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Longitudinal spirometry among patients in a treatment program for community members with World Trade Center (WTC)-related illness

Mengling Liu, PhD^{1,2}, Meng Qian, MS^{1,2}, Qinyi Cheng, PhD^{1,2}, Kenneth I. Berger, MD³, Yongzhao Shao, PhD^{1,2}, Meredith Turetz, MD³, Angeliki Kazeros, MD³, Sam Parsia, MD³, Roberta M. Goldring, MD³, Maria Elena Fernandez-Beros, PhD³, Michael Marmor, PhD^{1,2,3}, and Joan Reibman, MD^{2,3,*}

¹Department of Population Health, New York University School of Medicine, USA

²Department of Environmental Medicine, New York University School of Medicine, USA

³Department of Medicine, New York University School of Medicine, USA

Abstract

Objective—The course of lung function in community members exposed to World Trade Center (WTC) dust and fumes remains undefined. We studied longitudinal spirometry among patients in the WTC Environmental Health Center (WTCEHC) treatment program.

Methods—Observational study of 946 WTCEHC patients with repeated spirometry measures analyzed on the population as a whole and stratified by smoking status, initial spirometry pattern and WTC-related exposure category.

Results—Improvement in forced expiratory volume (FVC; 54.4 ml/year; 95% CI: 45.0-63.8) and forced expiratory volume in one second (FEV₁; 36.8 ml/year; 95% CI: 29.3-44.3) was noted for the population as a whole. Heavy smokers did not improve. Spirometry changes differed depending on initial spirometry pattern and exposure category.

Conclusions—These data demonstrate spirometry improvement in select populations suggesting reversibility in airway injury and reinforcing the importance of continued treatment.

Background

The destruction of the World Trade Center (WTC) towers on September 11, 2011 released massive amounts of dust, gas and fumes with potential environmental and occupational exposures for thousands of individuals, including community members who lived (Residents) and worked (Local workers) in the area, as well as for those involved in rescue, recovery (Rescue/recovery workers) and clean-up (Clean-up workers). These groups were potentially exposed via inhalation to the initial WTC dust clouds created as the buildings collapsed, resuspended dust from incompletely cleaned indoor and outdoor areas, fumes from fires that burned for four months, and exposure from prolonged clean-up activities. (1)

*To whom correspondence should be addressed: Joan Reibman, MD, 550 1st Avenue, Room NB7N24, New York University School of Medicine 011-New York, NY 10016, Tel: 212-263-6479, Fax: 212-263-8442, joan.reibman@nyumc.org.

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Settled outdoor and indoor dust was composed of highly alkaline materials (pH 11) consisting of pulverized concrete, fiberglass, glass, plastics and other building materials and contained polycyclic aromatic hydrocarbons, volatile organic compounds, lead, dioxin, and furans.(2), (3)

Persistent WTC-related adverse medical and mental health effects have been well described in rescue and recovery workers as well as in community members.(4-8) These studies suggest that most exposed symptomatic individuals have asthma-like symptoms. Detailed evaluations have suggested that many have bronchial hyperresponsiveness (9), small airways disease (10, 11) and some have interstitial lung disease. (12, 13) Two longitudinal studies of spirometry in rescue and recovery workers showed loss of lung function. (14, 15) Community members who lived or worked in the vicinity of the WTC towers have symptoms that are similar to those of rescue and recovery workers, but their potential exposures and demographics may differ (8). Longitudinal studies have not been reported for community members with potential WTC exposure.

The Bellevue Hospital Center World Trade Center Environmental Health Center (WTCEHC) is a medical and mental health treatment program that focuses on community members including Residents, Local workers, and Clean-up workers.(8) Launched in 2005 through joint efforts of the local communities, organized labor, and the medical community, the program was initially funded by philanthropic organizations and subsequently by city and federal agencies. Evaluation and treatment guidelines were developed for common health complaints, which were predominantly upper and lower respiratory symptoms, and patients were treated for asthma-like symptoms unless further evaluation revealed a different diagnosis. We now report longitudinal analyses of repeated spirometry measurements in patients who enrolled in the WTCEHC and presented for repeated treatment or monitoring between 2005 and 2011. Our objectives are to characterize longitudinal changes in spirometry in the WTCEHC population as a whole and after stratification for smoking, baseline lung function pattern, and WTC-related exposure category to assist our understanding of disease evolution in patients enrolled in the WTC medical treatment program.

Methods

Subjects

Individuals were self-referred to the WTCEHC with medical and/or mental health symptoms related to September 11, 2001 exposures as previously described. (8) The Institutional Review Board of New York University School of Medicine approved the research database (NCT00404898). Only data from patients who provided informed consent were used for analysis.

Procedures

Clinic protocol—At enrollment, patients responded to a multi-dimensional interviewer-administered questionnaire that included characterizations of WTC-related exposures and potential exposure category as Residents, Local worker, and Clean-up workers or Rescue/recovery, respiratory symptoms and history of tobacco use. The WTCEHC treatment protocol included treatment of asthma-like symptoms based on guidelines for asthma management,(16) with patients evaluated with additional studies if findings were inconsistent with an asthma diagnosis.

Spirometry—All individuals were referred for screening spirometry at time of enrollment, and at repeat clinic visits for therapeutic evaluation, or at visits scheduled for routine

monitoring. Spirometry was performed in accordance with American Thoracic Society/ European Respiratory Society standards (17) on a Viasys Vmax spirometer (Yorba Linda, CA). Spirometry data were electronically downloaded along with an automated quality assurance code. All studies were performed in the Bellevue Hospital Center Pulmonary Function (PFT) Laboratory. Data for pre-bronchodilator forced vital capacity (FVC) and forced expiratory volume in one second (FEV_1) are presented in liters (L) or as % of predicted (% FVC and % FEV_1) derived from NHANES III.(18) Patients were categorized as having a spirometry pattern of Normal, Low FVC, Obstructed, or Low FVC/obstructed at enrollment using definition based on the lower limits of normal as previously described.(19, 20)

Definitions—Individuals were classified as having WTC dust cloud exposure if they reported being in the initial WTC dust cloud from the collapsing buildings. Patients were characterized as Resident, Local worker, Rescue/recovery worker, Clean-up worker, or Other, based on their initial questionnaire responses about residence and workplace.(8) Patients who reported > 5 pack-year history of tobacco use were defined as heavy smokers. Respiratory symptoms were defined by at least one symptom of wheezing, chest tightness, dyspnea, or nasal or sinus symptoms with onset after September 11, 2001 and occurring with a frequency \geq 2 times per week in the month before enrollment in the WTCEHC. A patient's "last visit" was considered the visit closest to December 12, 2011 when the analysis was conducted.

Patient selection—Patients were included for analysis if they had no respiratory symptom prior to September 11, 2001, had valid longitudinal spirometry defined as an acceptable baseline spirometry measurement, and had at least one valid repeated measurement. Patients with follow-up spirometry duration less than 90 days were excluded.

Statistical methods

Continuous variables were summarized using mean and standard deviation (SD) and compared across groups using the nonparametric Kruskal-Wallis test. Categorical variables were summarized by counts and proportions and compared using Chi-square test. (21) Crude estimates of within-individual annual change of spirometry measurements were calculated as the difference between the last and first spirometry measurements divided by the time duration. Initial analyses using these crude estimates as outcomes examined potential factors that might be associated with the temporal change in spirometry, so these could subsequently be used as stratification variable in longitudinal analysis. We considered the following variables: age at baseline, gender, race/ethnicity, income, body mass index (BMI), smoking status, dust cloud exposure, WTC-related exposure category and baseline spirometry patterns. A linear mixed-effects model was used to investigate longitudinal changes of lung function using all repeated spirometry measurements.(22, 23) Separate models were fit with FVC, % FVC, FEV_1 and % FEV_1 as dependent variables for the entire longitudinal population and subsequently for strata defined by smoking status, baseline spirometry pattern or WTC exposure category. In each model, a fixed linear effect of the follow-up time (defined as duration since joining the program) was estimated with adjustment for potential confounders. Random intercept and slope were assumed to explain within-subject correlation among repeated measurements and among-subject heterogeneity. We used the alpha level of 0.05 to declare significance for tests and analyses in the whole population, and alpha level of 0.01 in the stratified analyses to account for multiple testing issue. Statistical analyses were conducted using SAS (version 9.2).

Results

Baseline characteristics

The final study population consisted of 946 subjects who enrolled in the WTCEHC between August 17, 2005 and December 14, 2011, had new onset and persistent respiratory symptoms, and fit criteria for longitudinal assessment (Figure 1). The population was diverse as can be seen by the characteristics of the study population (Table 1): 478 (50.5%) were female and 429 (52.5%) were Hispanic. Many, 445 (47.6%) reported being caught in the WTC dust cloud. The largest group with potential WTC exposure was the Local worker group (438; 46.3%). Heavy smokers comprised 21.1% of the group (196; 21.1%). Most (86.1%) had current respiratory symptoms.

When classified by baseline spirometry pattern (Table 1), the most common abnormal pattern was Low FVC, a finding similar to that previously published.(8, 15, 19, 24) Distributions of Race/ethnicity, BMI, and tobacco differed among spirometry patterns. WTC dust cloud exposure was not associated with baseline spirometry pattern, whereas the distribution of WTC exposure categories differed among spirometry patterns ($p = 0.014$) with the Low FVC/Obstructed group having the highest proportion of Local workers.

Longitudinal analysis

The average number of repeated observations was 3 (range 2-10). The average duration of follow-up was 2.43 years ($SD = 1.37$ years) and the longest was 5.94 years. Initial analyses, based on crude estimates of individual annual change (Appendix Table 1), indicated that smoking status, spirometry pattern and WTC-related exposure category might be associated with the temporal change of FVC, % FVC, FEV_1 , and % FEV_1 , but not WTC dust cloud exposure or the presence of respiratory symptoms. Therefore, we conducted longitudinal analyses in the population as whole and then stratified by smoking status, spirometry pattern and WTC-related exposure category.

Annual change of FVC, % FVC, FEV_1 , and % FEV_1 estimated via linear mixed-effects models adjusted for appropriate variables, as shown in Table 2, revealed significant improvement in spirometry parameters in the overall sample: with a 54.4 ml/year improvement in FVC (95% CI: 45-63.8) and a 36.8 ml/year in FEV_1 (95% CI: 29.3-44.3). These changes translated into a 1.37 %/year (95% CI: 1.11-1.64) improvement in % FVC and a 1.12 %/year (95% CI: 0.85-1.38) improvement in % FEV_1 . Temporal trends in lung function differed between heavy-smokers and non- or light- smokers (Table 2), with the non- or light- smoker group showing significant ($p < 0.0001$) improvement in all spirometry parameters, whereas no improvement was observed among heavy smokers.

Longitudinal analysis and spirometry pattern

Changes in longitudinal spirometry differed between each spirometry pattern group (Table 2). The Normal group demonstrated improvement in both FVC and FEV_1 : FVC (47.3; 95% CI: 36.5-58.2), % FVC (1.16; 95% CI: 0.85-1.47), FEV_1 (29.2; 95% CI: 20.7-37.7) and % FEV_1 (0.82; 95% CI: 0.5-1.14). In the groups with abnormal spirometry, the Low FVC group and the Low FVC/Obstructed group demonstrated significant increases in all spirometry parameters. In contrast, the Obstructed group only showed a significant increase in FEV_1 (50.2; 95% CI: 14.2-84.2). The Low FVC/Obstructed group had the greatest improvement for both FVC and FEV_1 .

We further examined spirometry measurements as predicted values at the participants' last visit to the WTCEHC stratified by initial spirometry patterns (Figure 2). At their last visit, the Low FVC group and the Low FVC/Obstructed group continued to have abnormal

spirometry with median values of % FVC and % FEV₁ less than 80%. The Obstructed group had lower FEV₁ measurements compared to the Normal group at the last visit.

Longitudinal spirometry and WTC exposure category

When stratified by WTC-related exposure category (Table 2), longitudinal analysis indicated statistically significant improvement in all lung function parameters among Local worker, Resident, Rescue/recovery and Clean-up workers, with the exception of % FEV₁ among the Clean-up group ($p = 0.01$). The Rescue/recovery group demonstrated the greatest annual improvement in all lung function parameters, whereas the Local worker group showed the least improvement.

Discussion

We report longitudinal assessment of spirometry parameters in a group of community members with potential for WTC dust exposures. Our analyses of a group of patients with heterogeneous exposures to environmental contaminants related to the September 11 disaster showed statistically significant temporal improvements in lung function in the group as a whole, with the degree of improvement varying by the pattern of spirometry at enrollment, the category of WTC-related exposure, and smoking status.

We demonstrated overall improvement in spirometry parameters during an average 2.4-year follow-up period in this community cohort enrolled in a treatment program about four years after the WTC disaster on September 11, 2001. These data are in contrast to longitudinal analyses of lung function in those involved in rescue and recovery.(15), 19 In firefighters, an initial loss of lung function was detected within the first year after exposure with subsequent decline in lung function approximating the expected age related change.(14) Analysis of an additional rescue and recovery population also demonstrated a reduction of lung function over time although analysis was limited by lack of antecedent data.(15) Our baseline observational period started four years after the event, and unlike data in firefighters, we did not have lung function data prior to or within the immediate years after exposure. The increase in spirometry measurements is also in contrast to the anticipated loss of 31 ml/year in FEV₁ that has been reported in longitudinal studies of non-smokers with asthma and the loss in lung function in patients with COPD.(25, 26)(27) The improvement in lung function in the WTCEHC population suggests the presence of a reversible component of lung injury in the population despite the delayed time of entry into the treatment program.

We demonstrated heterogeneity in response over time among the spirometry patterns. The distribution of the four spirometry patterns in our population was consistent with other populations (19) and with the spectrum of diseases that have been described for WTC exposed individuals, which include reactive airways dysfunction, irritant-induced asthma, sarcoidosis and other interstitial lung diseases.(4, 6, 12, 28, 29) The variability in improvement among the four spirometry patterns suggests different mechanisms for the development of each pattern. A Normal spirometry pattern can be seen in patients without any underlying lung disease, but is also consistent with asthma or airway hyper-responsiveness. Alternatively, exposed patients with the normal pattern may have distal airway disease that is not reflected by spirometry.(30, 31) Indeed, the improvement in FVC in this population, and the absence of decline in FEV₁, suggested the presence of reversible airway closure even in this group with a Normal spirometry pattern. Although most patients with WTC exposures and lower respiratory symptoms have been suggested to have asthma, an Obstructed pattern is uncommon in this as well as other reported WTC exposed cohorts. (19)The Obstructed group improved both FVC and FEV₁ parameters, consistent with airway disease. The Low FVC pattern has been described as the predominant spirometry abnormality in WTC-exposed populations.(19) A reduced FVC can be consistent with sub-

maximal effort, parenchymal disease, obesity, or alternatively can be associated with patchy peripheral air trapping.(20) Patients with the Low FVC pattern predominantly improved FVC over time, consistent with a component of reversible airflow obstruction. Patients with the combined Low FVC/Obstructed pattern had the most abnormal lung function at baseline yet the greatest improvement in spirometry. However, lung function in this group, as well as in the Low FVC group failed to return to the normal range. These findings suggest that all groups had some potential for reversibility of injury, however, the components that improved differed, reinforcing differences in type or location of injury or response to treatment.

The WTCEHC includes individuals with a variety of potential exposures to the WTC dust and fumes and we grouped patients according to their potential for exposure. Longitudinal changes in lung function differed in these WTC exposure categories, with the Local workers demonstrating the least improvement. The Local worker category included people who worked either within the WTC towers or worked in surrounding buildings, many of whom evacuated on September 11, 2001 and returned to work in the surrounding buildings one week later. This category also had the highest likelihood of dust cloud exposure. We did not see a relationship with dust cloud exposure and temporal longitudinal changes in lung function in the cohort as a whole, however, the possibility remains that this intense exposure combined with subsequent exposures influenced disease pattern and longitudinal change in a specific category. Further detailed characterization of the specific exposure within each of these categories is warranted to improve our understanding of a dose-response relationship. Moreover, the data suggest that close observation of the Local worker group is warranted.

The use of tobacco had an important influence in our findings. In contrast to the group as a whole, we did not detect temporal improvement in lung function among heavy smokers. The finding of a lack of improvement among heavy smokers may be particularly important as the information can be used in counseling WTC-affected patients regarding the importance of smoking cessation.

The longitudinal analyses were modeled with a linear time trend for changes in spirometry over time and showed improvement in lung function over the observation period. However, it would be unreasonable to expect that this improvement would be sustained over prolonged periods of time and we expect the positive changes to be counterbalanced by normal age-related decline in spirometry measurements. Our current data were too sparse at the longer time points for a definitive long-term analysis of this issue. Exploratory analysis using an alternate model with quadratic time trend (data not shown) suggested an initial improvement in lung function followed by a subsequent decline. Future studies with a longer follow-up period will fully characterize the long-term changes in lung function in this population.

There are several potential limitations to this study. Although patients were treated in a standardized manner and were provided medications free of charge, we do not have data to assess medication adherence and thus cannot directly attribute changes in lung function to treatment. Nevertheless, the improvements in FVC and FEV₁ suggest the potential for reversibility of airway injury. The possibility exists that the initial spirometry measurements were suboptimal and improved over time with patient experience. However, we used stringent criteria for inclusion of spirometry values making this explanation unlikely. Previous investigators have demonstrated that the reduced FVC pattern is associated with weight gain;(15) however, our patients with a reduced FVC improved despite the absence of significant changes in weight over time (data not shown). Our data are collected in the practical setting of a treatment program with patient's follow-up visits being clinically indicated or after a routine monitoring visit, and only 39% of symptomatic patients with valid baseline spirometry measurement had follow-up visits raising the possibility of bias in

our results. We acknowledge that the findings might differ for a general population but these findings appear to have important implications for WTC treatment programs.

In summary, we have observed an improvement in lung function among community members of the WTCEHC during their initial years of enrollment. The improvement in lung function suggests reversible injury amenable to treatment. The different initial lung function patterns and their varied responses over time reinforce the heterogeneity of potential lung injury. The effect of tobacco use on lung function reinforces that WTCEHC patients should be urged to avoid other potential sources of lung injury and in particular to stop all exposures to tobacco. Although subjects with abnormal lung function improved, predicted spirometry measurements did not return to normal values and differences were noted between exposure categories. The residual abnormalities reinforce the need for treatment and continued monitoring for populations affected by this environmental disaster and suggest the importance of clinical interventions for other environmental disasters.

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Appendix

Appendix Table 1:

Effects on average annual lung function change of FEV₁ and FVC (ml/year) and FEV₁% and FVC% (percent/year) in total longitudinal population¹

	FVC ^a		FVC% ^a		FEV ₁ ^a		FEV ₁ % ^a	
	Estimate	95% CI ^b	Estimate	95% CI ^b	Estimate	95% CI ^b	Estimate	95% CI ^b
Dust cloud								
Yes	7	(-14.3,28.4)	0.06	(-0.54, 0.66)	7.4	(-9.5,24.3)	0.32	(-0.30,0.95)
No	Ref		Ref		Ref		Ref	
Tobacco								
Heavy	-49.4 ^{***}	(-74.8, -24.0)	-1.13 ^{**}	(-1.85, -0.41)	-25.2 [*]	(-45.3,-5.1)	-0.97 [*]	(-1.72, -0.23)
Light/none	Ref		Ref		Ref		Ref	
Spirometry pattern								
Normal	Ref		Ref		Ref		Ref	
Low FVC	42.2 ^{**}	(16.7, 67.7)	0.85 [*]	(0.13, 1.58)	22.6 [*]	(2.1,43.0)	0.81 [*]	(0.53,1.56)
Obstructed	-15.8	(-52.9, 21.3)	-0.55	(-1.59, 0.49)	13	(-17.0, 42.9)	0.32	(-0.79,1.43)
Obstructed and low FVC	80.1 ^{**}	(25.2,135.0)	2.55 ^{**}	(1.04,4.07)	87.2 ^{**}	(42.1,132.4)	3.31 ^{***}	(1.68,4.94)
Exposure category								
Local workers	Ref		Ref		Ref		Ref	
Clean-up workers	35.2 [*]	(4.5, 65.9)	0.45	(-0.42,1.31)	31.7 [*]	(7.4,56.0)	0.65	(-0.26,1.55)

	FVC ^a		FVC% ^a		FEV ₁ ^a		FEV ₁ % ^a	
	Estimate	95% CI ^b	Estimate	95% CI ^b	Estimate	95% CI ^b	Estimate	95% CI ^b
Resident	45.8*	(16.6,75.0)	0.81	(-0.01,1.63)	21.6	(-1.6, 44.8)	0.6	(-0.27,1.46)
Rescue/recovery	22.5**	(38.1,126,2)	2.62***	(1.44,3.80)	43.4*	(8.7,78.1)	1.07	(-0.17,2.31)
other	-11.7	(-79.8,56.5)	-0.71	(-2.53, 1.10)	13.2	(-42.3,68.6)	-0.22	(-2.19,1.74)
Respiratory symptoms								
No	27.1	(-1.9,56.1)	0.71	(-0.11, 1.53)	8.5	(-14.5, 31.4)	0.39	(-0.46, 1.25)
Yes	Ref		Ref		Ref		Ref	

¹ All results were from linear regression models adjusted with age, gender, race, BMI, income, and lower respiratory symptom.

^a After removing outliers, regression models were fit with 820, 841, 828, and 825 subjects for FEV₁, FEV₁%, FVC, and FVC% respectively.

^b* : P-value<0.05;

** : P-value<0.005;

*** : P-value<0.0005

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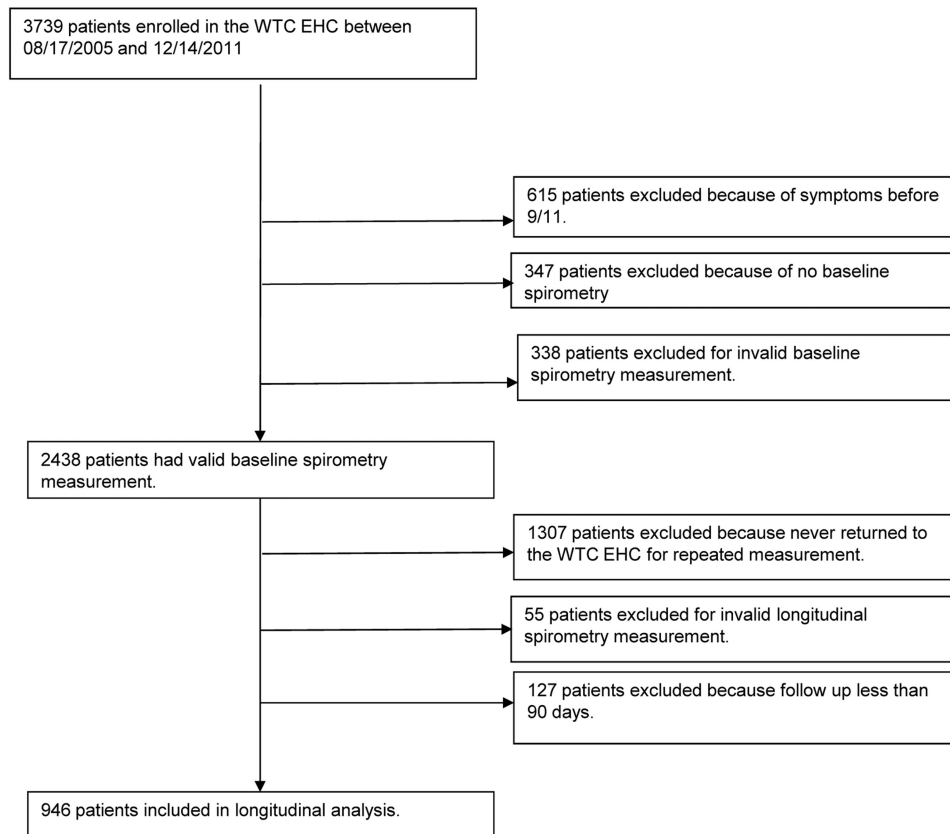


Figure 1. Flow diagram of patient identification for longitudinal analysis

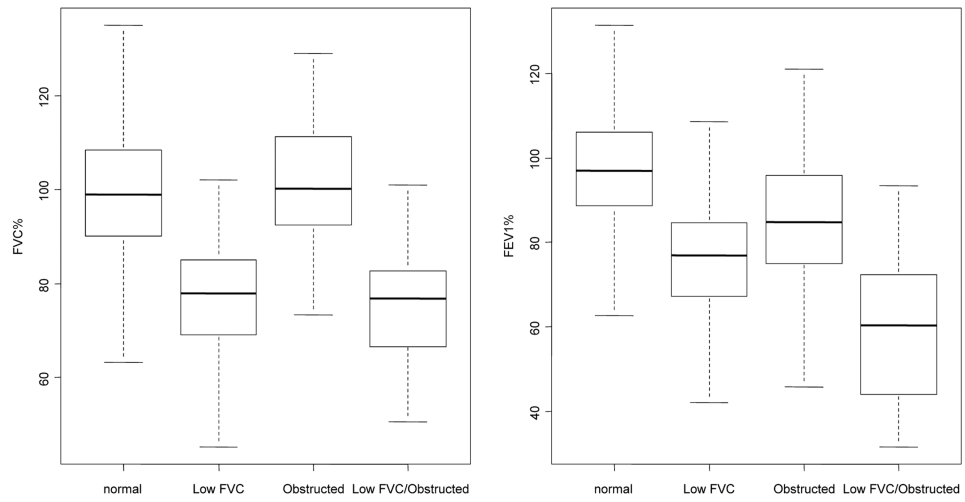


Figure 2. Boxplots of spirometry parameters at last-visit for each lung function subgroup¹
¹Outlier data points more than 1.5 times the interquartile range from the box are not shown.

Table 1

Baseline characteristics of the WTC EHC longitudinal population (N=946) as whole and stratified by baseline spirometry pattern.

Variable ^a	Longitudinal population N=946	Normal N=642	Low FVC N=193	Obstructed N=75	Low FVC/obstructed N=36	p-value ^b
Age						0.16
Mean(SD)	50.2(11.1)	50.0(11.4)	50.5(10.2)	49.6(10.8)	53.5(11.3)	
Gender N(%)						0.08
Female	478(50.5)	340 (53.0)	93 (48.2)	32(42.7)	13(36.1)	
Male	468(49.5)	302 (47.0)	100(51.8)	43(57.3)	23(63.9)	
Race/Ethnicity N(%)						<.0001
Hispanic	429 (52.5)	304(47.4)	88(45.6)	28(37.3)	9(25.0)	
White	257 (27.2)	173(27.0)	41(21.2)	25(33.3)	18(50.0)	
Black	163(17.2)	118(18.4)	26(13.5)	15(20.0)	4(11.1)	
Asian	68(7.2)	32(5.0)	28(14.5)	5(6.7)	3(8.3)	
Other	29(3.1)	15(2.3)	10(5.2)	2(2.7)	2(5.6)	
Income/year						0.05
15K	434(46.8)	284(45.0)	104(55.3)	33 (45.2)	13 (37.1)	
15K-30K	155(16.7)	108(17.1)	31(16.5)	13 (17.8)	3(8.6)	
>30K	338(36.5)	239(37.9)	53(28.2)	27(37.0)	19(54.3)	
BMI N(%)						0.0023
Normal	249(26.3)	164(25.6)	47(24.4)	33(44.0)	5(13.9)	
Over weight	353(37.3)	248(38.6)	63(32.6)	25(33.3)	17(47.2)	
Obesity	344(36.4)	230(35.8)	83(43.0)	17(22.7)	14(38.9)	
Tobacco N(%)						0.0016
5pack-year	733(78.9)	512(81.1)	151(79.5)	46(63.0)	24(68.6)	
>pack-year	196(21.1)	119(18.9)	39(20.5)	27(37.0)	11(31.4)	
Caught in WTC dust cloud N(%)						0.93
No	489(52.4)	330(52.1)	104(54.2)	37(50.0)	18(51.4)	
Yes	445(47.6)	303(47.9)	88(45.8)	37(50.0)	17(48.6)	
WTC exposure category N(%)						0.014

Variable ^a	Longitudinal population N=946	Normal N=642	Low FVC N=193	Obstructed N=75	Low FVC/obstructed N=36	p-value ^b
Local worker	438(46.3)	294(45.8)	94(48.7)	30(40.0)	20(55.6)	
Clean up worker	247(26.1)	185(28.8)	45(23.3)	12(16.0)	5(13.9)	
Resident	172(18.2)	108(16.8)	39(20.2)	20(26.7)	5(13.9)	
Rescue/recovery	67(7.1)	42(6.5)	13(6.7)	8(10.7)	4(11.1)	
Other	22(2.3)	13(2.0)	2(1.0)	5(6.7)	2(5.6)	
Respiratory symptoms						0.21
No	130(13.9)	94(14.8)	28(14.7)	6(8.2)	2(5.7)	
Yes	802(86.1)	540(85.2)	162(85.3)	67(91.8)	33(94.3)	

^aData were missing on "Caught in WTC dust cloud" for 12 subjects, "Income" for 19 subjects, "Tobacco" for 17 subjects, and "Lower respiratory symptom" for 14 subjects.

^bP-value were from Chi-square or ANOVA tests comparing baseline characteristic across spirometry pattern subgroups of the longitudinal population.

Estimates of the linear annual change of FEV₁ and FVC (ml/year) and FEV₁% and FVC% (percent/year) from the linear mixed-effects model for the total longitudinal population¹, each lung pattern subgroup¹, and each smoking category².

Table 2

	No.	FVC			FVC%			FEV ₁			FEV ₁ %		
		Estimate ^a	95% CI	Estimate ^a	95% CI	Estimate ^a	95% CI	Estimate ^a	95% CI	Estimate ^a	95% CI	Estimate ^a	95% CI
Total population¹	946	54.4 ^{***}	(45.0,63.8)	1.37 ^{***}	(1.11,1.64)	36.8 ^{***}	(29.3,44.3)	1.12 ^{***}	(0.85,1.38)				
Smoking status²													
Light/none	733	60.2 ^{***}	(50.3,70.2)	1.49 ^{***}	(1.21,1.78)	41.9 ^{***}	(33.7,50.2)	1.25 ^{***}	(0.96,1.54)				
Heavy	196	25.6	(-1.1,52.3)	0.81	(0.14,1.49)	13.1	(-4.7,30.9)	0.47	(-0.17,1.11)				
Spirometry pattern³													
Normal	642	47.3 ^{***}	(36.5,58.2)	1.16 ^{***}	(0.85,1.47)	29.2 ^{***}	(20.7,37.7)	0.82 ^{***}	(0.50,1.14)				
Low FVC	193	72.8 ^{***}	(52.9,92.6)	2.02 ^{***}	(1.50,2.54)	37.8 ^{***}	(22.9,52.8)	1.31 ^{***}	(0.81,1.82)				
Obstructed	75	17.9	(-26.5,62.4)	0.14	(-1.18,1.45)	50.2 [*]	(14.2,84.2)	1.45	(0.21,2.69)				
Low FVC/Obstructed	36	122.4 [*]	(45.0,199.8)	3.93 ^{***}	(2.18,5.68)	153.1 ^{***}	(82.0,224.3)	5.13 ^{***}	(2.98,7.28)				
Exposure category⁴													
Local worker	438	28 ^{**}	(13.0,43.0)	0.91 ^{***}	(0.51,1.32)	24.6 ^{***}	(12.3,36.8)	0.81 ^{**}	(0.39,1.23)				
Resident	172	59.5 ^{***}	(31.3,87.7)	1.71 ^{***}	(0.84,2.57)	44.5 ^{***}	(25.4,63.5)	1.57 ^{**}	(0.76,2.38)				
Rescue/recovery	67	122.8 ^{***}	(86.1,159.6)	3.01 ^{***}	(2.22,3.81)	80.4 ^{***}	(52.3,108.5)	2.47 ^{***}	(1.68,3.26)				
Clean-up worker	247	61.3 ^{***}	(46.3,76.3)	1.42 ^{***}	(1.02,1.82)	36.3 ^{***}	(24.2,48.4)	0.95	(0.53,1.37)				

¹Linear mixed effects model with a linear term of time was adjusted for baseline age, BMI, gender, race/ethnicity, income, dust-cloud exposure, WTC exposure category, smoking status, and baseline spirometry pattern.

²Linear mixed effects model with a linear term of time within each smoking group was adjusted for baseline age, BMI, gender, race/ethnicity, income, dust-cloud exposure, WTC exposure category, and baseline spirometry pattern.

³Linear mixed effects model with a linear term of time within each spirometry pattern group was adjusted for baseline age, BMI, gender, race/ethnicity, income, dust-cloud exposure, WTC exposure category, and smoking status.

⁴Linear mixed effects model with a linear term of time within each exposure category was adjusted for baseline age, BMI, gender, race/ethnicity, income, dust-cloud exposure, smoking status, and baseline spirometry pattern.

^a : P-value<0.01;

** : P-value<0.001;
*** : P-value<0.0001