Evaluation of the 3-minute chair rise test as part of preoperative evaluation for patients with non-small cell lung cancer

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Abstract

Background: Peak oxygen uptake (VO2peak) measured by a cardiopulmonary exercise test (CPX) is the gold-standard for predicting surgical risk in patients with non-small cell lung cancer (NSCLC). The 3-minute chair rise test (3CRT) is a simple test requiring minimal resources. This study aimed to determine the ability of 3CRT to predict VO2peak in patients with NSCLC.

Methods: Retrospective data from CPX and 3CRT carried out in 36 patients with NSCLC between March 2018 and February 2019 were included. A multivariate analysis was undertaken to derive a predictive VO2peak equation based on performance on the 3CRT. In addition, sensitivity-specificity analysis was carried out to estimate a threshold 3CRT value for the prediction of VO2peak ≥ 15 mL/kg/minute.

Results: The following equation was obtained: VO2peak predicted = (0.04765 × FEV1) - (0.207 59 × BMI) - (0.115 89 × age) + (0.386 09 × vertical distance) + 16.628 69; r² = 0.75, P<0.01. The bias between the VO2peak values predicted and measured during CPX was 0.0 ± 1.7 mL/kg/minute (95% limits of agreement [−3.5 to 3.5]). A performance ≥49 chair rises predicted VO2peak ≥ 15 mL/kg/minute with a sensitivity of 0.75 and a specificity of 0.81.

Conclusions: The level of error in the prediction of VO2peak from 3CRT performance was too great to recommend that 3CRT should replace CPX as the sole measurement of VO2peak. Nevertheless, the 3CRT could help to identify those patients that require CPX prior to lung resection surgery for NSCLC, larger prospective study is needed to confirm this hypothesis.

Key points
Significant findings of the study: Cardiopulmonary exercise tests can stratify the surgical risk. Prediction of the peak oxygen uptake (VO2peak) value from the
3CRT yields an unacceptable level of error. However, a performance of 49 chair rises or more during the 3CRT could indicate a $VO_2\text{peak} \geq 15 \text{ mL/kg/minute}$. **What this study adds:** The 3CRT is a useful screening tool to determine the necessity for a comprehensive cardiopulmonary exercise test, whose access is limited in clinical practice. It could also allow early screening of patients requiring specific prehabilitation programs.

**Introduction**

Pulmonary resection is the most effective treatment for non-small cell lung cancer (NSCLC).1 Unfortunately, many patients with lung cancer also have numerous comorbidities that increase the risk of postoperative complications.2 The incidence of postoperative complications after lung cancer surgery is very heterogeneous according to the studies, it is estimated to be around 5% to more than 40%.3 These complications are associated with significantly poorer long-term outcomes.4 For this reason, all patients with NSCLC undergo comprehensive preoperative assessment to determine the benefits and risks of surgery in order to make appropriate therapeutic decisions.

Cardiopulmonary exercise test (CPX) is an important part of the preoperative assessment. It is used to measure and calculate variables that have been shown to be powerful indicators of the postoperative risk of morbidity and mortality. These variables include maximal oxygen uptake ($VO_2\text{max}$) or, if maximal effort is not achieved, peak oxygen uptake ($VO_2\text{peak}$).5 6 Although the oxygen uptake capacity is now considered as the gold standard measurement to determine postoperative risk, access to CPX may be limited in some centres due to limitations in equipment and resources. As a result, other simpler tests have been developed.

The shuttle walking test (SWT) and the stair climbing test (SCT) are both currently recommended as screening tools to determine the necessity to perform CPX.7 Performance of 25 shuttles (ie, 5 minutes or 250 meters) in the SWT has been shown to be predictive of a $VO_2\text{peak} \geq 15 \text{ mL/kg/minute}$.8 For the SCT, an ascension of more than 22 meters has been associated with a $VO_2\text{peak} \geq 15 \text{ mL/kg/minute}$ and fewer postoperative complications.9 10 Therefore, these tests are useful to identify patients at low risk of postoperative complications. Patients who are at higher risk of postoperative complications and who do not achieve these test results often have a $VO_2\text{peak} < 15 \text{ mL/kg/minute}$.5 11 Those patients require CPX in order to determine their risk more precisely; a $VO_2\text{peak} < 10 \text{ mL/kg/minute}$ is considered a high surgical risk.7 12

Although the SWT and SCT are relatively simple tests, they still require space and stairs respectively. A new test, the semi-paced 3-minute chair rise test (3CRT), has recently been standardized and validated in patients with chronic obstructive pulmonary disease.13 It has the advantage of requiring very little space and only a chair and a pulse oximeter. Therefore, the 3CRT can be performed in nearly any care setting, including very limited spaces such as consultation rooms. However, the results of 3CRT have not yet been compared with the definitive data obtained by CPX in patients with NSCLC, as has been undertaken for the SWT and SCT methods. This could allow the method to be used as a practical preoperative screening tool.

The aim of this study, therefore, was to investigate the relationship between the measurement of metabolic power by CPX ($VO_2\text{peak}$) and the measurement of mechanical power developed during the 3CRT. More specifically, the aim was to determine the ability of 3CRT to predict the $VO_2\text{peak}$ in patients with NSCLC and/or its ability to identify patients with a $VO_2\text{peak} < 15 \text{ mL/kg/minute}$ in order to provide indications regarding either suitability for surgery, or the need for further testing.

**Methods**

**Population and design**

All patients in this paper were retrospectively included in the analysis. The institutional ethics committee of the Rouen University Hospital approved the data extraction for this study (n°E2019-67). Patients all had been diagnosed with NSCLC and had either a predicted postoperative forced expiratory volume in 1 second (ppoFEV1) or a predicted postoperative diffusing capacity for carbon monoxide (ppoDLCO) < 60%,7 and been referred for complementary tests by thoracic surgeon or pneumologist. The predicted postoperative values of FEV1 and DLCO were estimated by preoperative value/total number of functioning segments before the operation, andx the residual number of functioning segments after the operation.12 The patients had undergone CPX as part of their preoperative assessment for pulmonary resection between March 2018 and February 2019. In addition, they had all also been assessed using 3CRT in
the physiology department of Rouen university hospital, France, since the department was trialing the test to determine if it was clinically feasible. All tests were carried out by the physiology medical team trained in carrying out stress assessment tests.

**Assessments**

**Pulmonary function**

Pulmonary function tests were carried out according to the American Thoracic Society and the European Respiratory Society guidelines using plethysmography (Masterscreen, Jaeger, Wittsburg, Germany). Values were expressed as percentages of established theoretical values for European populations.

**Cardiopulmonary exercise test**

The CPX room was air-conditioned and kept at a constant temperature of 19°C for all tests, and the hygrometry was not controlled in current practice. The measuring instruments were recalibrated every half day with a reading of the atmospheric pressure. Patients were advised to avoid strenuous exercise and the consumption of alcohol, tobacco or caffeine for 24 hours prior to examination. CPX was performed on an electromagnetic ergometer (Ergoselect 200, Ergoline, Bitz, Germany) using a standardized, incremental protocol. Following a three-minute warm-up, incremental ramp exercise (5 to 15 W/minute) was maintained until exhaustion. Markers used to consider the CPX as maximal were determined according to current guidelines. A face mask (Hans Rudolph, Inc., Kansas City, MO), pneumotach and gas analyzer (Ergocard, Medisoft, Louvain, Belgium) were used to measure breath-by-breath oxygen uptake (VO₂) and carbon dioxide production (VCO₂). Heart rate (HR) was continuously monitored with a 12-lead electrocardiogram (Ergocard, Medisoft, Louvain, Belgium).

**3-minute chair rise test (3CRT)**

The 3CRT was performed in the same condition (CPX room) according to the methodology validated by Aguilaniu et al. Test instructions and instructions for rating the 0 to 10 points Borg scale (dyspnea and fatigue) were given while the patient was sitting at rest on a standard chair (floor to seat height 48 cm); this chair was then used for the test: “The aim of the exercise is to stand up and sit down repeatedly for three minutes. You must keep your hands on your hips all the time. I’ll give you the pace for the first minute by saying: “sit”, “stand” or by doing the sit-to-stand with you. During the last two minutes try to stand up and sit down as many times as possible. I’ll let you know how much time is left, for example: “1 minute and 30 seconds to go”, “1 minute to go”, “30 seconds to go”. You can slow down or speed up at any time. If the exercise is too difficult, you can have a little rest then continue. You can also stop the test before the end of three minutes if you feel too tired or breathless to finish it.” A chair rise was defined as one complete movement from sitting to standing and then sitting again (Fig 1).

Depending on the patient’s FEV1 value, the pace during the first minute was set to ensure that either 12 (FEV1 < 1200 mL) or 20 (FEV1 ≥ 1200 mL) rises were performed, in order to reduce interindividual variability at the start of the test. At the end of the test, physiological data...
(SpO₂, HR) and symptoms (dyspnea and leg fatigue with the Borg scale) were recorded. The number of chair rises was also recorded and two variables were calculated: (i) the vertical distance reached (in meters) = (patient’s standing height - sitting height) x number of chair rises and (ii) the weight displaced (patient’s weight x the number of chair rises).

**Statistical analysis**

Categorial data are expressed as numbers (percentages). Continuous data are expressed as means ± standard deviation or medians (interquartile range) according to their distribution; normality was assessed using the Shapiro-Wilk test. Paired t-tests or Wilcoxon matched-pairs signed rank tests were used for the comparison of continuous data depending on the normality of the distributions.

In order to predict the VO₂peak, a multiple regression was undertaken using VO₂peak as the dependent variable. First, the univariate relationships between the VO₂peak measured during CPX and each demographic variable (age, BMI and FEV1) and each performance outcomes of the 3CRT (number of rises, vertical distance reached and weight displaced) were assessed using a Pearson’s correlation test. The variables identified as related to VO₂peak with a P-value <0.10 were assessed for interrelationships and entered in a multivariate analysis. In the case of a strong interrelationship (r > 0.80), only the variable most related with VO₂peak was used. Finally, a multivariate forward/backward stepwise regression was used to derive the best model to predict VO₂peak. Agreement between the measured VO₂peak (during CPX) and predicted VO₂peak was evaluated using a Bland-Altman analysis and the mean absolute difference (MAD) between both results was calculated. Clinical equivalence between measured and predicted VO₂peak was defined a priori: limits of agreements (upper and lower bounds) < 1.5 mL/kg/minute, corresponding to less than 10% of risk of bias.

The ability of the 3CRT to predict a cutoff VO₂peak ≥ 15 mL/kg/minute was evaluated by sensitivity, specificity and receiver operating characteristic (ROC) curve analyses. A P-value <0.05 was considered statistically significant in all cases. GraphPad Prism 5.03 and R 3.6.1 software were used for all analyses.

**Results**

**Patients**

Between March 2018 and February 2019, 51 patients were referred for CPX as part of a preoperative evaluation for NSCLC. Of these, 13 did not undergo the 3CRT because of orthopedic limitations associated with disabling pain. A further two CPXs were uninterpretable due to submaximal effort. The flow chart of these results is presented in Fig. 2. Patient characteristics for the 36

![Flow-chart of patients included in the study](image-url)

*Figure 2* Flow-chart of patients included in the study; NSCLC, non-small cell lung cancer; CPX, cardiopulmonary exercise test; 3CRT, 3-minute chair rise test.
The CPX and 3CRT results are summarized in Table 2. The correlation between VO2peak and 3CRT is significant (r² = 0.37, P < 0.01) as well as the vertical distance reached during the 3CRT (r² = 0.62, P < 0.01) (Fig 3). The other variables that were significantly correlated with VO2peak were: age (r² = 0.29, P < 0.01), BMI (r² = 0.18, P = 0.01) and FEV1 (r² = 0.12, P = 0.04).

### Capacity of the 3CRT to predict VO2peak

The best model determined by the multivariate analysis to predict VO2peak was: VO2peak predicted = (0.04765 × FEV1 in % predicted) − (0.20759 × BMI in kg/m2) − (0.11589 × age in years) + (0.38609 × vertical distance in m) + 16.62869; r² = 0.75, P < 0.01. The mean VO2peak measured during CPX and predicted by the previous equation was not significantly different 14.6 ± 3.6 vs. 14.6 ± 3.1, P = 0.99 (Table 2).

The bias between the VO2peak predicted using this equation and the measured VO2peak was 0.0 ± 1.7 mL/kg/minute, but the 95% limits of agreement were −3.5 to 3.5. The corresponding Bland-Altman (B&A) plots are presented in Fig 4. The mean absolute difference (MAD) was 1.5 ± 1.0 mL/kg/minute. These results indicate that even if the r-square of the predictive model was high, it did not provide an adequate fit for our data.

### Capacity of the 3CRT to predict a VO2peak > 15 mL/kg/minute

During CPX, there were 16 patients with a VO2peak ≥ 15 mL/kg/minute and 20 who did not reach this capacity. The ROC curve analysis is presented in Fig 5 and detailed in Table S2.
The area under the curve (AUC: 95% CI) was 0.82 (0.67 to 0.96). A number of chair rises ≥49 predicted a $\text{VO}_2\text{peak} \geq 15 \text{ mL/kg/minute}$ with a sensitivity of 81% and a false-positive rate of 25% (specificity: 75%). The false-positive rate reduced to 15%, 10% and 5% for patients who achieved 54, 62 and 65 rises, respectively.

In this cohort, the threshold of 49 chair rises was associated with a positive predictive value (PPV) and a negative predictive value (NPV) to detect a $\text{VO}_2\text{peak} \geq 15 \text{ mL/kg/minute}$ of 72% and 83%, respectively.

### Discussion

The incidence of lung cancer is increasing worldwide.\(^{17}\) Access to curative surgical treatment for patients with NSCLC requires an assessment of functional capacity in order to identify patients at risk.\(^{7, 12}\) Since $\text{VO}_2\text{peak}$ is a powerful predictive marker in many chronic pathologies, the demand for CPX is exploding in many institutions.

The need to develop field tests to estimate oxygen uptake is particularly topical. This study demonstrated a significantly positive relationship between performance on the 3CRT and $\text{VO}_2\text{peak}$ measured during CPX in patients with NSCLC. The vertical distance reached during 3CRT was particularly strongly correlated with $\text{VO}_2\text{peak}$ results. According to the multivariate analysis, the four independent variables related to $\text{VO}_2\text{peak}$ during the 3CRT were the vertical distance reached, age, BMI and FEV1. Although the equation derived from the multivariate regression did not accurately predict specific $\text{VO}_2\text{peak}$, the number of 49 or more rises performed during the 3CRT predicted a $\text{VO}_2\text{peak} \geq 15 \text{ mL/kg/minute}$ with a sensitivity of 81% and a specificity of 75%.

To our knowledge, this is the first study to assess the ability of the 3CRT method as a tool to determine the need for presurgical CPX in patients with NSCLC. The 3CRT is one of a set of many sit-to-stand-tests (STST). Although they all evaluate the ability to get up from a chair without using the upper limbs as a measure of physical capacity, the specific aims and methods of each differ. The 1-minute STST (1-STST) particularly is used in patients with COPD. It is well correlated to quadriceps strength,\(^{18}\) as well as distance walked during the 6-minute walk test (6MWT), thus providing an estimate of functional capacity with less hemodynamic stress.\(^{19, 20}\) However, such quick tests do not reflect cardiorespiratory capacity, especially as $\text{VO}_2$ kinetics are known to be slowed and related to the severity of the respiratory disease.\(^{21, 22}\) For patients with NSCLC, a longer evaluation over 3 minutes seems more appropriate since respiratory function is often altered and it may take 3 minutes for cardiorespiratory function to adapt to a constant load test.

We believe this test is valuable since it can easily be performed during a consultation to provide data to guide further therapeutic decisions. Similarly, to tests such as the SWT, the SCT and CPX, the main contraindication to its use is disabling lower limb pain. Patients with peripheral arteriopathy and hip prostheses were able to perform the 3CRT in the present study.
The lack of accuracy of the prediction of specific VO$_{2peak}$ value by the multivariate regression equation prohibits its use in clinical practice to predict VO$_{2peak}$. The fact HR$_{max}$ and Borg rating of dyspnea were significantly higher during the CPX suggests that the 3CRT was physiologically less stressful, or the evaluators did not motivate the patient as much as during the CPX during this new test. It is also possible that the sample size was too small and thus the dispersion of the data was relatively high. Larger studies should therefore be conducted to refine the equation and assess the influence of the evaluation conditions.

The ROC analysis showed that performance of 49 rises or more during the 3CRT was predictive of a VO$_{2peak}$ $\geq$ 15 mL/kg/minute, with a high level of sensitivity (81%) and a moderate false-positive rate (25%) (specificity: 75%). To compare with other field tests used in screening NSCLC patients in the preoperative assessment, the PPV and NPV for the threshold of 49 rises during the 3CRT were calculated (72% and 83%, respectively). A study of the SWT found higher PPV and NPV (90% each) for a distance $\geq$250 meters to predict a VO$_{2peak}$ $>$ 15 mL/kg/minute. For the SCT, a high PPV (86%) was found for a climb $\geq$22 meters to predict a VO$_{2peak}$ $>$ 15 mL/kg/minute; however, the NPV was only 62%. The SWT is therefore the most accurate of these tests in determining which patients need to be referred for CPX. However, it is also the most difficult to carry out in everyday practice, as none of the thoracic surgeons interviewed in a recent American survey prescribe a SWT. Comparatively to the SWT and SCT, the 3CRT can be performed in nearly any care setting, including very limited spaces such as consultation rooms by surgeons themselves, or allied health professionals, and could be sufficient to determine the need for presurgical CPX in patients with NSCLC.

During the 3CRT validation study in COPD patients, Aguilaniu et al. reported that patients who were able to perform more than 50 rises during the 3 minutes had a lower score on the disability related to COPD tool (DIRECT) questionnaire, suggesting less impaired function. Similarly, Lévesque et al. reported a mean number of 51.9 ± 13.1 rises during the 3CRT in patients with COPD whose BODE index score was in the first favorable quartile (0–2 /10). These results are consistent with the threshold of 49 rises found in the present study to predict a VO$_{2peak}$ $\geq$ 15 mL/kg/minute in the present cohort.

This study had several limitations. First, there may have been selection bias due to the retrospective design. Second, even if the multivariate regression analysis model (r$_{2}$ = 0.75, P < 0.01) and ROC curve analyses (AUC 0.82 [0.67 to 0.96]) were statistically significant, the study was not powered a priori and was probably underpowered. The measurement of post hoc statistical power is questionable and has not been recommended by several authors. We took the option not to include this measure in the present paper, as we felt that it would not strengthen the level of evidence of this preliminary retrospective study. Third, a separate validation with an independent cohort of patients is necessary to confirm the PPV and NPV for the threshold of 49 rises during the 3CRT to predict a VO$_{2peak}$ $\geq$ 15 mL/kg/minute. Finally, this study only focused on the evaluation of performance on the 3CRT in relation to the VO$_{2peak}$ as a postoperative risk factor, and did not directly evaluate postoperative complications.

Nevertheless, the results of this preliminary, retrospective study suggest that this simple tool is promising as a screening test prior to CPX in patients with NSCLC and thus larger, prospective studies are justified. Moreover, the data relating to the relationship between 3CRT performance and VO$_{2peak}$ can be used to determine the appropriate design of a prospective study with a larger number of patients.

In conclusion, performance of at least 49 chair rises during the 3CRT indicated a VO$_{2peak}$ $\geq$ 15 mL/kg/minute, similar to results from the SCT and the SWT. However, prediction of the specific VO$_{2peak}$ value based on the 3CRT performance was insufficiently accurate or sensitive to be used as a standalone method for determining the specific

![Figure 5](image-url)
VO₂peak value. The 3CRT is a simple field test that could be very useful as a screening test to indicate patients with NSCLC who require further testing with CPX and those who can be safely referred for surgery. Prospective studies with larger sample sizes are needed to validate these results.

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Disclosure

The authors report having no conflict of interest to disclose in relation to the present study.

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Supporting Information

Additional Supporting Information may be found in the online version of this article at the publisher’s website:

Table S1: Comorbidities and treatments \((n = 36)\). Table S2: Receiver operating characteristic analysis of the number of chair rises versus peak oxygen consumption \((\text{VO}_{2\text{peak}}) \geq 15 \text{ mL/kg/minute}; \text{binary outcome})\.

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