

Accuracy of forced oscillation technique to assess lung function in geriatric COPD population

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Introduction: Performing lung function test in geriatric patients has never been an easy task. With well-established evidence indicating impaired small airway function and air trapping in patients with geriatric COPD, utilizing forced oscillation technique (FOT) as a supplementary tool may aid in the assessment of lung function in this population.

Aims: To study the use of FOT in the assessment of airflow limitation and air trapping in geriatric COPD patients.

Study design: A cross-sectional study in a public hospital in Hong Kong. [ClinicalTrials.gov](https://clinicaltrials.gov/ct2/show/study/NCT01553812) ID: NCT01553812.

Methods: Geriatric patients who had spirometry-diagnosed COPD were recruited, with both FOT and plethysmography performed. “Resistance” and “reactance” FOT parameters were compared to plethysmography for the assessment of air trapping and airflow limitation.

Results: In total, 158 COPD subjects with a mean age of 71.9±0.7 years and percentage of forced expiratory volume in 1 second of 53.4±1.7 L were recruited. FOT values had a good correlation ($r=0.4-0.7$) to spirometric data. In general, X values (reactance) were better than R values (resistance), showing a higher correlation with spirometric data in airflow limitation ($r=0.07-0.49$ vs $0.61-0.67$), small airway ($r=0.05-0.48$ vs $0.56-0.65$), and lung volume ($r=0.12-0.29$ vs $0.43-0.49$). In addition, resonance frequency (Fres) and frequency dependence (FDep) could well identify the severe type (percentage of forced expiratory volume in 1 second <50%) of COPD with high sensitivity (0.76, 0.71) and specificity (0.72, 0.64) (area under the curve: 0.8 and 0.77, respectively). Moreover, X values could stratify different severities of air trapping, while R values could not.

Conclusion: FOT may act as a simple and accurate tool in the assessment of severity of airflow limitation, small and central airway function, and air trapping in patients with geriatric COPD who have difficulties performing conventional lung function test. Moreover, reactance parameters were better than resistance parameters in correlation with air trapping.

Keywords: airflow limitation, air trapping, chronic obstructive pulmonary disease

Introduction

COPD is a common and morbid disease worldwide, affecting a significant proportion of geriatric patients.¹ Though spirometry is a mandatory tool for diagnosis and assessing severity in patients with COPD, it is underused in geriatric patients.^{2,3} The higher prevalence of cognitive and functional impairment in elderly patients has rendered many of them incapable of performing a satisfactory forced expiratory maneuverer in spirometry.⁴⁻⁶ In fact, ~18% of advanced-age patients with COPD failed to perform adequate spirometry,⁵ and the quality of spirometric performance is also jeopardized in elderly patients with COPD.⁴

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Forced oscillation technique (FOT) may be an alternative tool to assess lung function in geriatric patients. FOT is less time-consuming and technically easier to perform, as it is measured when patients effortlessly breathe-in their tidal volume, requiring minimal patient cooperation. In fact, current evidences suggest that FOT parameters correlated well with forced expiratory volume in 1 second (FEV_1) and could reflect airway resistance accurately.^{7–10} It was also suggested to be a more sensitive marker for detection of airway hypersensitivity^{11–14} as well as early airway disease.^{15,16}

Despite these benefits, FOT was not widely applied in the diagnosis and monitoring of patients with geriatric COPD. Besides, the use of FOT in the assessment of air trapping in “geriatric COPD patients” was not well investigated in earlier studies, although close relationship exists between air trapping and patients’ symptomatology.¹⁷ Furthermore, previous studies did not explicitly demonstrate the use of FOT in identifying the “severe type” ($\%FEV_1 < 50\%$) of elderly patients with COPD, which is important since the current Global Initiative for Chronic Obstructive Lung Disease (GOLD) guideline recommends combined assessment approach to manage patients with COPD with $\%FEV_1 < 50\%$ being regarded as a “high-risk” group.

Therefore, our study aims to investigate the use of various FOT parameters (both reactance and resistance) in assessment of airflow limitation, small airway disease, and air trapping in patients with geriatric COPD. We also aim to investigate the accuracy of FOT in identifying the “severe type” of elderly patients with COPD ($\%FEV_1 < 50\%$), who are regarded as belonging to a high-risk group according to the GOLD guideline.

Aims

The study aims to investigate the use of FOT in the assessment of different degrees of air trapping as well as airflow limitation in patients with geriatric COPD.

Methods

Study design and population

This is a cross-sectional study conducted in a public hospital in Hong Kong (Kwong Wah Hospital, Hong Kong, People’s Republic of China). The study population included stable COPD subjects who attended the specialized COPD clinic or the COPD ambulatory rehabilitation clinic from January 1, 2011 to December 31, 2014, in Kwong Wah Hospital.

Geriatric COPD subjects aged between 60 and 85 years with a clinical history compatible with COPD and the spirometry showing obstructive pattern ($FEV_1/\text{forced vital capacity [FVC]} < 0.7$) were recruited. Subjects were excluded from the study

if they had coexisting pulmonary diseases such as asthma, interstitial lung diseases, and lung malignancy or if they were uncooperative or too dyspneic for lung function test.

Patients’ demographic data including age, sex, smoking status, comorbidities, and dyspnea level (Modified Medical Research Council Dyspnea Scale) were recorded. Both FOT and plethysmography were performed, and the results from both methods were compared for assessment of airflow limitation and air trapping.

Lung function tests

Both lung function tests (FOT and plethysmography) were performed by trained respiratory specialist nurses in the same visit. Since FOT would not affect the bronchial tone, it was performed before plethysmography for every patient. The details of the lung function tests are as follows.

FOT

FOT was performed using the instrument i2M (Chess Medical, Ghent, Belgium) with the aid of nose clip and mouthpiece that stabilized patients’ tongue position. It measured the respiratory mechanics (“resistance” and “reactance”) by producing a multifrequency (4–48 Hz) pseudorandom noise impulse. The test was done with the patient in sitting position and the cheek supported by both hands to reduce airway shunting. The machine was calibrated before using for each COPD subject, according to the instructions of the machine manufacturer. The test was done based on the European Respiratory Society Task Force recommendations on forced oscillation testing.¹⁸ Moreover, only measured values with a minimal coherence function of 95% were considered valid. Three valid measurements were taken, and the final result of the test was calculated as the mean of these three measurements.

The resistance and reactance components of the respiratory system over various frequencies (4–48 Hz) were obtained at every 2 Hz and were expressed as two distinct curves. The FOT parameters that were included for analysis included resistance (R) and reactance (X) at various oscillation frequencies (4–48 Hz), frequency resonance (F_{res} – the frequency at which reactance equals to zero), frequency dependence (FDep – the slope of resistance plotted against frequency), average reactance (X_{Avr}), and average resistance (R_{Avr}). The predicted values of FOT parameters were calculated and provided by the manufacturer according to the European Respiratory Society recommendation.¹⁸

Plethysmograph

This was performed by the machine MedGraphics Elite series™ Plethysmograph (Medical Graphics Corporation,

St Paul, MN, USA) with acceptable repeatability and in concordance with the American Thoracic Society standard for lung function test.¹⁹ Spirometry data (FEV₁, FVC, FEV₁/FVC ratio, %FEV₁, forced expiratory flow between 25% and 75% of forced vital capacity [FEF_{25-75%}]), lung volume (inspiratory capacity [IC], vital capacity, total lung capacity [TLC], residual volume [RV], RV/TLC ratio), as well as airway resistance (Raw) and conductance (Gaw) were measured. The Chinese reference values for spirometric parameters were used for analysis.²⁰

Statistical analysis

For continuous parametric data, values were expressed as mean ± standard error of mean unless stated otherwise. Comparison between two continuous variables was done by correlation test and expressed as Spearman correlation coefficient (*r*). Analysis of variance test was used to compare continuous variables among three or more groups with post hoc analysis estimated by Bonferroni test. Receiver operating characteristic curve was plotted with Jouden Index being used to determine the cutoff points of various FOT parameters in predicting severe type of COPD (%FEV₁ <50%). All tests were two sided and regarded as significant if *P*-value was <0.05. Predictive Analytics Software 18.0 (IBM Corp., Armonk, NY, USA) was used for statistical analysis.

Ethics

Written informed consent was obtained from every eligible subject. The study was approved by the Hong Kong Hospital Authority Kowloon West Cluster Research Ethic Committee (reference: KW/EX-12-030 [48-15]).

Results

Among the 177 eligible COPD subjects, 19 patients were excluded (five patients refused to participate, 14 patients defaulted the lung function tests). Finally, 158 COPD subjects, with a mean age of 71.9±8.3 years and %FEV₁ of 53.4±21.4, completed the study. Demographic and lung function data of the subjects are shown in Tables 1 and 2, respectively.

Assessment of airflow limitation/COPD severity

Reactance (*X*) was superior to resistance (*R*) parameters in the assessment of airflow limitation, with a higher correlation to FEV₁ (Figure 1), FEF_{25-75%} (Figure 2) and RV/TLC ratio (Figure 3). Regarding the COPD severity, all *X* parameters (Fres, XAvr, X6Hz, and X20Hz) had a good correlation with GOLD stages, while the correlation to GOLD stages was much diverse among *R* parameters (Figure 4). Among the *R*

Table 1 Demographic data of the COPD subjects

Demographics, N=158	Mean ± SEM
Sex	Male: 91.1%; female: 8.9%
Age (years)	71.9±0.66
Body weight (kg)	56.8±0.91
Height (cm)	160.3±0.54
BMI (kg/m ²)	22.0±0.31
Smoking status	Active smoker: 18.9%; ex-smoker: 65.7%; nonsmoker: 15.4%
Alcohol	Chronic drinker: 6.3%; social drinker: 17.9%; nondrinker: 56.3%; ex-drinker: 19.6%
Education status	Primary school: 70.4%; secondary school: 26.8%; tertiary school: 2.8%
Working status	Unemployed: 6.3%; retired: 82.4%; working: 10.6%
MMRC	Grade 0: 1.5%; Grade 1: 12.5%; Grade 2: 46.3%; Grade 3: 38.2%; Grade 4: 1.5%
GOLD stages	Stage 1: 28/158 (17.7%) Stage 2: 53/158 (33.5%) Stage 3: 66/158 (41.8%) Stage 4: 11/158 (7%)

Abbreviations: BMI, body mass index; GOLD, Global Initiative for Chronic Obstructive Lung Disease; SEM, standard error of mean; MMRC, Modified Medical Research Council Dyspnea Scale.

Table 2 Lung function parameters of the COPD subjects

Lung function parameters	Mean ± SEM
Plethysmography data	
FEV ₁ (L)	1.12±0.04
%FEV ₁	53.4±1.72
FVC (L)	2.29±0.05
%FVC	80.3±1.49
FEV ₁ /FVC ratio	48.4±1.06
IC (L)	1.52±0.04
%IC	56.8±1.5
RV (L)	3.6±0.16
%RV	169.3±7.48
TLC (L)	6.02±0.16
%TLC	110±3.61
Raw (kPa·L·S ⁻¹)	7.82±0.64
%Raw	529.9±44.4
Gaw (L·s·kPa ⁻¹)	0.26±0.02
%Gaw	25.5±1.78
RV/TLC	0.57±0.01
FOT data	
Reactance (X)	
Fres	26.4±0.71
X6	-2.05±0.12
X20	-0.59±0.06
XAvr	-1.10±0.08
Resistance (R)	
FDep	-0.07±0.005
R6	3.69±0.11
R20	2.61±0.08
RAvr	2.98±0.08

Abbreviations: FDep, frequency dependence; FEV₁, forced expiratory volume in 1 second; Fres, resonance frequency; FVC, forced vital capacity; Gaw, conductance; IC, inspiratory capacity; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAvr, average resistance; Raw, airway resistance; RV, residual volume; TLC, total lung capacity; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

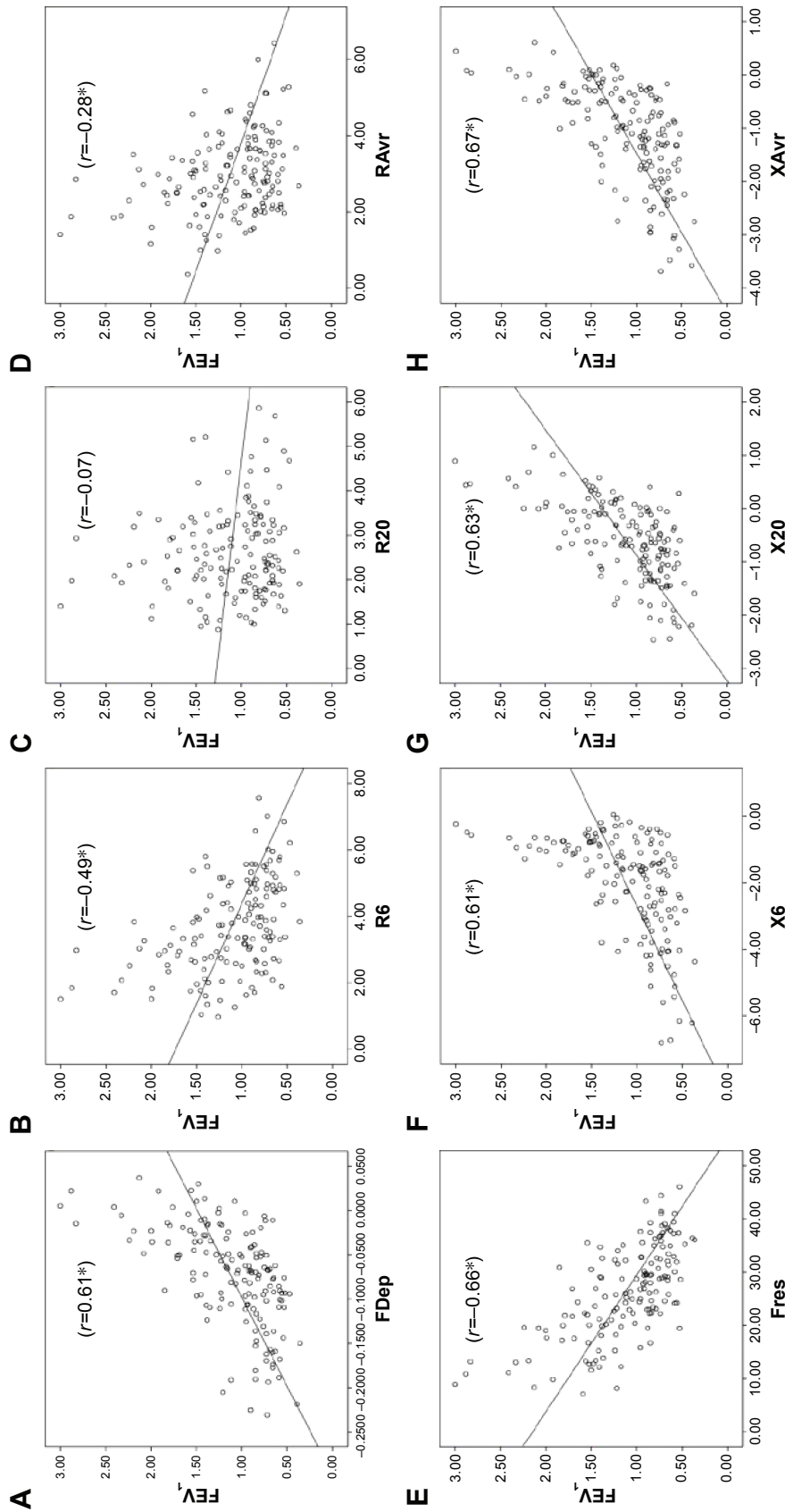


Figure 1 Correlation between FEV₁ and various FOT parameters.

Notes: (A) FDep, (B) R6, (C) R20, (D) RAVr, (E) X6, (F) X20, and (H) XAvr. * $p < 0.05$.

Abbreviations: FDep, frequency dependence; FEV₁, forced expiratory volume in 1 second; FOT, forced oscillation technique; Fres, resonance frequency; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAVr, average resistance; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

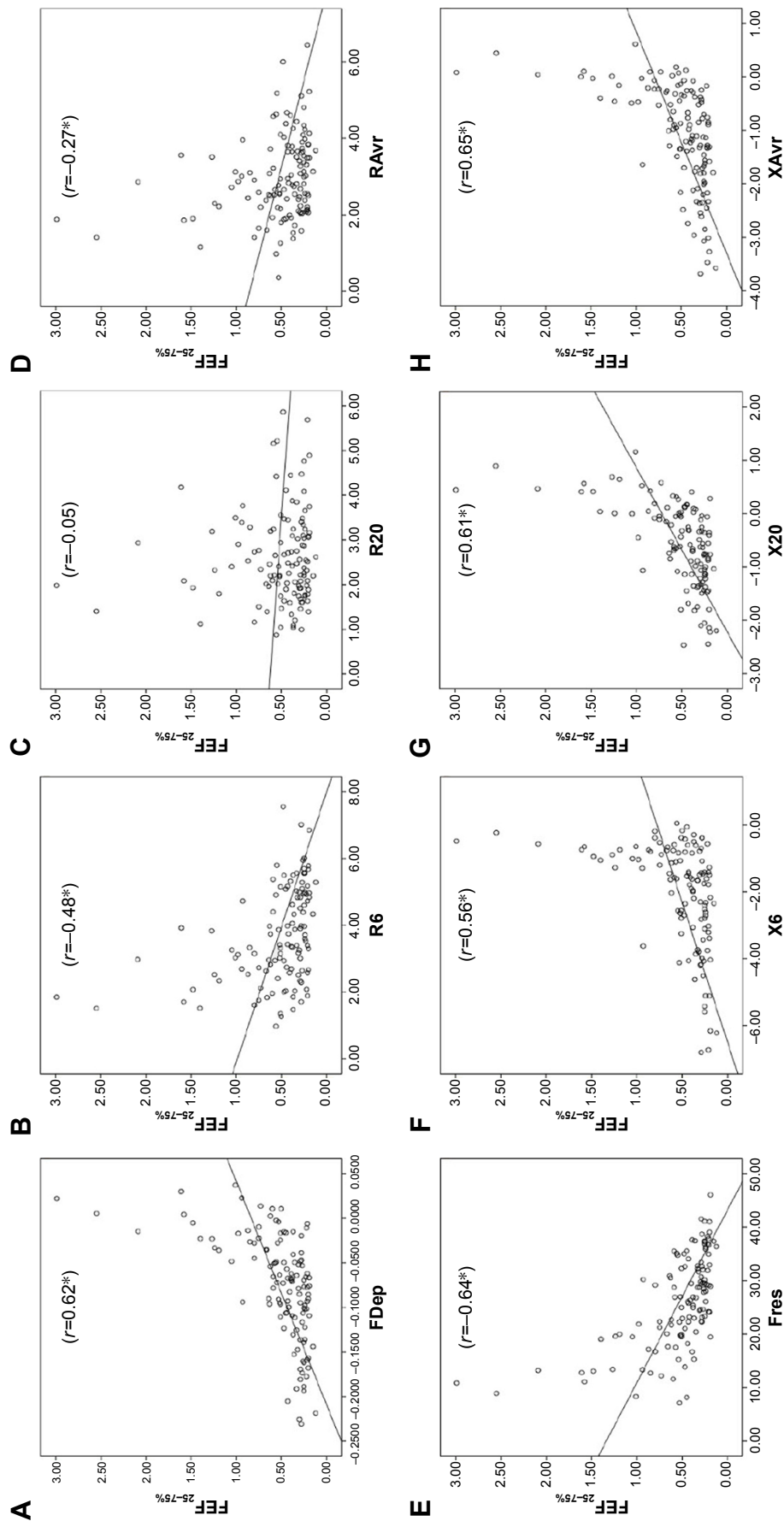


Figure 2 Correlation between $FEF_{25-75\%}$ and various FOT parameters.
Notes: (A) FDep, (B) R6, (C) R20, (D) RAVr, (E) Fres, (F) X6, (G) X20, and (H) XAVr. * $P < 0.05$.
Abbreviations: FDep, frequency dependence; FOT, forced oscillation technique; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAVr, average resistance; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAVr, average reactance.

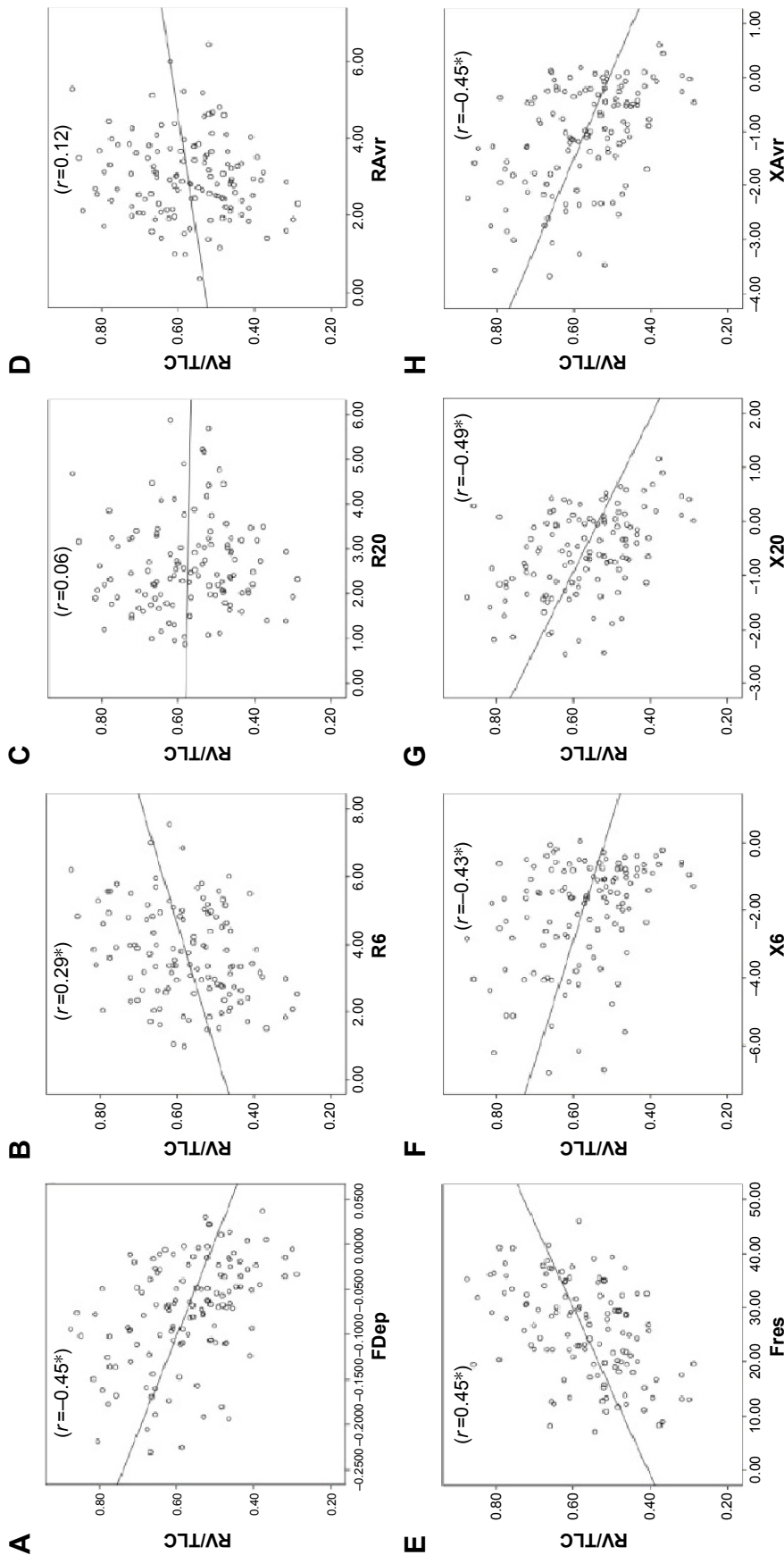


Figure 3 Correlation between RV/TLC ratio and various FOT parameters.

Notes: (A) FDep, (B) R6, (C) R20, (D) RAVr, (E) Fres, (F) X6, (G) X20, and (H) XAVr. * $P < 0.05$.

Abbreviations: FDep, frequency dependence; FOT, forced oscillation technique; Fres, resonance frequency; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAVr, average resistance; RV, residual volume; TLC, total lung capacity; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAVr, average reactance.

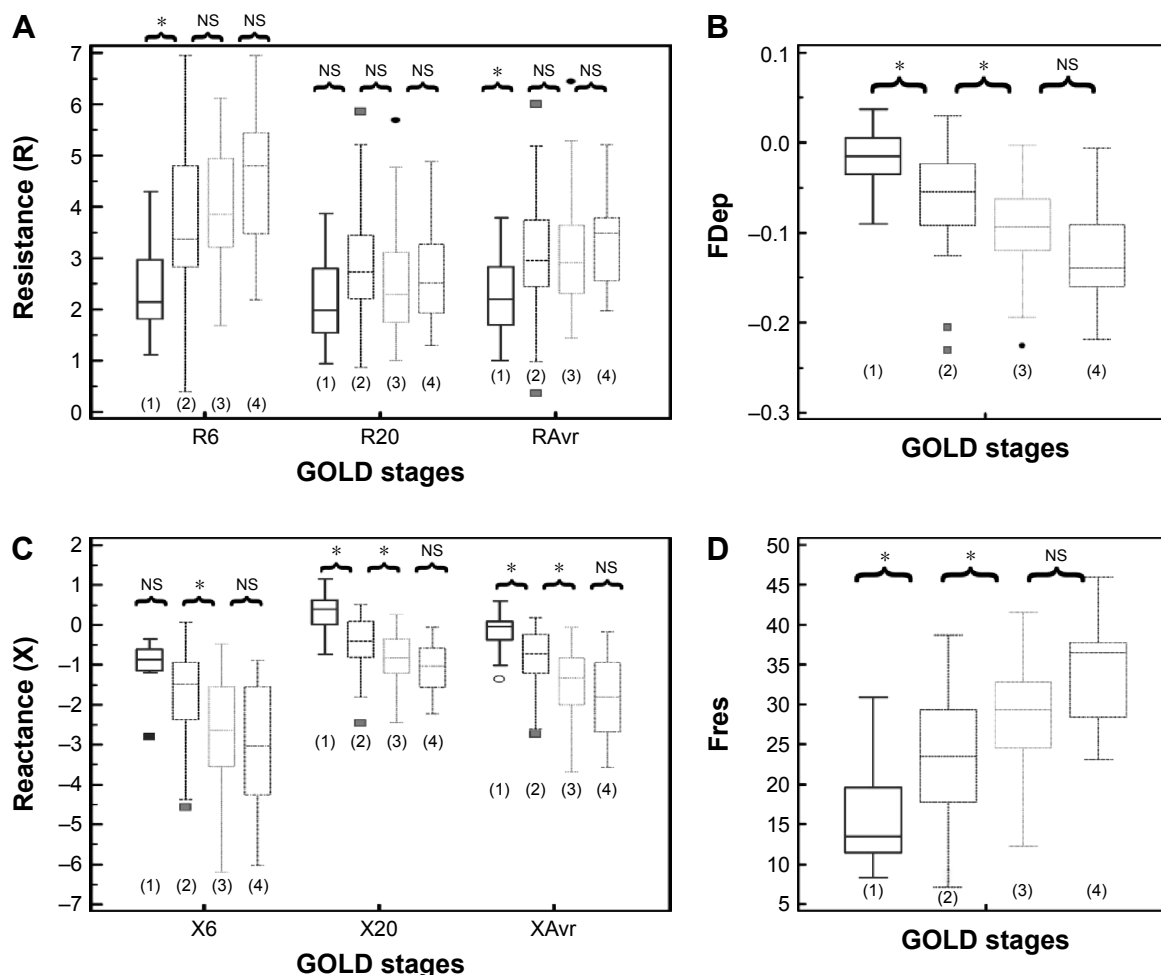


Figure 4 Different “resistance” and “reactance” parameters in various GOLD stages (stages 1–4). **Notes:** (A) Resistance (R6, R20, RAvr); (B) FDep; (C) reactance (X6, X20, XAvr); and (D) Fres. * $P < 0.05$. Outliers are displayed as various symbols to represent the different groups. **Abbreviations:** FDep, frequency dependence; Fres, resonance frequency; GOLD, Global Initiative for Chronic Obstructive Lung Disease; NS, not significant; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAvr, average resistance; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

parameters, FDep correlated well with GOLD stages, while R20 and RAvr correlated poorly (Figure 4).

Predicting severe type of patients with COPD ($FEV_1 < 50\%$)

Reactance values (X), in general, had a high sensitivity (70%–76%) and specificity (64%–72%) in identifying severe type of COPD, among which Fres showed the highest area under the curve (AUC: 0.80). Of the resistance parameters, FDep was good in identifying severe patients with COPD (AUC: 0.77), while isolated R values (R6, R20, and RAvr) were not (AUC: 0.4–0.6) (Table 3; Figure 5).

Assessment of air trapping

Since there is no standardized stratification system for the severity of air trapping in patients with COPD, air trapping

Table 3 The cutoff points of various FOT parameters (reactance and resistance) in diagnosing severe type of patients with COPD ($\%FEV_1 < 50\%$)

FOT parameters	AUC	Cutoff points	Sensitivity	Specificity
Resonance (X)				
Fres	0.80	25.8	0.76	0.72
X6	0.73	-1.61	0.70	0.65
X20	0.77	-0.62	0.72	0.64
XAvr	0.79	-1.0	0.74	0.70
Resistance (R)				
FDep	0.77	-0.070	0.71	0.64
R6	0.66	3.35	0.69	0.6
R20	0.45	2.42	0.46	0.47
RAvr	0.57	2.88	0.54	0.53

Abbreviations: AUC, area under the curve; FDep, frequency dependence; FEV_1 , forced expiratory volume in 1 second; FOT, forced oscillation technique; Fres, resonance frequency; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAvr, average resistance; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

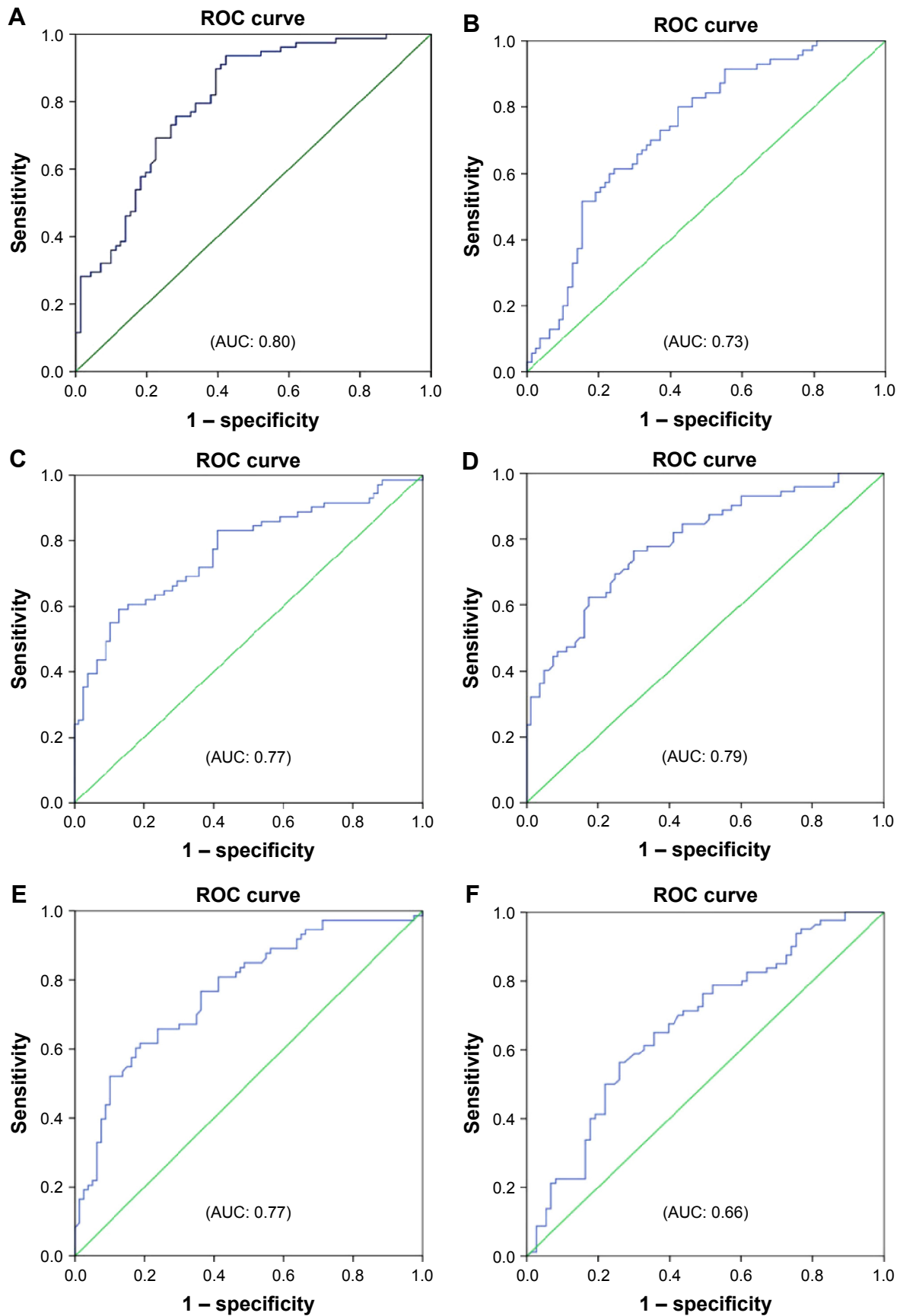


Figure 5 (Continued)

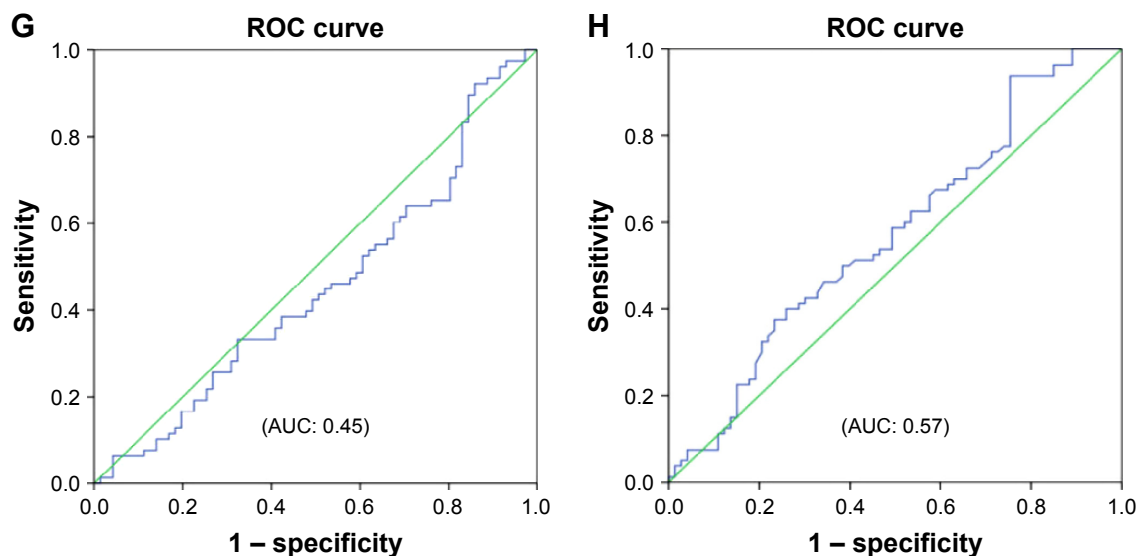


Figure 5 ROC curves for various FOT parameters in prediction of severe type of patients with COPD ($\%FEV_1 < 50\%$).

Notes: (A) Fres, (B) X6, (C) X20, (D) XAvr, (E) FDep, (F) R6, (G) R20, and (H) RAvr.

Abbreviations: AUC, area under the curve; FDep, frequency dependence; FEV₁, forced expiratory volume in 1 second; FOT, forced oscillation technique; Fres, resonance frequency; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAvr, average resistance; ROC, receiver operating characteristic; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

was arbitrarily divided into three different degrees according to the RV/TLC ratio in this study. The cutoff points were chosen to be close to the 25th and 50th percentile of RV/TLC ratio. Group 1 (mild air trapping) consisted of patients with RV/TLC ratio < 0.45 , group 2 (moderate air trapping) patients had the ratio between 0.45 and 0.6, and group 3 (severe air trapping) patients had RV/TLC ratio > 0.6 .

The correlation of FOT parameters to various degrees of air trapping is depicted in Figure 6. Reactance was better than resistance parameters in stratifying various degrees of air trapping. There was a significant difference of reactance values (XAvr, Fres, and X20Hz) among the three different severities of air trapping ($P < 0.05$ between each group), while isolated resistance FOT parameters (RAvr and R6Hz) did not differ ($P > 0.05$) (Figure 6). Moreover, reactance (X) parameters showed a stronger correlation with other lung volume parameters (IC, FVC, RV, RV/TLC ratio) when compared to the isolated R parameters (r around 0.45 vs $r < 0.3$) (Figure 3; Table S1).

Discussion

Assessment of the severity of airflow limitation and central airway function

Previous studies have shown a rather good correlation between FOT and spirometric parameters.^{7,8,21–29} Moreover, our study has further highlighted that such good correlation could be maintained even at an advanced age among

geriatric group patients (mean age of our study population was 72 years), who are more vulnerable to cognitive and functional impairment that results in difficulties and underuse of conventional lung function test.

Furthermore, we also demonstrated that FOT might have a role in the assessment of the severity of airflow limitation in geriatric patients, especially in those elderly patients who have difficulty in performing conventional spirometry. In particular, FEV₁, generally regarded as a marker to reflect central airway function, was shown to be highly correlated to both X values and FDep ($r = 0.61–0.67$). Furthermore, the role of X values and FDep in the assessment of severity of airflow obstruction was further highlighted by their strong correlation with various GOLD stages in the current study. These results are compatible with previous findings and are of paramount importance since current COPD guideline recommends combined assessment approach in the management of patients with COPD, in which patients with COPD with severe airflow limitation ($\%FEV_1 < 50\%$) are regarded as a high-risk group. Our study showed not only the superiority of X to R in the classification of various GOLD stages but also its high sensitivity and specificity in identifying the high-risk patients. In particular, X values, in general, have a high sensitivity (70%–76%) and specificity (64%–72%) in identifying the severe type of COPD ($\%FEV_1 < 50\%$), among which Fres showed the highest AUC (0.80). In contrast, R values were insensitive to identify severe type of COPD (AUC: 0.4–0.6). Likewise,

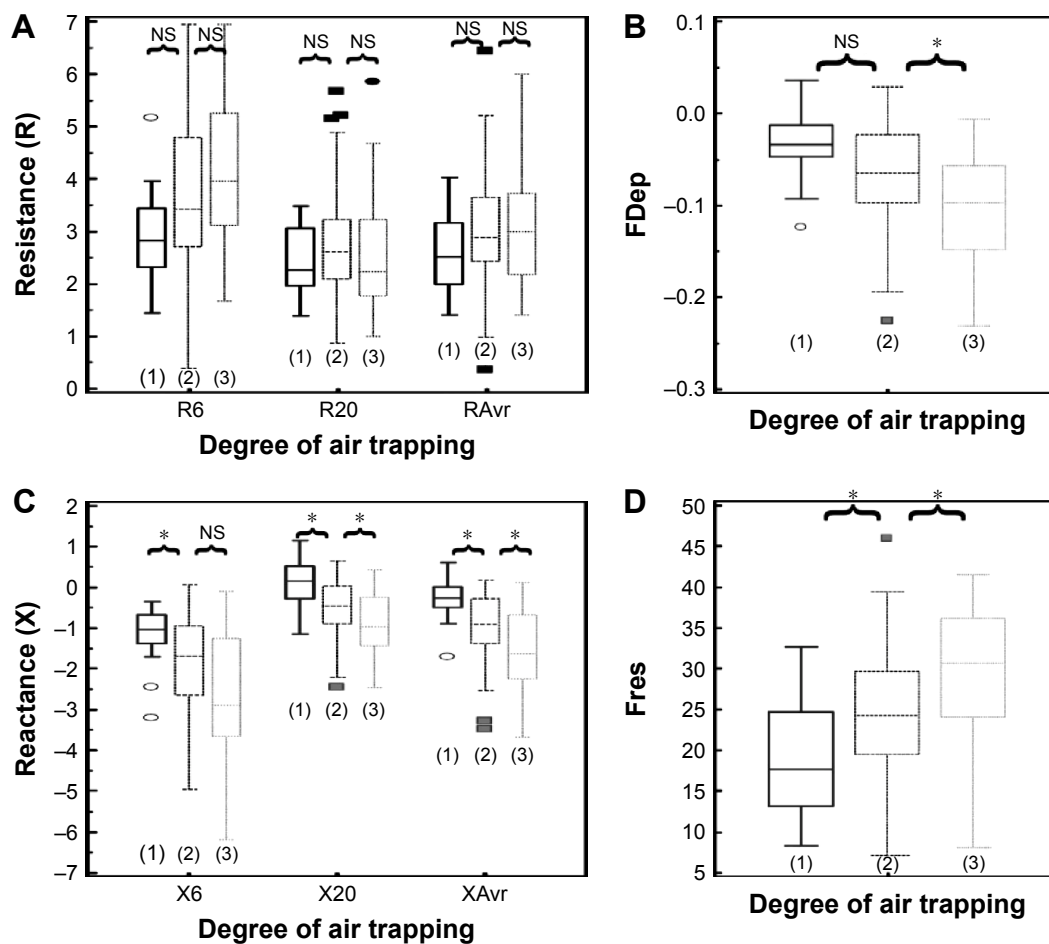


Figure 6 Different “resistance” and “reactance” parameters in various degrees of air trapping (grades 1–3).

Notes: (A) Resistance (R6, R20, RAvr); (B) FDep; (C) reactance (X6, X20, XAvr); and (D) Fres. * $P < 0.05$. Outliers are displayed as various symbols to represent the different groups.

Abbreviations: FDep, frequency dependence; Fres, resonance frequency; NS, not significant; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAvr, average resistance; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

previous studies^{26,30,31} had suggested that isolated R values like RAvr were a poor parameter to assess COPD severity, while X values (especially Fres) as well as FDep were better markers.³² In addition, the various FOT cutoff values for severe airflow limitation ($\%FEV_1 < 50\%$) were also determined in our study.

Assessment of small airway function

Small airway measurement is challenging and it was expressed in terms of $FEF_{25-75\%}$ in our study. A high correlation between X and $FEF_{25-75\%}$ signified that X is valuable in small airway assessment. X is the imaginary part of respiratory impedance (Zrs), comprising the effects of inertia (I) and compliance (C). Compliance (C), which represents the ability of the respiratory system to store energy, is primarily located in the lung periphery. Therefore, X at a low frequency can reflect tissue recoil in distal airway. Furthermore, within-breath analysis

had further suggested that respiratory phase difference in reactance (ΔXrs) could detect expiratory flow limitation (EFL), which is a characteristic feature in patients with moderate-to-severe COPD with dynamic hyperinflation.^{33–35}

However, R values, in general, had a lower correlation to $FEF_{25-75\%}$, when compared to X values. R20 assesses small airway function poorly because of its damping effect in distal airways. Moreover, our study suggested that FDep was the best indicator among R values for assessment of peripheral airway obstruction ($FEF_{25-75\%}$) ($r=0.48$ vs 0.62). FDep is the slope of resistance–frequency curve, it is very sensitive to detect small airway disease because a small rise in small airway resistance would significantly reduce the slope-gradient in FDep. In fact, the use of FDep has also been proposed by previous studies to be highly correlated with peripheral airway disease^{32,36} and detection of early airway disease.^{16,37} Furthermore, FDep had been shown to

be closely linked to the activity-related dyspnea in patients with moderate-to-severe COPD.³⁸

In fact, evidences suggested that FOT may detect subtle changes in distal airway function earlier than with conventional spirometry, even in the setting of normal spirometry.³⁹ Shi et al⁴⁰ suggested that small airway impulse oscillometry index was even better than FEV₁ and FEF_{25–75%} in the detection of small airway dysfunction. Besides, in the trial of Frantz et al,⁴¹ self-reported chronic bronchitis, emphysema, or COPD was associated with higher impulse oscillometry pulmonary resistance and lower pulmonary reactance, even in patients with normal spirometry. The high sensitivity of FOT in small airway function renders its value in clinical application to monitor disease progress and treatment response. It has already been utilized to monitor disease progress in patients with posttransplant bronchiolitis obliterans and treatment effect on small airway in patients with COPD.^{42,43}

Assessment of the degree of air trapping

Measurement of air trapping is difficult, although it contributes significantly to symptoms in patients with COPD.¹⁷ Conventional plethysmography is labor intensive and time-consuming, while a quantitative computed tomography (CT) scan is expensive with significant radiation risk.^{44,45} FOT, a convenient tool with good correlation with small airway function, may have a role in the assessment of air trapping. However, previous trials on the correlation between FOT and air trapping parameters showed conflicting results. For instance, the prospective study of Kolsum et al⁸ showed only a weak association between FOT parameters and lung volume parameters such as TLC and IC ($r < 0.25$). Likewise, the large cohort study of Crim et al⁴⁶ that utilized “percentage of low attenuation area” (%LAA) in CT thorax scan for assessing degree of air trapping/emphysema failed to demonstrate a significant direct relationship between FOT parameters (R5, R5–R20, reactance area) and %LAA ($r < 0.16$), despite the fact that both FOT impedance parameters and %LAA worsen as GOLD stage increases.

In contrary to these studies, our study did demonstrate a higher correlation between FOT parameters and IC (r around 0.4–0.46). We also demonstrated that X parameters (eg, Fres and XAvr) had a good correlation with RV/TLC ratio (r around 0.45), while isolated “resistive” parameters (RAvr and R6Hz) did not show the correlation ($r < 0.3$).

Our study is consistent with the study of Van Noord et al,⁴⁷ which showed the most prominent decrease in Xrs (resonance frequency) and frequency dependence in emphysema patients when compared to chronic bronchitis and asthma

patients. In a recent study, Mikamo et al⁴⁸ reported that a high EFL index ($0.55 \text{ kPa} \cdot \text{L}^{-1} \cdot \text{S}^{-1}$) in FOT was independently predicted by emphysema extent measured by high-resolution computed tomography as well as the degree of hyperinflation as expressed by functional residual capacity. These findings could be explained by the superiority of X in the assessment of small airway dysfunction that results in air trapping. Furthermore, X represents lung compliance, which is the ability of the respiratory system to store energy. Compliance is significantly reduced in emphysema or air trapping because of loss of lung elastic recoil. It was, therefore, proposed that the degree of emphysema or air trapping correlated to the FOT reactance (X) parameters.⁴⁹ Apart from this, respiratory phase difference in reactance (ΔX_5) could also detect EFL,^{33,34} as demonstrated in the within-breath analysis. This EFL and small airway obstruction would result in air trapping. We thus believe that reactance parameters may have a role in the assessment of severities of air trapping in patients with COPD.

Strengths and limitations

The present study suggests that FOT may be a simple and effective means to assess airflow limitation (both central and small airway) as well as air trapping in patients with geriatric COPD, even at an advanced age (mean age > 72 years), which may be important because this population has the most difficulties in performing conventional spirometry. Besides, we performed a comprehensive assessment of FOT in our study, showing that reactance parameters (X) are better than resistance parameters (R) in the assessment of the degree of airflow limitation, small and central airway function, as well as air trapping, which is significantly related to patients’ symptomatology. Furthermore, we also contributed to the management of patients with geriatric COPD by demonstrating FOT as a simple tool with high sensitivity and specificity in identifying the patients with high-risk COPD (%FEV₁ $< 50\%$) in the geriatric group.

However, there were a number of limitations in our study. First, the COPD subjects were predominantly Chinese male; while reflecting the local epidemiology of predominant male smokers, this affects the study’s generalizability.

Second, since there was no reference value for the FOT parameters in Chinese patients, it was difficult to compare the “adjusted” FOT values to the “adjusted” spirometric values such as %FEV₁. Regarding the assessment of air trapping, though RV/TLC ratio was correlated with hyperinflation of lung, it may not be the most accurate or stable parameter to determine the degree of air trapping. Other measurements such as quantitative CT scan may be a more robust method

for determining the degree of air trapping. Moreover, within-breath analysis of FOT parameter was not performed in our study, though previous studies had suggested that higher impedances were observed in the expiratory phase of patients with COPD. Nevertheless, our study did confirm the high correlation between FOT and spirometric values, and we also demonstrated its role in the assessment of small airway function and air trapping.

Further studies on the role of FOT in the assessment of air trapping as well as identifying different COPD phenotypes are needed, especially using within-breath analysis of different FOT parameters and more robust techniques, such as quantitative CT scan, to quantify the severity of air trapping.

Conclusion

FOT may act as a simple and accurate tool in the assessment of severity of airflow limitation, small and central airway function, and air trapping in patients with geriatric COPD, especially in those who have difficulties performing conventional lung function test. Moreover, reactance parameters were better than resistance parameters in correlation with air trapping as well as identifying high-risk patients (%FEV₁ <50%).

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Author contributions

All authors made substantial contributions to conception and design, acquisition of data, or analysis and interpretation of data; took part in drafting the article or revising it critically for important intellectual content; gave final approval of the version to be published; and agree to be accountable for all aspects of the work.

Disclosure

The authors report no conflicts of interest in this work.

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Supplementary material

Table S1 Correlation coefficient (*r*) between FOT parameters and conventional lung function test

	Resistive parameters				Reactive parameters			
	R6	R20	RAvr	FDep	X6	X20	XAvr	Fres
Airflow parameters								
FEV ₁	-0.49*	-0.07	-0.28*	0.61*	0.61*	0.63*	0.67*	-0.66*
Raw	0.41*	0.06	0.26*	-0.49*	-0.48*	-0.54*	-0.55*	0.57*
FEF _{25-75%}	-0.48*	-0.05	-0.27*	0.62*	0.56*	0.61*	0.65*	-0.64*
Lung volume parameters								
FVC	-0.46*	-0.21*	-0.36*	0.43*	0.47*	0.48*	0.52*	-0.47*
IC	-0.40*	-0.22*	-0.32*	0.32*	0.44*	0.44*	0.46*	-0.44*
RV	0.07	0.24*	-0.10	-0.31*	-0.24*	-0.31*	-0.26*	0.28*
TLC	-0.11	-0.31*	-0.24*	-0.11	-0.05	-0.08	-0.05	0.06
RV/TLC	0.29*	0.06	0.12	-0.45*	-0.43*	-0.49*	-0.45*	0.45*

Notes: **P*<0.05. The figures illustrate Spearman correlation coefficient (*r*).

Abbreviations: FDep, frequency dependence; FEV₁, forced expiratory volume in 1 second; FOT, forced oscillation technique; Fres, resonance frequency; FVC, forced vital capacity; IC, inspiratory capacity; R6, resistance at 6 Hz; R20, resistance at 20 Hz; RAvr, average resistance; Raw, airway resistance; RV, residual volume; TLC, total lung capacity; X6, reactance at 6 Hz; X20, reactance at 20 Hz; XAvr, average reactance.

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