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Determining Factors of Distance During 6-Minutes Walk Test in COPD Patients

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KEYWORDS	A B S T R A C T
COPD; 6-minute walk test; Inspiratory capacity; Hyperinflation; Quality of life.	In stable COPD, identify and analyze the factors determining six minutes walking distance and dyspnea the end of the test. Patients were evaluated by test of lung function. St Georeges respiratoiry questionnaire (SGRQ) and 6 MWT with inspiratory capacity measurements (IC) continuous oxymetry. Eighty-two patients (mean FEV ₁ : $56\pm19\%$ predicted) were studied. Mean 6-minutes walking distance was $477 \pm 89m$ ($72 \pm 14PV$). Walking distance during 6MWT(m) was correlated with FEV ₁ , CI/TLC ratio, TLC, pré-test CI and DLCO/VA. When expressed as percent of predicted values, walking distance was correlated with FRC, pré-test IC and SGRQ activity score. End test dyspnea was correlated with FRC, pré-test dyspnea and SGRQ activity and total scores. The factors determining 6-minutes walking distance and end-test dyspnea are complex and include both functional and non functional factors. In COPD, 6MWT is thus an investigation that has additional integrative value.

Introduction

Chronic obstructive pulmonary disease (COPD) is characterized by not fully reversible airway obstruction. This obstruction, the most common cause is smoking, is gradually and severity is assessed primarily by the GOLD guidelines (Global initiative for chronic obstructive lung disease.). The degree of bronchial obstruction, appreciated by the expiratory volume per second (FEV) is a prognostic factor (Anthonisen *et al.*, 1994; Anthonisen *et al.*, 1986).

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But it has recently emerged that other factors, in particular functional, were decisive for evaluating the prognosis and survival of these patients. The index body mass index, airflow obstruction, dyspnea functional, exercise capacity (BODE) (Celli et al., 2004) composite index taking FEV, distance from the six-minute walk test (CT), degree of dyspnea index body mass is maintained routinely used to assess the prognosis of these patients; but also the impact of therapeutic interventions as the revalidation (Beaumont et al., 2011). TDM also reflects, more than ergospirometry on bicycle, activities of daily living (Guyatt et al., 1991; Garcia-Rio et al., 2009). It emerged recently that the poor performance in CT was associated with increased mortality in COPD with a lower FEV 50% predicted (VP) (Casanova et al., 2008).

The measures needed to perform quality YDM, as recommended by the American Thoracic Society (ATS) (Statement, 2002), are the qualification of the number of meters made and dyspnea before and end of the test. In this study, we evaluated in COPD in stable condition if respiratory functional factors, the development of a dynamic destruction or distension during the test (Callens *et al.*, 2009) or even the quality of life determine the distance and end of the test dyspnea.

Methods

Patients

From September 2013 to September 2014, all patients in stable condition BPCIO of GOLD stage I to IV, presenting an outpatient basis at the pulmonology department of the Brazzaville University Hospital for lung function tests were evaluated prospectively. They were selected based on their medical history, their habits and tabagismes spirometry results showing FEV / vital capacity (VC) less than 70%. None had presented exacerbation in the previous month.

Informed consent was obtained. The study had previously been accepted by the ethics committee of the Medical Society of the Congo.

Measures

First Evaluation

Pulmonary function tests were performed in all patients. They completed the Respiratory Questionnaire St. George's (SGRQ) (Piperno *et al.*, 2000), followed by TDM habituation, conducted according to the ATS guidelines. The standards used to evaluate performance TDM are those of Troosters *et al.* (1999)

Second Evaluation

During the same week, a second CT scan was performed, with additional measures of inspiratory capacity (IC) and continuous oximetry.

A first IC measurement was performed at rest, before MDA. The CI was measured using a portable spirometer (Spiro BT 100, Medical Electronic Construction, Brussels, Belgium) after patient education and training. After obtaining from April to June volumes end expiration successive stable, we asked them to inspire total lung capacity (CTP). then breathe normally. The measurements were repeated for a maximum variation of 5% on two consecutive measurements, and we retained the highest value (Marin et al., 2001). At the end of TDM, a measure of CI was also carried out in the corridor, immediately after walking,

using the same portable spirometer. Dyspnea was assessed before and after MDD with the Borg Scale (Borg *et al.*, 1982).

The distance of the CT and end of the test dyspnea were correlated with the severity of airway obstruction (expiratory volume in one second) to distention (functional capacity (FRC), total lung capacity (CPT) and CI report / CPT), the dynamic distension during the test (modification of the CI during the test), the oxygen desaturation and quality of life (SGRQ).

Statistical Analysis

The results are expressed as mean \pm standard deviation. Comparing the quantitative data was carried out by a paired t test. Univariate analyzes were performed by calculating the Pearson correlation coefficient, in order to study the existence of a relationship between various clinical and functional variables and walking distance (in meters and percentage VP) or the end of dyspnea test.

Multivariate analysis was performed using the method of logistic regression to determine step by step what the variables independently correlated with walking distance (in meters and percentage VP) and test end dyspnoea.

The differences were considered statistically significant when the critical value of uncertainty p is less than 0.05.

Results and Discussion

A total of 82 patients (59 men and 23 women) were studied COPD. All were (former) smoking and classified GOLD stages I to IV. None were oxygeno dependent. The average age was 62 ± 10 years. The mean FEV was 1.6 ± 0.6 (56 ± 18

of the VP). The SGRQ score was $41 \pm 20\%$ for the item " symptoms " and 59 ± 23 for " activities ". Main functional and anthropological data are summarized in Table 1.

The distance at the end of CT was 477 ± 89 m, or $72 \pm 14\%$ of the VP and CT was associated with a significant increase in dyspnea (0.4 ± 0.7 at the beginning of test 3, 9 ± 2.1 end of the test; p<0,0001) and a significant increase in O2 saturation ($94 \pm 11-91 \pm 5\%$; p 0.0001). For the population as a whole, the TDM was accompanied by the development of a dynamic distension since the CI decreased from 2.2 ± 0.6 L standing at 1.8 ± 0.7 L (p<0,0001) late test, or $111 \pm 34-89 \pm 30\%$ of the VP (p<0,0001). Nineteen of 81 patients (23%) did not develop dynamically distension to the effort.

Predictors level dynamic distension in Δ CI efforts as percentage of VP) were FEV, a percentage of the VP (p = 0.0026) and the FIU in liters (p = 0.0029). All parameters measured during MDA are summarized in Table 2.

Table 3 summarizes the significant relationships in univariate analysis, and Table 4 in multivariate analysis with the distance in meters and percentage of the VP, and dyspnea end CT.

The end dyspnea MDD was, she, well correlated with FEV, VP percentage (r = -0.37), with the FIU, VP percentage (r = 0.33) with the Δ CI early CT; L (r = 0.26) and with the carbon monoxide diffusing capacity (DLco), percentage VP (r = -0.25).

In multivariate analysis, the distance covered in meters CT correlated with FEV (L), CPT (L), the IC (L) before the test and the report DLCO / VA (Table 4); these four variables explained 35% of variance of CT in meters. This same distance as a percentage of the VP was significantly correlated to the FIU (L and percent predicted), the CI (L) before the test, and the score of Saint George's activity. These four variables explained 38% of variance of CT as a percentage of the VP.

Dyspnea at the end of TDM was about that correlated with the FIU (L), with the start of test and dyspnea with activities and total SGRQ scores (Table 4); and 35% of the variance was accounted for by these variables 4.

This study suggests that the co-variables that are related to the distance covered during the 6-minute CT in patients with COPD include FEV, the FRC and TLC, IC resting DLCO / VA and score Saint-Georges on. For dyspnea at the end of TDM, the study found as covariates of the FIU, dyspnea before the test and score Saint-Georges and total activity.

Our study, however, suffers from limitations due to the small sample size, but also because some parameters that could affect the results of the walk test as exacerbations, the degree of emphysema, body composition, muscle strength devices, cardiac dysfunction induced by distension have investigated. not been These limitations invoked do not affect completely the power of our observations. This study is in any case the first of its kind in our environment, including proven patients.

Some differences in the variables related to the distance to the TDM absolute or percentage of the VP called a comment. For example, FEV is a determinant of the distance traveled in meters; but it is no longer when it is expressed as a percentage of the VP. In reality, the walk in the normal individual is related to age, gender and size, factors that also determine lung volumes so it is not surprising that their decisive less evident in the distance expressed in percentage of the VP.

Apart from the effect p = 0,0026t expected FEV on the walking distance, we observed that the parameters indicating a distension as the CPT, the FIU, the IC off and the IC / TLC ratio are correlated negatively with the walk, in agreement with observations of other authors. Similarly, DLCO / VA correlates with distance. Together, these observations suggest that the emphysema phenotype is associated with worse performance in the walk test. In our study, no emphysema evaluation CT data is available to confirm this; but analysis of da ECLIPSE cohort as other studies (Spruit et al., 2010; Lee et al., 2008) show the scope or extent of emphysema measured by CT scan are correlated negatively with the distance. We found no correlation between desaturation during the TDM and distance (or dyspnea); but note that no patient suffering from resting hypoxemia and desaturation than the average observed in the effort was modest.

We have not found any correlation between the parameters that reflect a dynamic test at the end of distension as the CI end TDM or ΔCI over TDM and distance or the end of the test dyspnea although the majority of our population with COPD (77%) actually develops a dynamic distension during the walk test. This observation is consistent with the results obtained by Callens et al., in 52 patents COPD, but contradict the data of Marin et al., obtained in 72 COPD patients where ΔCI correlated with the performance and dyspnea associated with the walking test. Regarding the dynamics distension, this difference seems apparent to the extent that the CI rest is raised by all these studies, such as ours (CI rest is same the only significant parameter in absolute and VP) as a predictor of distance.

Table.1 Anthropometric characteristics and plethysmographic of 82 patients
bronchoneumopathie chronic obstructive pulmonary disease (COPD) and results in respiratory
questionnaire St. George's

	Average	Standard Deviation
Age (years)	61.91	9.92
BMI kg/m ²	24.95	5.004
Smoking (pack-years)	50.5	27.61
FEV (L)	1.569	0.5794
FEV (%% predicted)	56.37	18.67
CRF (L)	4.71	1.381
CRF (%% predicted)	146	40.17
CPT (L)	6.835	1.32
CPT (% predicted)	114.9	20.06
Saint-George's TOTAL (%)	41.61	19.56
Saint-George's syndromes	45.61	22.94
Saint-George's activity	58.65	23.43
DCLO (% predicted)	52.60	17.03
DCLO/VA	61.22	20.82
Sex	59/23=2,6	

Abbreviations: BMI, body mass index; FEV, maximum expiratory volume; FRC, functional residual capacity, CPT, total lung capacity; DCLO, carbon monoxide diffusing capacity.

Table.2 Results of the measurements taken during the walk test (CT) scans of 82 patients with chronic obstructive bronchoneumopathie (COPD)

	Average	Standard Deviation
Distance (m)	477.3	89.43
Distance (% prédit)	72.25	13.82
CI max rest (L)	2.234	0.624
CI max rest (% predited)	112.2	34.43
CI end TDM (L)	1.814	0.6557
CI end TDM (% predited)	89.19	29.66
$\Delta CI (L)$	0.4922	0.3509
∆CI (% predited)	25.73	20.77
Borg pré-TDM	0.4074	0.7379
Borg post-TDM	3.938	2.117
∆Borg	3.556	2.055
Saturation end pre-TDM (%)	90.98	5.02
Saturation pré-TDM (%)	95.02	1.844

Abbreviations: CI, inspiratory capacity; Borg, Borg scale.

	Distance traveled TDM (m)		Distance traveled TDM (%)		Dyspnea and TDM (Borg)	
	r	р	r	Р	r	р
FEV (L)	0.54	< 0.001	0.33	0.003	0.27	0.01
FEV (% predited)	0.49	< 0.001	0.53	< 0.001	0.40	< 0.001
CRF (L)	0.28	0.01	0.41	< 0.001	0.35	< 0.001
CRF (% predited)	0.27	0.02	0.31	0.005	0.35	0.001
CPT (L)	0.13	0.24	0.34	0.002	0.30	0.002
CPT (%)	0.22	0.05	0.20	0.08	0.01	0.007
CI rest (L)	0.40	< 0.001	0.22	0.05	0.23	0.93
CI rest (%)	0.04	0.70	0.08	0.48	0.19	0.04
CI end TDM (L)	0.42	< 0.001	0.26	< 0.001	0.01	0.92
CI end TDM (%)	0.13	0.27	0.09	0.02	0,26	0.02
ΔCI (L)	0.06	0.59	0.07	0.45	0.33	0.003
ΔCI (%)	0,21	0.06	0.21	0.52	0.19	0.09
CI/CPT	0.48	< 0.001	0.44	0.06	0.48	0.05
Sa0 ₂ pré-TDM	0.30	0.007	0.20	< 0.001	0.22	0.05
DCLO (%)	0.21	0.10	0.25	0.07	0.25	0.31
DCLO/VA	0.01	0.93	0.01	0.04	0.13	0.01
Borg pre-TDM	0.22	0.05	0.28	0.92	0.28	0.14
Borg post-TDM	0.14	0.21	0.20	0.07	/	/
∆Borg	0.09	0.45	0.12	0.27	/	/
QRSG activities	0.45	< 0.001	0.52	< 0.001	0.47	< 0.001
QRSG symptomes	0.24	0.03	0.33	0.003	0.30	0.006
QRSG total	0.42	< 0.001	0.53	< 0.001	0.36	0.001
IMC	0.20	0.07	0.04	0.72	0.20	0.84

Table.3 Correlations significant in univariate analysis between the distance to the walking test (CT) in meters or percentage of predicted or dyspnea at the end of TDM, assessed by the Borg scale and various parameters

Abbreviations: CI, inspiratory capacity; CPT, total lung capacity; FEV, maximum expiratory volume; FRC, functional residual capacity; Sa02 pre-TDM, oxygen saturation; DCLO, carbon monoxide diffusing capacity; DCLO / VA, diffusing capacity for carbon monoxide reduced to alveolar volume; SGRQ, Respiratory Questionnaire of St. George; BMI, body mass index.

	Distance traveled TDM (m)	Distance traveled TDM (%)	Dyspnea and TDM (Borg)
FEV (L)	0.0036	NS	NS
CRF (L)	NS	0.0009	0.034
CRF (% predited)	NS	0.0151	/
CPT (L)	0.0016	NS	NS
CI rest (L)	0.0035	0.0215	NS
DCLO/VA	0.0070	NS	NS
Dyspnea pre-TDM (Borg)	NS	NS	0.0562
QRSG Activities	NS	< 0.0001	0.0006
QRSG Total	NS	NS	0.0258

Table.4 Correlations significant in univariate analysis between the distance to the walking test (CT) in meters or percentage of predicted or dyspnea at the end of TDM, evaluated by Borg scale and various parameters

Abbreviations: CI, inspiratory capacity; CPT, total lung capacity; FEV, maximum expiratory volume per second; FRC, functional residual capacity; DCLO, carbon monoxide diffusing capacity; SGRQ, Respiratory Questionnaire St. George.

Indeed, the rest of CI also integrates dynamic distension as illustrated by the fact that salbutamol in our population of COPD, improves CI rest when patients have a limitation of expiratory flow and does not affect it in the condition reverse (Boni *et al.*, 2002).

Several factors could explain then that the test end IC has no predictive value in our study. The explanation does not seem linked to a greater propensity to develop a dynamic distension in the study by Marin et al., On the contrary, the decrease of our population CI late effort was greater (-492mL versus -271mL). is possible that It the methodological differences in the measurement of the end of the test ICs explain these conflicting results. End of the test, a single measurement was performed in our study while Marin et al. obtained the average of two measurements. L a single measure, however, seems better in this particular circumstance, because it was (Marin 2001) assumed et al., and demonstrated that CI is normalized in less than 3 minutes after exercise and therefore repeated measures were training a sub-

estimation of dynamic hyperinflation. The team of Yan et al., has also demonstrated by esophageal pressure measurements at rest and at year end, in 15 COPD, that the efforts made during the IC maneuvers were reliable, reproducible and allowed to properly estimate changes in volume endtidal during exercise. Recent observations of Guenette et al., are in agreement with our results and serve to dispel commonly held opinion that the development of a dynamic effort to distension is related to the endurance time and dyspnoea. In this study, the authors compared 65 COPD who develop dynamic distension exercise and 65 did not develop in FEV equivalent, and found that dyspnea and endurance time were similar in both groups. Dyspnea was determined by the level of the slope inflection VT / VE, otherwise similar between the two groups and not by the level of distension (Guenette et al., 2001).

Our study also found, in agreement with recent data (Spruit *et al.*, 2010), a correlation between the quality of life (essentially the score of the SGRQ activity) and the distance of the walk test or dyspnea. This observation

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is not surprising inasmuch as the assessment of quality of life brings together into a single variable the overall impact of COPD. Multivariate analyzes also show that dyspnea score assessed by the Medical Research Council and the distance traveled in CT are part of the five factors (the others are coughing, wheezing and anxiety score) that help explain 50 % of the variance of the SGRQ (Jones, 2001).

In conclusion, the factors that determine the performance and dyspnea during the sixminute walk test are complex and depend on functional and non-functional factors. The six-minute walk test thus appears as a test of exploration integral value in COPD.

Declaration of Interest

The authors declare having no conflicts of interest related to this article.

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