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## CAN HEALTH STATUS QUESTIONNAIRES BE USED AS A MEASURE OF PHYSICAL ACTIVITY IN COPD PATIENTS?

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### Take home message

Health status questionnaires provide only limited insight in the physical activity of patients with COPD

*To the editor*

Acting to address the amount of physical activity (PA) of patients with Chronic Obstructive Pulmonary Disease (COPD) is recommended as part of the GOLD recommendations [1]. A level lower than approximately 5000 steps ('sedentary lifestyle index') is associated with significantly increased health risks [2,3]. Actively screening to identify patients below this threshold could be an important step towards targeting interventions to increase PA. Potential screening tools include activity monitors (objective assessment) or self-reported questionnaires (subjective assessment) to measure the amount of PA [4]. The latter method, feasible in clinical practice, may not result in an accurate representation in an individual patient, as questionnaire responses tend to misclassify PA [5]. Although more accurate and widely used in research, activity monitoring is not yet commonly included in patients' routine assessment. Several health status questionnaires in routine clinical use, contain a domain or dimension related to PA [6]. In the analytical framework of Leidy, et al. functional performance has been defined as the physical, psychological, social, occupational and spiritual activities that people actually do in the normal course of their lives to meet basic needs, fulfill usual roles, and maintain their health and well-being [7]. A review by Kocks, et al., proposed that these could be used in the measurement of functional performance (defined as 'what a patient is actually doing') and that this would be a more practical alternative to PA monitoring [8]. Whether this approach is sufficiently valid as a representation for PA levels in clinical practice remains to be established.

The aims of the present analyses were (i) to describe the relationship between objectively measured PA and responses to health status questionnaires and (ii) to assess the utility of these questionnaires to screen for severe physical inactivity (SPI) in this population.

Data from two hundred thirty-five COPD patients (diagnosis confirmed with a post-bronchodilator spirometry) recruited from five centers across Europe [Leuven(Belgium), Athens (Greece), Groningen (The Netherlands), London and Edinburgh (UK)], as part of the PROactive project

([www.proactivecopd.com](http://www.proactivecopd.com)) were included in this analysis. Patients were current or ex-smokers (+10 pack years) without comorbidities significantly interfering with their ability to exercise and without respiratory conditions other than COPD. Further study details are available in the paper reporting the primary analyses of the present study [9]. PA was measured for 14 days during waking hours using the Dynaport Movemonitor (©McRoberts; The Hague, the Netherlands), which has been validated in this population [10,11]. Patients with a minimum of four valid days, defined as days with at least eight hours of wearing time, were included in this analysis [12]. SPI was defined as a step count lower than 5000 per day [2]. At the end of this measurement period, several questionnaires were administered including: Chronic Respiratory Disease Questionnaire Self-administered Standardised Format (CRDQ-SAS), COPD assessment test (CAT), Clinical COPD questionnaire (CCQ) and modified Medical Research Council dyspnea questionnaire (mMRC). Relations between PA (number of daily steps, STEPS) and different health status questionnaires were investigated using Spearman correlations. The ability of questionnaires to predict SPI was analyzed using logistic regression analysis with SPI as outcome and different questionnaires as explanatory variables, each included in a separate analysis. Area under the curve (AUC) was retrieved if the regression analysis was significant. AUC values are a measure of accuracy and are considered as excellent ( $\geq 0.90$ ), good (0.80-0.89), fair (0.70-0.79) or poor ( $< 0.70$ ). ROC curves were drawn for the different questionnaires, including total scores and the subdomain with the best discriminative property. A cutoff was suggested for each of the questionnaires (using the total score or subdomain, giving the highest AUC) putting an equal weight to sensitivity and specificity. Likelihood ratios were calculated as the ratio between sensitivity and (1-specificity). Likelihood ratios above 10 are considered to provide strong evidence for diagnostic purposes [13].

Nine patients did not have a valid PA measurement and were excluded from the analysis, one patient did not complete the questionnaires, resulting in 225 patients (67% male, mean $\pm$ SD of 67 $\pm$ 8 years, FEV<sub>1</sub> 56 $\pm$ 20%<sub>pred</sub>, 6MWD 426 $\pm$ 129m (68 $\pm$ 19%<sub>pred</sub>), with a median [Q1-Q3] step count of 4287 [2971-6331] steps.day<sup>-1</sup> representing all GOLD stages I/II/III/IV (12%/47%/32%/9% respectively) included in

the present analyses. Sixty percent of patients were defined as severely inactive. The health status questionnaires (and their subdomains)  $CCQ_{\text{symptoms}}$ ,  $CRDQ-SAS_{\text{mental}}$ ,  $CRDQ-SAS_{\text{mastery}}$  were poorly correlated ( $abs(r) < 0.3$ ); CAT,  $CCQ_{\text{total}}$ ,  $CCQ_{\text{mental}}$ ,  $CCQ_{\text{functional state}}$ ,  $CRDQ-SAS_{\text{total}}$ ,  $CRDQ-SAS_{\text{dyspnea}}$ ,  $CRDQ-SAS_{\text{fatigue}}$  were weakly related ( $0.5 < abs(r) \leq 0.3$ ); the mMRC score was moderately related ( $r = -0.52$ ) to STEPS ( $p < 0.01$  for all). All scores, except from the  $CRDQ-SAS_{\text{mental}}$  score, were significant predictors of SPI ( $p \leq 0.01$  for all). Only mMRC and  $CCQ_{\text{functional state}}$  scores showed a fair discriminative property (AUC respectively 0.719 and 0.724). Other questionnaires resulted in a poor discrimination ( $AUC < 0.7$ ). ROC curves were built for the different questionnaires including the total score and the best discriminative subdomain (see Figure 1). To predict SPI, a mMRC score  $\geq 2$  resulted in a positive predictive value (PPV) of 79%, accuracy (ACC) of 67% and positive likelihood ratio (LR+) of 2.27. A CAT score  $\geq 13$  resulted in a PPV of 73%, ACC of 64% and LR+ of 1.78. A  $CCQ_{\text{functional state}}$  score  $\geq 1.5$  results in a PPV of 76%, ACC of 67% and LR+ of 2.06. A  $CRDQ-SAS_{\text{dyspnea}}$  score  $\leq 5.3$  results in PPV of 76%, ACC of 68% and LR+ of 2.10.

The present data show that responses to health status questionnaires are only loosely related to the amount of PA and their use cannot therefore be recommended as a *standalone* screening tool for SPI because of low sensitivity and specificity. Two explanations can be proposed for the very low to moderate associations observed between these questionnaires which mainly capture symptoms and PA. First, none of these questionnaires use PA as a main concept and thus they are not designed to give a reliable assessment of the PA level [6]. The questionnaires mainly reflect symptoms that are indirectly associated with PA. Second, it is plausible that there is an interplay between the volume (amount\*intensity) of PA and the symptom experience. Patients may decrease their level of PA to avoid symptoms. This leads to the hypothesis that symptoms can depend on the PA level and therefore fail to give a reliable estimate of the actual underlying PA level. Indeed, patients' perception of PA includes not only the amount of PA but also symptoms experienced during and adaptations related to PA [14,9].

A good screening tool for inactive patients would be able to identify truly inactive patients (sensitivity) without including too many active patients (specificity). Depew, et al. concluded that the mMRC is the best predictor of SPI, compared to two PA questionnaires, self-efficacy and the ADO-index [15]. These authors proposed a mMRC score of 3 as a triage for SPI (defined as  $PAL < 1.4$ , i.e. the ratio of active to total energy expenditure ) [15]. Above this cutoff, 84% of patients identified as severely inactive were indeed inactive (PPV 84%). However, it resulted in a sensitivity of 36% meaning that only the minority of inactive patients were identified by this screening tool. Based on the ROC analysis, for mMRC we chose a cutoff of 2 points, which also resulted in a majority of patients above this threshold to be inactive (PPV 79%) with a sensitivity of 65%. These results suggest that this cutoff could be used in clinical practice as a first screening tool to identify severely inactive patients. However, a significant proportion of patients with an mMRC of 0 or 1 are inactive (NPV 56%). Therefore, PA should still be measured to identify inactive patients with a mMRC below 2. A comparable conclusion can be drawn based on the  $CCQ_{functional\ state}$  domain, using a cut off  $\geq 1.5$  (PPV 76%, NPV 57%). In the review of Kocks, et al., [8] the questionnaires were judged mainly based on the use in a primary care setting (e.g. practical use, responsiveness). These authors also concluded the mMRC and  $CCQ_{functional\ state}$  to be the most suited questionnaires to measure functional performance [8]. The present study shows that this conclusion only holds true in terms of PPV whereas NPV is poor.

The use of simple clinical tests, e.g. six minute walk test, were also shown to fail in predicting SPI [16]. Taking all this into account we can conclude that neither health status questionnaires, nor simple clinical tests can replace objective measurement of PA in COPD patients. The mMRC dyspnea and  $CCQ_{functional\ state}$  score could be recommended as easy first screening tests to identify severely inactive patients but will misclassify patients as 'not severely inactive' below the proposed thresholds. Therefore, objective PA measurement should be recommended in the clinical routine assessment of COPD patients.

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### Figure caption

**Figure 1** ROC analyses of modified Medical Research council dyspnea questionnaire (mMRC), with a range 0 to 4 [panel A], COPD assessment test (CAT), with a range 0-40 [panel B], total score (dotted line) and functional state domain score (solid line) of the Clinical COPD questionnaire (CCQ), with a range 0 to 6 for both [panel C]; total score (dotted line) and dyspnea domain score (solid line) of the Chronic Respiratory Disease Questionnaire self-administered standardized format (CRDQ-SAS), with a range 1 to 7 for both [panel D]. mMRC $\geq 2$  resulted in a sensitivity (SENS) of 65%, specificity (SPEC) of 71%; CAT $\geq 13$  resulted in 64% SENS and 64% SPEC, CCQ<sub>functional state</sub>  $\geq 1.5$  in 66% SENS and 68% SPEC, CRDQ-SAS<sub>dyspnea</sub>  $\leq 5.3$  in 67% SENS and 68% SPEC.