* The authors have no conflicts of interest  
 254 to declare. 255

256 *Ethical Statement:* The study was approved by Institutional  
 257 Review Board and written informed consent was obtained  
 258 from all patients. 259

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