Association between Emergency Visits with Acute Exacerbation and Exercise Performance in Chronic Obstructive Pulmonary Disease

İlknur Naz¹, Hülya Şahin², Işıl Karasu³, Nimet Aksel³

¹Department of Physiotherapy and Rehabilitation, İzmir Katip Çelebi University Faculty of Health Sciences, İzmir, Turkey ²Department of Pulmonary Rehabilitation, University of Health Sciences Dr. Suat Seren Chest Diseases and Surgery Training and Research Hospital, İzmir, Turkey

³Department of Chest Diseases, University of Health Sciences Dr. Suat Seren Chest Diseases and Surgery Training and Research Hospital, İzmir, Turkey

Cite this article as: Naz I, Şahin H, Karasu I, Aksel N. Association between Emergency Visits with Acute Exacerbation and Exercise Performance in Chronic Obstructive Pulmonary Disease. Eurasian J Pulmonol 2017; 19: 71-5.

Abstract

Objective: The relationship between acute exacerbation and exercise performance in chronic obstructive pulmonary disease (COPD) is unclear. The aim of this study was to compare exercise performance between patients with COPD who visited and did not visit an emergency department in the last year; we also aimed to investigate the correlation between frequency of emergency visits and exercise performance.

Methods: We recorded the number of emergency department visits of 206 COPD patients. The six-minute walk test, the modified Medical Research Council Dyspnea Scale, body plethysmographs, carbon monoxide diffusion tests, and arterial blood gas analysis were practiced.

Results: We had one hundred twenty seven participants visited an emergency department at least once in the last year. Exercise performance, dyspnea, pulmonary functions, and oxygenation were significantly poorer in those patients (p<0.05). The number of emergency department visits was correlated with exercise performance, dyspnea, oxygenation, forced expiratory volume in one second forced expiratory volume in one second/forced vital capacity, and carbon monoxide diffusion capacity (p<0.05).

Conclusion: Exercise performance was poor and negatively related to the frequency of emergency department visits in COPD patients. We concluded that exercise performance-enhancing implementations may contribute to reduce the frequency of emergency department visits.

Keywords: Acute exacerbation, COPD, emergency visit, exercise performance, six-minute walk test



Received Date: 16.09.2016 Accepted Date: 31.12.2016 DOI: 10.5152/ejp.2017.98159

Corresponding Author İlknur Naz E-mail: ilknurnaz4@gmail.com

Available online at www.eurasianjpulmonol.com



This work is licensed under a Creative Commons Attribution Nor C Commons Attribution-NonCommercial 4.0 International License

INTRODUCTION

Chronic obstructive pulmonary disease (COPD) is one of the leading causes of mortality and morbidity worldwide; however, it is also a chronic disease with increasing prevalence despite medical treatment (1, 2). Acute exacerbation of COPD, which is the most common cause of hospital visits, may appear at any stage of the disease; its frequency and severity increases with disease severity (3, 4). Accelerated reduction in forced expiratory volume in one second (FEV,), which is the primary determinant of pulmonary function and exacerbation, increases the severity of the disease, reduces the quality of life of patients, and increases health expenditures (5-7). Serious exacerbations requiring hospitalization are the leading causes of mortality and morbidity in COPD (8).

Physical decondition and pulmonary dysfunction are the primary causes of exercise intolerance, which is the main clinical feature of COPD (9). Assessment of exercise tolerance is necessary to evaluate the impact of COPD; however, measurement of pulmonary functions such as FEV, is not adequate to determine exercise tolerance (10). The six-minute walk test is a submaximal exercise test that assesses the functional status of COPD patients. It is easily applicable, inexpensive, and repeatable. In addition, this test allows simultaneous evaluation of the patient's pulmonary, cardiovascular, musculoskeletal, and neuromuscular systems (11, 12).

There is no study in the literature investigating the correlation between exercise performance and emergency visits with acute exacerbations in patients with COPD. In the present study, we investigated COPD patients in our clinic in two groups, those who visited and did not visit an emergency department in the last year; we aimed to compare exercise performance between these groups as well as to investigate the correlation between frequency of emergency department visits and exercise performance in cases who visited an emergency department.

METHODS

A total of 206 symptomatic COPD patients (189 males, 91.7%) who were referred to the University of Health Sciences Dr. Suat Seren Chest Diseases and Surgery Training and Research Hospital, Depart-ment of Chest Diseases between June 2013 and June 2015 were enrolled in our cross-sectional study. The demographic and clinical characteristics of the patients were recorded. They were asked about their emergency department visits due to dyspnea, as well as the increases in amount and purulence of their sputum in the last year. The study was approved by the local institutional review board (number: 320). Written consent was obtained from all patients. The physical and demographic data, disease duration, and history of smoking of the patients were recorded. All patients underwent pul-monary and cardiac system examination and pulmonary function testing (PFT). Lung radiographs and arterial blood gases (ABG) were evaluated. Each patient performed the sixminute walk test.

Pulmonary functions: The pulmonary functions of the patients were evaluated by body plethysmography (Zan 500, Germany) and carbon monoxide diffusion capacity (DLCO) (Zan 300, Germany).

Perception of dyspnea: The five-item Medical Research Council (mMRC) dyspnea scale was used to determine the severity of the pa-tients' shortness of breath (13). "1" indicates the lowest severity, and "5" indicates the highest severity.

Exercise performance: Exercise performance was assessed by the six-minute walk test, which measures the longest distance patients can walk in six minutes (14). Heart rate, peripheral oxygen saturation, dyspnea, and leg fatigue, which was measured by the modified Borg scale pretest and post-test, were recorded.

The modified Borg scale, which is scored from 0 to 10, is used to as-sess exertional dyspnea; "0" indicates no dyspnea, and "10" indicates severe dyspnea (15). The study participants were divided into in two groups: patients who had visited and who had not visited an emergency department in the last year.

Statistical Analysis

We performed statistical analyzes using SPSS 15.0 (Statistical Pack-age for the Social Sciences Inc.; Chicago, IL, USA). Descriptive statisti-cal tests were performed for all the recorded variables, and the data were expressed as median (interquartile range) or percentage (%). We used the Mann-Whitney U test to compare continuous data and the chi-square test for categorical data. We determined the relationships between continuous variables by cal-

culating Spearman correlation coefficient. We considered p<0.05 to indicate significance.

RESULTS

hotwoon group

Of the 206 study participants, 127 (61.6%) (116 males, 91.3%) had visited an emergency department at least once, whereas 79 (73 males, 92.4%) had not visited an emergency department in the last year. The median number of exacerbation was 3(1, 6) in patients who visited an emergency department. Of those 127 patients, 73 (57.5%) were hospitalized and 54 (42.5%) were prescribed drug treatment after exacerbation.

The median age and the number of patients with stage 4 COPD were significantly higher in patients who visited an emergency department (p=0.032 and p<0.001, respectively; Table 1). Both groups were similar in terms of body mass index (BMI) and cigarette consumption (p=0.99 and p=0.91, respectively; Table 1). Perception of dyspnea, pulmonary functions, and oxygenation were significantly poorer in patients who visited an emergency department compared to patients who did not visit an emergency department in the last year (p<0.05, Table 1).

When exercise performance was compared between the groups, the six-minute walk distance and changes in peripheral oxygen Saturation between pre and post-test were significantly lower and changes in dyspnea and leg fatigue were higher in patients who visited an emergency department (p<0.05, Table 2).

Table 1. Comparison of demographic and clinical characteristics

between groups							
Variable	Emergency department visit (+) (n=127)	Emergency department visit (-) (n=79)	р				
Age (years)	64 (58, 70)	61 (55,67)	0.032*				
BMI (kg/m ²)	26 (22, 30)	26 (23, 30)	0.99				
Smoking (p-y)	50 (30, 80)	50 (35, 80)	0.91				
STAGE n (%)							
Stage 1	0 (%0)	4 (%5.1)	0.021*				
Stage 2	23 (%18.1)	29 (%36.7)	0.005*				
Stage 3	56 (%44.1)	35 (%44.3)	0.545				
Stage 4	48 (%37.8)	11 (%13.9)	<0.001*				
FEV ₁ (% predicted)	34 (26, 47)	44 (35, 63)	<0.001*				
FEV ₁ /FVC	54 (44, 62)	60 (52, 70)	0.001*				
DLCO (%)	34 (24, 44)	43 (30, 52)	0.008*				
PaO ₂ (mmHg)	67 (60, 76)	75 (69, 82)	<0.001*				
PaCO ₂ (mmHg)	41 (38, 47)	39 (36, 44)	0.014*				
SaO ₂ (%)	94 (91, 95)	95 (94, 96)	<0.001*				
mMRC	4 (3, 5)	3 (2, 4)	<0.001*				

Data are expressed as median (interquartile range) or percentage (%)

*p<0.05 for comparison of changes between groups based on Mann-Whitney U test or chi-square test

BMI: Body mass index; p-y: packs-year; FEV; forced expiratory volume in one second; FVC: forced vital capacity; DLCO: carbon monoxide diffusing capacity; PaO₂; partial arterial oxygen pressure; PaCO₂: partial arterial carbon dioxide pressure; SaO₂: arterial oxygen saturation; mMRC: Medical Research Council dyspnea scale Correlation analysis performed for the patients who visited an emergency department revealed a weak negative correlation between the number of emergency department visits and exercise performance (r=-0.301; p=0.001), FEV₁ (%) (r=-0.254; p=0.004), FEV₁/forced vital capacity (FVC) (r=-0.300; p=0.001), and DLCO (%) (r=-0.264; p=0.012); weak positive correlation with perception of dyspnea (r=0.367; p<0.001); and very weak negative correlation with partial arterial oxygen pressure (PaO2) and arterial saturation (r=-0.179; p=0.046 and -0.178; p=0.047, respectively; Table 3).

DISCUSSION

In this study, we found that exercise performance was lower in COPD patients who visited an emergency department at least once in the last year. Moreover, high numbers of emergency visits were correlated with poor exercise performance.

In the present study, elderly males accounted for most of the COPD patients who were admitted to an emergency department with acute exacerbation (16). According to a study that evaluated the spirometric parameters of patients, FEV, was lower than 40% in COPD patients who visited a hospital with acute exacerbation (17). Decrease in FEV, (%) is a factor that leads to hospitalization of COPD patients for acute exacerbation (18). In another study, impaired oxygenation was a factor that led to emergency department visits (19). In the present study, FEV, (%) and partial oxygen pressure were also significantly lower in COPD patients who visited an emergency department for acute exacerbation compared to the other group. In another study, which investigated risk factors related to the mortality of COPD patients with exacerbation, it was found that a change in partial arterial carbon dioxide pressure (PaCO₃) between the first measurement on admission and during hospitalization was associated with mortality (20). Emerman et al. (21) showed that although FEV, was lower than 35% in patients who visited an emergency department with acute exacerbation and whose PaCO, levels were higher than 45 mmHg, no correlation was determined between FEV, and PaO,. However, in the present study, PaCO, was significantly higher in patients who visited an emergency department.

In a study where COPD patients were followed for at least one year, increased dyspnea score was regarded to be an independent predictor of acute exacerbations (22). It was stated that COPD patients with low pulmonary functions and poor dyspnea scores returned to hospital within 30 days after hospital discharge (23). In another study,

dyspnea was considered to be a significant risk factor for hospital revisits (24). In the present study, dyspnea scores were also significantly higher in patients who visited an emergency department compared to the other group.

Chronic obstructive pulmonary disease patients experience a loss of exercise capacity compared to healthy controls independent of the disease stage (25, 26). The simple and valuable six-minute walk test is the most frequently used method to assess exercise performance (27). Patient compliance with the six-minute walk test is higher than with the cardiopulmonary exercise test (28). Studies have revealed that the body-mass index, airflow obstruction, dyspnea, and exercise (BODE) index is the best indicator to determine not only the number and severity of exacerbations and hospital visits, but also revisits and mortality; however, no existing study focuses directly on the correlation between acute exacerbation of COPD and six-minute walk distance (29-31). In the present study, the six-minute walk distance was significantly shorter in COPD patients who visited an emergency department compared to the other group. In one study, the dominant symptoms at peak exercise were dyspnea and leg fatigue in COPD patients (32). Consistent with this study, we found changes in dyspnea perception were higher and changes in peripheral oxygen saturation were lower post-test in patients who visited an emergency department.

Table 2. Comparison of exercise capacity between groups								
Variable	Emergency department visit (+) (n=127) Emergency department visit (n=79)		р					
6 MWD (meters)	320 (217, 375)	395 (315, 440)	<0.001*					
ΔHR (beats/minute)	19(12, 25)	14(9,22)	0.062					
ΔSpO ₂ (%)	-7(-10, -4)	-4(-6,-2)	0.029*					
ΔDyspnea (MB)	2(1, 3)	1(1, 3)	0.003*					
ΔLeg Fatigue (MB)	3(2, 4)	1(1,2)	<0.001*					

Data are expressed as median (interquartile range). Changes between post-test and baseline levels are shown as Δ values

*p<0.05 for comparison of changes between groups based on Mann-Whitney U test 6MWD: six-minute walk distance; HR: heart rate; SpO₂: peripheral oxygen saturation; MB: Modified Borg Scale

Table 3. Factors correlated with frequency of emergency department visits									
	6MWD	mMRC	FEV ₁ (%)	FEV ₁ /FVC	DLCO (%)	PaO ₂	SaO ₂		
EV	-0.301**	0.367**	-0.254**	-0.300**	-0.264**	-0.179*	-0.178*		
6MWD		-0.633**	0.368**	0.237**	0.320**	0.305**	0.261**		
mMRC			-0.410**	-0.353**	-0.424**	-0.380**	-0.343**		
FEV ₁ (%)				0.053**	0.557**	0.363**	0.373**		
FEV ₁ /FVC					0.524**	0.123	0.127		
DLCO(%)						0.148	0.221*		
PaO ₂							0.914**		

Spearman Correlation Test. *p<0.05, **p<0.01

EV: number of emergency department visits; 6MWD: six minute walking distance; mMRC: modified Medical Research Council; FEV,: forced expiratory volume in one second; FVC: forced vital capacity; DLCO: carbon monoxide diffusing capacity; PaO,: partial arterial oxygen pressure; SaO,: arterial oxygen saturation

Chronic obstructive pulmonary disease is characterized by restriction in progressive air flow, and pulmonary function tests are frequently used to evaluate COPD patients. FEV, is one of the best indicators of pulmonary emphysema and is traditionally used to grade the severity of COPD (33). However, exercise tolerance must be determined to assess the impact of COPD on the patient and to monitor alterations in exercise performance due to disease progression. The six-minute walk test is the most appropriate, simple, and reliable test for evaluation of exercise tolerance (29). A strong positive correlation has been determined between the six-minute walk test and FEV, (10, 11). Another study that determined the correlation between the six-minute walk test and spirometric parameters stated that the walk test can be used to monitor alterations in pulmonary function in patients with severe and very severe COPD (34). In another study, a correlation was found between the six-minute walk test and FEV,, but not FEV,/ FVC, after bronchodilation. It was suggested that the six-minute walk test can be used instead of a spirometer to determine the severity of COPD (12). Studies investigating the correlation of the six-minute walk test with spirometric parameters demonstrated significant correlations with FEV, (31) and DLCO (35, 36). In the present study, the six-minute walk test showed a positive correlation with FEV, (%), FEV,/FVC, and DLCO (%) values.

No study investigating the correlation between arterial blood gases values and the six-minute walk test is available. Most previous studies focused on the impact of the six-minute walk test on gas exchange (37, 38). It was demonstrated that six-minute walk distance is correlated with baseline O_2 saturation before and after testing (37). In a study suggesting that the six-minute walk test is a good determinant of prognosis in COPD patients, it was concluded that short walking distance and increased variability of O_2 saturation are independent and significant determinants of mortality (38). Similar to our study, changes in peripheral oxygen saturation, which reflect desaturation, were higher in patients who visited an emergency department.

In a study comparing hypercapnic and normocapnic COPD patients, the six-minute walk distance was found to be shorter in hypercapnic patients (39). In the present study, six-minute walk distance showed positive correlations with arterial saturation and PaO_2 but showed a negative correlation with $PaCO_2$. As in the present study, two other studies determined strong negative correlations between the mMRC scale and six-minute walk distance (40, 41).

This study has some limitations. Particularly, our study population has severe and very severe COPD; it does not include many patients with mild disease. Therefore, the results may not reflect all COPD patients. Investigating the correlation between emergency department visits and exercise performance in patients according to their disease stages may provide more accurate results and stronger correlations.

We could not record the last time of exacerbation because our patients visited other hospitals. Furthermore, some of our patients could not remember their last exacerbation event well. Therefore, we could not provide any information about the patients' most recent exacerbations, which may be important to determine final functional capacity.

Assessment of exacerbation severity is an important outcome to define the clinical features of patients. However, we could not use an effective tool to evaluate the severity of exacerbation and could only report the patients' number of hospitalizations, which indicated severe exacerbation.

The literature contains no study investigating the correlations between patient visits to an emergency department and acute exacerbation and exercise performance. However, some studies suggest strong correlations between regular physical activity and hospital visits (8) and revisits (42). It is stated that non-pharmacological interventions, such as education, smoking cessation, immunization, and pulmonary rehabilitation, in addition to pharmacological therapies, may prevent acute exacerbations in COPD patients (43).

CONCLUSION

Given the correlation between exercise performance and the number of emergency department visits determined in the present study, we conclude that non-pharmacological interventions such as regular daily physical activity and pulmonary rehabilitation may reduce the number of acute exacerbations in COPD patients.

Ethics Committee Approval: Ethics committee approval was received for this study from the ethics committee of University of Health Sciences Dr. Suat Seren Chest Diseases and Surgery Training and Research Hospital.

Informed Consent: Written informed consent was obtained from patients who participated in this study.

Peer-review: Externally peer-reviewed.

Author Contributions: Concept - İ.N., H.Ş.; Design - İ.N., H.Ş.; Supervision - İ.N., H.Ş.; Resources - H.Ş., I.K., N.A.; Materials - H.Ş., I.K., N.A.; Data Collection and/ or Processing - H.Ş., I.K., N.A.; Analysis and/or Interpretation - İ.N.; Literature Search - İ.N., H.Ş.; Writing Manuscript - İ.N., H.Ş.; Critical Review - İ.N., H.Ş., I.K., N.A.; Other - İ.N., H.Ş., I.K., N.A.

Conflict of Interest: No conflict of interest was declared by the authors.

Financial Disclosure: The authors declared that this study has received no financial support.

REFERENCES

- Wilkinson T, Wedzicha JA. Strategies for improving outcomes of COPD exacerbations. Int J Chron Obstruct Pulmon Dis 2006; 1: 335-42. [CrossRef]
- Wedzicha JA, Wilkinson T. Impact of chronic obstructive pulmonary disease exacerbations on patients and payers. Proc Am Thorac Soc 2006; 3: 218-21. [CrossRef]
- Dusser D. Chronic obstructive pulmonary disease exacerbations and their impact on the long- term natural history of the disease. Presse Med 2008; 37: 1599-603. [CrossRef]
- Chan FWK, Wong FYY, Yam CH, Cheung WL, Wong EL, Leung MC, et al. Risk factors of hospitalization and readmission of patients with COPD in Hong Kong population: Analysis of hospital admission records. BMC Health Serv Res 2011; 11: 186. [CrossRef]
- 5. Arnedillo MA. Impact of exacerbations and admissions in chronic obstructive pulmonary disease. Arch Bronconeumol 2010; 46: 8-14.
- Chabot F, Gomez E, Guillaumot A, Kheir A, Chaouat A. Acute exacerbations of chronic obstructive pulmonary disease. Presse Med 2009; 38: 485-95. [CrossRef]
- 7. Marchetti N, Criner GJ, Albert RK. Preventing acute exacerbations and hospital admissions in COPD. Chest 2013; 143: 1444-54. [CrossRef]
- 8. Bourbeau J. Preventing hospitalization for COPD exacerbations. Semin Respir Crit Care Med 2010; 31: 313-20. [CrossRef]

- Van Stel HF, Bogaard JM, Rijssenbeek-Nouwens LH, Colland VT. Multivariable assessment of the 6-min walking test in patients with chronic obstructive pulmonary disease. Am J Respir Crit Care Med 2001; 163: 1567-71. [CrossRef]
- Sachin RA, Rajnish J, Ajitprasad J. Correlation of severity of chronic obstructive pulmonary disease with health-related quality of life and six-minute walk test in a rural hospital of central India. Lung India 2015; 32: 233-40 [CrossRef]
- Kodavala AK, Dash S. Correlation between forced expiratory volume in first second (FEV1) and 6 minute walk distance in moderate, severe and very severe chronic obstructive pulmonary disease. IOSR-JDMS 2013; 2: 72-6. [CrossRef]
- Kundu A, Maji A, Sarkar S, Saha K, Jash D, Maikap M. Correlation of six minute walk test with spirometric indices in chronic obstructive pulmonary disease patients: A tertiary care hospital experience. J Assoc Chest Physicians 2015; 1: 9-13.
- Bestall JC, Paul EA, Garrod R, Garnham R, Jones PW, Wedzicha JA. Usefulness of the Medical Research Council (MRC) dyspnoea scale as a measure of disability in patients with chronic obstructive pulmonary disease. Thorax 1999; 54: 581-6. [CrossRef]
- 14. ATS Committee on Proficiency Standards for Clinical Pulmonary Function Laboratories. ATS statement: guidelines for the six-minute walk test. Am J Respir Crit Care Med 2002; 66: 111-7.
- 15. Borg GA. Psychophysical bases of perceived exertion. Med Sci Sports Exerc 1982; 14: 377-81. [CrossRef]
- Garcia-Sanz MT, Canive-Gomez JC, Alonso AS, Barreiro GA, Lopez VE, Senin RE, et al. Description of Hospital Admissions for Acute Exacerbation of COPD. J Pulm Respir Med 2014; 4: 200.
- Emerman CL, Effron D, Lukens TW. Spirometric criteria for hospital admission of patients with acute exacerbation of COPD. Chest 1991; 99: 595-9. [CrossRef]
- Mohapatra PR, Janmeja AK. Factors associated with hospital admission in patients with acute exacerbation of chronic obstructive pulmonary disease. Indian J Chest Dis Allied Sci 2010; 52: 203-6. [CrossRef]
- Garcia-Sanz MT, Pol-Balado Carlos, Abellas C, Canive-Gomez JC, Anton-Sanmartin D, et al. Factors associated with hospital admission in patients reaching the emergency department with COPD exacerbation. Multidisciplinary Respiratory Medicine 2012; 7: 6. [CrossRef]
- 20. Niksarlıoğlu EY, Arsava BE, Demir AU, İskit AT, Çöplü L. Risk Factors Associated with Mortality of COPD Patients Hospitalised for Exacerbation. Turk Toraks Derg 2013; 14: 134-40. [CrossRef]
- Emerman CL, Connors AF, Lukens TW, Effron D, May ME. Relationship arterial blood gases and spirometry in acute exacerbations of chronic obstructive pulmonary disease. Ann Emerg Med 1989; 18: 523-7. [CrossRef]
- 22. Müllerova H, Shukla A, Hawkins A, Quint J. Risk factors for acute exacerbations of COPD in a primary care population: a retrospective observational cohort study. BMJ open 2014; 4: e006171. [CrossRef]
- Guerrero M, Crisafulli E, Liapikou A, Huerta A, Gabarrús A, Chetta A, et al. Readmission for acute exacerbation within 30 days of discharge is associated with a subsequent progressive increase in mortality risk in COPD patients: a long-term observational study. PLoS One 2016; 11: e0150737. [CrossRef]
- Bahadori K, Fitzgerald JM. Risk factors of hospitalization and readmission of patients with COPD exacerbation- systematic review Int J Chron Obstruct Pulmon Dis 2007; 2: 241-51.
- 25. Çalık-Kutukcu E, Savci S, Saglam M, Vardar-Yagli N, Inal- Ince D, Arikan H, et al. A comparison of muscle strength and endurance, exercise capacity, fatigue perception and quality of life in patients with chronic obstructive

pulmonary disease and healthy subjects: a cross-sectional study. BMC Pulmonary Medicine 2014, 14: 6. [CrossRef]

- Akkoca Ö, Öner F, Saryal S, Şen E, Karabıyıkoğlu G. Hafif ve Orta Şiddetteki Kronik Obstrüktif Akciğer Hastalıklarında Egzersiz Kapasitesi. Tuberk Toraks 2001; 49: 5-12.
- 27. Spruit MA, Watkins ML, Edwards LD, Vestbo J, Calverley PM, Pinto-Plata V, et al. Determinants of poor 6-min walking distance in patients with COPD: The ECLIPSE cohort. Respir Med 2010; 104: 849-57. [CrossRef]
- Çiftçi F, Şen E, Akkoca-Yıldız Ö, Saryal S. A comparison of cardiopulmonary exercise test and six minute walking test in determination of exercise capacity in chronic obstructive pulmonary disease. Tuberk Toraks 2014; 62: 259-66. [CrossRef]
- 29. Marin JM, Carrizo SJ, Casanova C, Martinez-Camblor P, Soriano JB, Agusti AG, et al. Prediction of risk of COPD exacerbations by the BODE index. Respir Med 2009; 103: 373-8. [CrossRef]
- Kian-Chung O, Arul Earnest, Suat-Jin L. A multidimensional grading system (BODE Index) as predictor of hospitalization for COPD. Chest 2005; 128: 3810-16. [CrossRef]
- Ko FWS, Tam W, Tung AH, Ngai J, Ng SS, Lai K. A longitudinal study of serial BODE indices in predicting mortality and readmissions for COPD. Respir Med 2011; 105: 266-73. [CrossRef]
- 32. Ganju AA, Fuladi AB, Tayade BO, Ganju NA. Cardiopulmonary exercise testing in evaluation of patients of chronic obstructive pulmonary disease. Indian J Chest Dis Allied Sci 2011; 53: 87-91.
- 33. Sariaydın M, Altıntaş N, İnce Ö. Relationship between Lung Functions and Extent of Emphysema in Patients with Chronic Obstructive Pulmonary Disease. Eurasian J Pulmonol 2014; 16: 159-63. [CrossRef]
- Chen Hong, Liang B, Tang Y, Xu ZB, Wang K, Yi Q, et al. Relationship between 6-minute walk test and pulmonary function test in stable chronic obstructive pulmonary disease with different severities. Chinese Med J 2012; 125: 3053-8.
- Fujimoto H, Asai K, Watanabe T, Kanazawa H, Hirata K. Association of six-minute walk distance with resting pulmonary function in patients with chronic obstructive pulmonary disease. Osaka City Med J 2011; 57: 21-9.
- 36. Wijkstra PJ, TenVergert EM, Mark ThW, Postma DS, Van Altena R, Kraan J, et al. Relation of lung function, maximal inspiratory pressure, dyspnea, and quality of life with exercise capacity in patients with chronic obstructive pulmonary disease. Thorax 1994; 49: 468-72. [CrossRef]
- Agrawal MB, Awad NT. Correlation between six minute walk test and spirometry in chronic pulmonary disease. J Clin Diagn Res 2015; 9: 1-4. [CrossRef]
- Takigawa N, Tada A, Soda R, Date H, Yamashita M, Endo S, et al. Distance and oxygen desaturation in 6-min walk test predict prognosis in COPD patients. Respir Med 2007; 101: 561-7. [CrossRef]
- Sahin H, Naz I, Varol Y, Aksel N, Tuksavul F, Ozsoz A. Is a pulmonary rehabilitation program effective in COPD patients with chronic hypercapnic failure? Expert Rev Respir Med 2016; 28: 1-6. [CrossRef]
- 40. Manescu V. The relevance of the 6 minutes walking test and of dyspnea measured with mMRC scale in evaluating COPD severity. Pneumologia 2012; 61: 153-9.
- Vishteh HRK, Hajilari A, Masoudnia M, Ghorbani F, Shafaghi S, Najafizadeh K, et al. Comparison of patients' exercise capacity according to the MRC classifycation and 6-minute walking test. Eur Respir J 2012; 40: P3490.
- 42. Garcia-Aymerich J, Farrero E, Felez MA, Izquierdo J, Marrades RM, Antó JM, et al. Risk factors of readmission to hospital for a COPD exacerbation:a prospective study. Thorax 2003; 58: 100-5. [CrossRef]
- Criner GJ, Bourbeau J, Diekemper RL, Ouellette DR, Goodridge D, Hernandez P, et al. Prevention of acute exacerbations of COPD: American college of chest physicians and Canadian Thoracic Society guideline. Chest 2015; 147: 894-942. [CrossRef]