

Canadian prediction equations of spirometric lung function for Caucasian adults 20 to 90 years of age: Results from the Canadian Obstructive Lung Disease (COLD) study and the Lung Health Canadian Environment (LHCE) study

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BACKGROUND: Currently, no reference or normative values for spirometry based on a randomly selected Canadian population exist.

OBJECTIVE: The aim of the present analysis was to construct spirometric reference values for Canadian adults 20 to 90 years of age by combining data collected from healthy lifelong nonsmokers in two population-based studies.

METHOD: Both studies similarly used random population sampling, conducted using validated epidemiological protocols in the Canadian Obstructive Lung Disease study, and the Lung Health Canadian Environment study. Spirometric lung function data were available from 3042 subjects in the COLD study, which was completed in 2009, and from 2571 subjects in the LHCE study completed in 1995. A total of 844 subjects 40 to 90 years of age, and 812 subjects 20 to 44 years of age, were identified as healthy, asymptomatic, lifelong nonsmokers, and provided normative reference values for spirometry. Multiple regression models were constructed separately for Caucasian men and women for the following spirometric parameters: forced expiratory volume in 1 s (FEV₁), forced vital capacity (FVC) and FEV₁/FVC ratio, with covariates of height, sex and age. Comparison with published regression equations showed that the best agreement was obtained from data derived from random populations.

RESULTS: The best-fitting regression models for healthy, never-smoking, asymptomatic European-Canadian men and women 20 to 90 years of age were constructed. When age- and height-corrected FEV₁, FVC and FEV₁/FVC ratio were compared with other spirometry reference studies, mean values were similar, with the closest being derived from population-based studies.

CONCLUSION: These spirometry reference equations, derived from randomly selected population-based cohorts with stringently monitored lung function measurements, provide data currently lacking in Canada.

Key Words: Canadian; Prediction equations; Reference values; Spirometry

Les équations prévisionnelles canadiennes de la fonction pulmonaire spirométrique des adultes de race blanche de 20 à 90 ans : les résultats de l'étude COLD sur la maladie pulmonaire obstructive au Canada et de l'étude LHCE sur la santé pulmonaire dans les milieux canadiens

HISTORIQUE : Il n'existe pas de référence ou de valeur normative de la spirométrie selon une population canadienne sélectionnée au hasard.

OBJECTIF : La présente analyse visait à établir des valeurs de référence spirométriques pour les adultes canadiens de 20 à 90 ans en combinant des données colligées auprès de personnes qui n'avaient jamais fumé tirées de deux études en population.

MÉTHODOLOGIE : De façon similaire, les deux études faisaient appel à un échantillon aléatoire en population et avaient été menées au moyen de protocoles épidémiologiques validés dans l'étude COLD sur la maladie pulmonaire obstructive au Canada et dans l'étude LHCE sur la santé pulmonaire dans les milieux canadiens. L'étude COLD, terminée en 2009, possédait les données de fonction pulmonaire spirométrique de 3 042 sujets, et l'étude LHCE, terminée en 1995, les données de 2 571 sujets. Au total, 844 sujets de 40 à 90 ans et 812 sujets de 20 à 44 ans étaient considérés comme des personnes asymptomatiques en santé qui n'avaient jamais fumé et ont fourni les valeurs de référence normatives de la spirométrie. Les chercheurs ont créé des modèles de régression multiple distincts pour les hommes et les femmes de race blanche à l'égard des paramètres de spirométrie suivants : volume expiratoire maximal par seconde (VEMS), capacité vitale forcée (CVF) et rapport VEMS/CVF, incluant les covariables de la taille, du sexe et de l'âge. La comparaison avec les équations de régression publiées a démontré que le meilleur appariement provenait de données tirées de populations aléatoires.

RÉSULTATS : Les meilleurs modèles de régression ont été créés à partir d'hommes et de femmes euro-canadiens asymptomatiques en santé de 20 à 90 ans qui n'avaient jamais fumé. Lorsque la VEMS, la CVT et le rapport VEMS/CVF corrigés selon l'âge et la taille étaient comparés avec d'autres études de référence de la spirométrie, les valeurs moyennes étaient similaires, les plus près étant dérivées d'études en population.

CONCLUSION : Ces équations de référence de la spirométrie, dérivées de cohortes en population sélectionnées de manière aléatoire, dont les mesures de la fonction pulmonaire avaient été rigoureusement surveillées, fournissent des données encore inexistantes au Canada.

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Spirometry is the most common pulmonary function test used in the objective assessment of patients with respiratory symptoms in clinical practice because it is readily accessible, relatively easy to perform and reliable under conditions of strict quality control (1,2).

Correct interpretation of spirometry in clinical practice requires comparison with previously published reference values (3). The American Thoracic Society (ATS) criteria for constructing reference values recommended that they be based on a large sample of representative populations, the use of internationally accepted equipment and methods, and appropriate statistical analysis (4). Ideally, normative values should be derived from healthy individuals identified in population-based surveys such as in the The National Health and Nutrition Examination Survey (NHANES) Study (5). However, population surveys with standardized and quality-assured lung function testing remains uncommon (5-8). In Canada, a recent publication (9) reported values for Caucasian Canadians 20 to 80 years of age that were based on lung function tests in 637 healthy nonsmokers selected from relatives and friends of patients and employees of the six participating sites. To date, no population-based spirometry reference values for Canada have been published. The availability of data from two large population-based studies – the Lung Health and Canadian Environment (LHCE) study (10,11), and the Canadian Obstructive Lung Disease (COLD) study – based on the Burden of Obstructive Lung Disease (BOLD) protocol (12-14) provided the opportunity to generate reference values for spirometry from randomly selected populations 20 to 90 years of age from across Canada.

The aim of the present analysis was to construct spirometric reference values for Canada in adults 20 to 90 years of age with no history of respiratory disease, by combining data from healthy, lifelong nonsmokers based on these two population-based studies, and to test how the prediction equations for spirometric measurements compare with other published American equations (5,9,15-18).

METHODS

Patients

LHCE study: The details of the LHCE study are fully described in previous publications (10,11). Briefly, 18,616 individuals randomly selected from the community from six sites across Canada, answered the European Community Respiratory Health Survey (ECRHS) questionnaire. Preselected random subsamples were invited to undergo lung function testing in a research laboratory. Spirometry was performed using a dry rolling-seal spirometer (Grasbe-Andersen, Spirotech Division, USA) using the Lung Health study protocol and computer software (19). The study was conducted between March 1993 and November 1994.

COLD study: Details regarding the subjects and methodology in the COLD study have been described in previous publications (12,14,20,21). This epidemiological study to determine the prevalence of chronic obstructive pulmonary disease (COPD) was a nationwide, cross-sectional, multisite and population-based analysis that used spirometry as the main diagnostic tool. It was initiated in Vancouver (British Columbia) in August 2005, and completed at five sites (Vancouver, Montreal [Quebec], Toronto [Ontario], Halifax [Nova Scotia] and Calgary [Alberta]) by May 2009. Briefly, the strategy and instruments were the same as those used in the international BOLD initiative (12,14,20,21). Trained and certified technicians conducted interviews using the BOLD questionnaires, which contained information on respiratory symptoms and diseases, smoking history and other risk factors for COPD, medication use, health care use, activity limitation and health-related quality of life. Lung function data were obtained with the use of the ndd EasyOne spirometer (ndd Medical Technologies, USA), in which spirometry was performed before and 15 min after administration of 200 µg of salbutamol/albuterol (12-14,20). Strict quality assurance of spirograms involve scoring of all tracings based on acceptability and reproducibility criteria of the ATS and European Respiratory Society (ERS) (1,22). The present analysis used

data acquired from 3042 individuals who were 40 years of age or older, and identified healthy, never-smokers with no respiratory disease.

Inclusion criteria for reference values

To generate reference values, prebronchodilator spirometric values were used. Subjects were selected using questionnaire and anthropometric data. The exclusion and inclusion criteria were similar to those described in the study by Hankinson et al (5). Healthy subjects were never-smokers without a respiratory diagnosis of current asthma (present in the preceding 12 months), current chronic bronchitis, emphysema, COPD, tuberculosis, no hospitalization for respiratory illness in childhood, and no chronic symptoms of cough, sputum or wheeze. In addition, subjects were required to have three acceptable spirometric tracings of forced expiratory volume in 1 s (FEV₁) and forced vital capacity (FVC), with a variation of <200 mL between the two greatest values (1). The exclusion criteria for the two cohorts are shown in Tables 1 and 2.

Statistical analysis

Statistical analyses were performed using SAS version 9.1.3 (SAS Institute, USA). Reference equations and equations to calculate the 5th percentile lower limit of normal (LLN) criteria were developed during the SAS procedures PROC REG and PROC UNIVARIATE. Each of the spirometry measures was entered into a multiple regression model using age, height (cm) and sex as predictors. Nonlinear relationships were also tested by including the square of age, height, their interactions, weight and body mass index (BMI) in the regression equations. Predictor variables were retained in the regression model only if their addition significantly improved R², and the SE of the differences decreased. Men and women were analyzed separately. For each spirometric variable, in addition to the mean value, the SE of the estimate (SEE or SD of the residuals) was determined to enable the computation of the LLN, which is estimated using the following equation:

$$\text{LLN} = \text{Predicted value} - (1.645 \times \text{SEE})$$

The spirometric values generated by the study were also compared with those of seven previous spirometry reference studies: two population-based studies with randomly sampled populations, namely the North American population in the NHANES study by Hankinson et al (5), and the South American population in the Chronic Obstructive Pulmonary Disease in Five Latin American Cities (PLATINO) study (18); one Canadian study that used a selected convenience sample of subjects (Gutierrez et al [9]); and four studies that are approximately 18 to 40 years old, which generated reference values that are still widely used in Canada (2,15-17).

RESULTS

A total of 812 participants (20 to 44 years of age) from the total sample of 2892 participants (28%) in the LHCE study, and 844 (40 to 90 years of age) from the total sample of 3042 participants (28%) in the COLD study were identified as healthy, asymptomatic, lifelong Caucasian nonsmokers in Canada, and provided normative reference values for spirometry. These individuals had never smoked, had not been diagnosed with respiratory diseases and did not exhibit respiratory symptoms. The method of selection from the two study cohorts are shown in Tables 1 and 2. Table 3 summarizes the general characteristics of the subjects in the two studies.

All healthy subjects included in each study met the quality criteria proposed by the ATS and ERS (three acceptable manoeuvres and repeatability defined as readings between the two best values for FEV₁ and FVC <200 mL) (4).

Figure 1 presents scatterplots of FEV₁ and of FVC for men and women from the LHCE study (20 to 44 years of age) and the COLD study (40 to 90 years of age). The figures demonstrate the continuity of the FEV₁ and FVC values for the different age ranges examined in the two studies, and overlapping values for the ages of 40 to 44 years from the two cohorts.

TABLE 1
Exclusion criteria for the selection of healthy, nonsmoker Caucasians in the Lung Health and Canadian Environment study (age range 20 to 44 years)

Criteria	Men (n=1163)		Women (n=1408)		Total (n=2571)	
	Excluded	Remaining	Excluded	Remaining	Excluded	Remaining
FEV ₁ or FVC data missing	33	1130	65	1343	98	2473
Smoker or exsmoker	576	558	689	654	1261	1212
History of asthma	60	498	88	566	148	1064
Lung problem before 16 years of age	33	465	47	519	80	984
Cough for >3 months	21	444	42	477	63	921
Sputum production for >3 months	12	432	13	464	25	896
Wheeze apart from colds	26	406	30	434	56	840
Woken by shortness of breath	11	395	17	417	28	812

Data presented as n. FEV₁ Forced expiratory volume in 1 s; FVC Forced vital capacity

TABLE 2
Exclusion criteria for the selection of healthy, nonsmoker Caucasian subjects in the Canadian Obstructive Lung Disease study (age range 40 to 90 years)

Criteria	Men (n=1234)		Women (n=1808)		Total (n=3042)	
	Excluded	Remaining	Excluded	Remaining	Excluded	Remaining
Smoker or exsmoker	706	528	893	915	1599	1443
MD diagnosis of emphysema	2	526	6	909	8	1435
MD diagnosis of current asthma	43	483	107	802	150	1285
MD diagnosis of COPD	2	481	5	797	7	1278
MD diagnosis of tuberculosis	6	475	16	781	22	1256
MD diagnosis of current chronic bronchitis	2	473	6	775	8	1248
Non-Caucasian	67	406	151	624	218	1030
Unacceptable spirometry data	17	389	15	609	32	998
Hospitalized for breathing problems before 10 years of age	18	371	24	585	42	956
Cough for >3 months	22	349	39	546	61	895
Sputum for >3 months	7	342	16	530	23	872
Wheeze apart from colds	8	334	20	510	28	844

Data presented as n. COPD Chronic obstructive pulmonary disease; MD Physician

TABLE 3
Baseline characteristics and spirometric measurements for the healthy, nonsmoker Caucasian subjects in the Lung Health Canadian Environment (LHCE) and Canadian Obstructive Lung Disease (COLD) study populations

Variable	LHCE		COLD	
	Women (n=417)	Men (n=395)	Women (n=510)	Men (n=334)
Age, years	31.5±6.9	31.4±6.9	56.5±10.9	54.8±10.3
Height, cm	164.2±6.8	176.8±6.4	162.4±6.7	176.1±7.3
Weight, kg	66.1±13.7	81.6±13.4	71.2±16.4	85.8±16.1
Body mass index, kg/m ²	24.5±4.9	26.1±4.0	27.0±6.2	27.6±4.6
FEV ₁ , L	3.30±0.48	4.40±0.62	2.51±0.56	3.60±0.68
FVC, L	3.92±0.58	5.37±0.77	3.32±0.71	4.82±0.87
FEV ₆ , L	*	*	3.19±0.68	4.58±0.82
FEV ₁ /FVC, %	84.5±6.0	82.3±5.5	75.7±5.8	74.9±6.1
FEV ₁ /FEV ₆ , %	*	*	78.9±4.3	78.6±4.5

Data presented as mean ± SD. *Data not available. FEV₁ Forced expiratory volume in 1 s; FEV₆ Forced expiratory volume in 6 s; FVC Forced vital capacity

The prediction equations for men and women between 20 and 90 years of age compiled from the two study cohorts are shown in Table 4. The height-squared and age-squared terms were not significant for predicting FEV₁, FVC or FEV₁/FVC and, therefore, the final equations used linear terms.

Comparison of spirometry with previous studies

Table 5 summarizes the mean spirometry values for Caucasians in our combined study population, expressed as percentage of normal predicted according to previously published regression equations: four commonly used sources of spirometry prediction equations (5,15-17), and with two more recently published regression equations (9,18).

Values for spirometric variables from our study were approximately 8% higher than the FEV₁ predicted by Morris et al (17) and for FVC reported by Knudson et al (16), and approximately 5% lower than those predicted by a previous Canadian study of a healthy convenience sample reported by Gutierrez et al (9). However, they were very similar to those predicted by the regression equations reported in three other studies: for Caucasian Americans derived from the NHANES population-based study by Hankinson et al (5); for South American subjects from another population-based study (18); and for Caucasian Americans reported by Crapo et al (15). Figures 2A and 2B present FEV₁ values for Caucasian men and women, with increasing age at a constant height for our cohort and as predicted according to other published

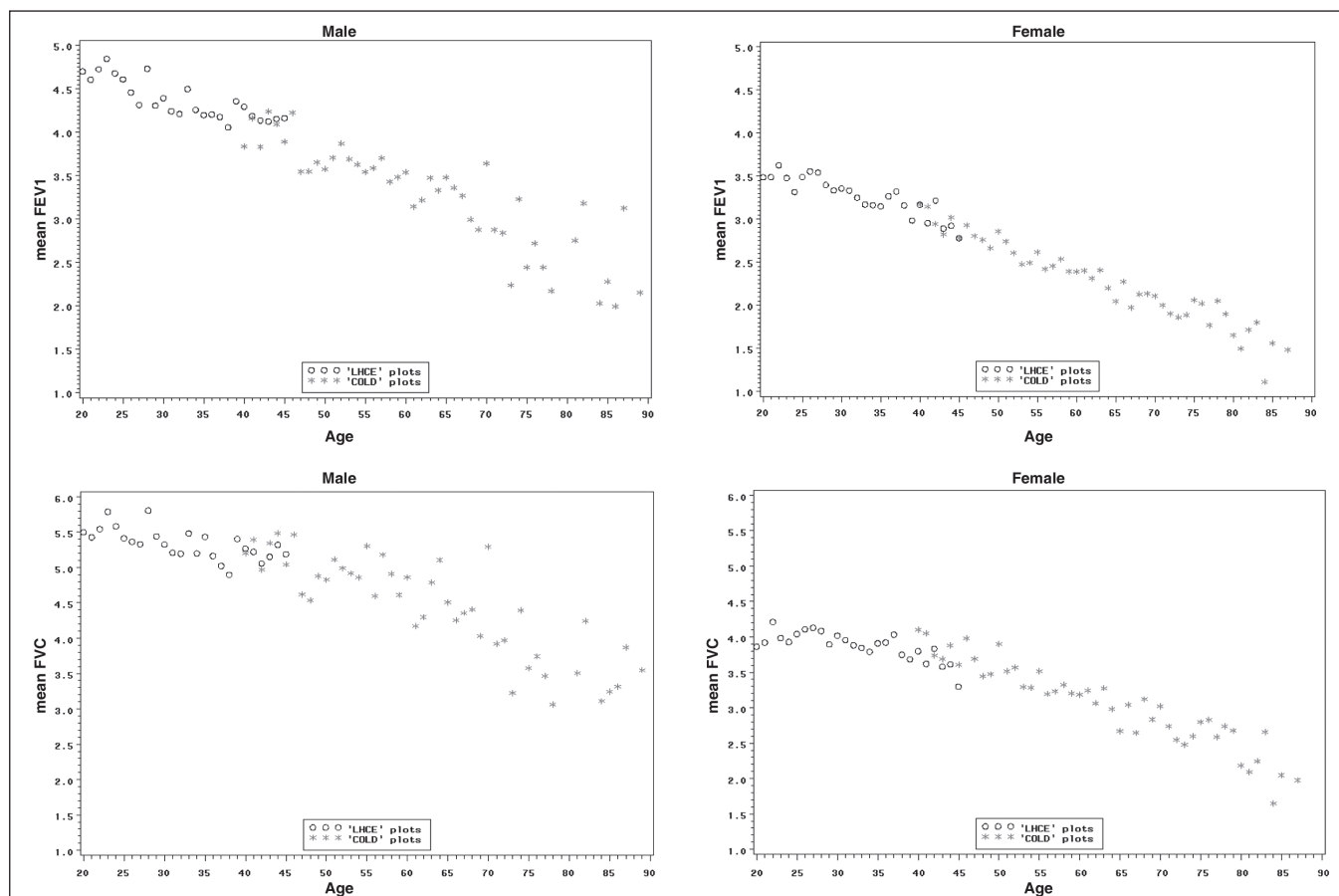


Figure 1 Scatterplots of the spirometric variables forced expiratory volume in 1 s (FEV₁) (upper panel) and forced vital capacity (FVC) (lower panel) versus age for men (upper left and lower left) and women (upper right and lower right) from the collated data of the Lung Health and Canadian Environment (LHCE) study (20 to 44 years of age) and the Canadian Obstructive Lung Disease (COLD) study (40 to 90 years of age)

TABLE 4
Reference equations for men and women between 20 and 90 years of age for predicting normal FEV₁, FVC and FEV₁/FVC ratio (n=1656)

Sex	Parameter	Intercept	Age	Height	R ²	SD of difference (SEE)*	Observations, n
Male	FEV ₁	-2.06961	-0.03167	0.04215	0.56010	0.50586	722
	FVC	-5.39383	-0.02286	0.06500	0.45720	0.63557	708
	FEV ₁ /FVC	122.0893	-0.27977	-0.17794	0.35770	5.48656	703
Female	FEV ₁	-1.68697	-0.02773	0.03557	0.68390	0.36973	926
	FVC	-4.11886	-0.02124	0.05310	0.52160	0.49381	917
	FEV ₁ /FVC	120.59691	-0.30805	-0.16531	0.39550	5.68808	916

Men: FEV₁ = 0.04215 × height (cm) - 0.03167A - 2.06961; FVC = 0.06500 × height (cm) - 0.02286A - 5.39383; FEV₁/FVC = -0.17794 × height (cm) - 0.27977A + 122.0893; Women: FEV₁ = 0.03557 × height (cm) - 0.02773A - 1.68697; FVC = 0.05310 × height (cm) - 0.02124A - 4.11886; FEV₁/FVC = -0.16531 × height (cm) - 0.30805A + 120.59691. *The 5th percentile lower limit of normal (LLN) of the population can be computed for these lung function parameters using the following formula: LLN = predicted value - (1.645 × SEE). FEV₁ Forced expiratory volume in 1 s; FVC Forced vital capacity; SEE Standard error of the estimate

regression equations. Again, the FEV₁ values at different ages for Caucasian men and women of a fixed height in the present study were most similar to those predicted by the two population-based studies (ie, NHANES [5] and the PLATINO study [18]), with overestimation by the convenience sample of the Gutierrez et al (9) regression equation and underestimation by the oldest reference equations still in use (17).

DISCUSSION

We have constructed reference regression equations for spirometry based on healthy subjects derived from two population-based studies in Canada, for predicting the normative spirometric values of FEV₁, FVC and FEV₁/FVC ratio. For the predicted values of these variables, we included the usual mean values that are most widely used for lung function interpretation, but we also provided the SEE for the

simplified determination of the 5th percentile as the LLN, which has become increasingly accepted in recent years as an alternative measure for defining airway obstruction (23-26).

The present study, which merged two studies of continuous age ranges, has the advantage of being based on representative population samples from eight major cities/sites in Canada, and of having been conducted using validated and standardized methods in accordance with international recommendations. It represents an advance in the pre-existing values used in Canada, which were calculated from convenience samples (9).

Merging the two data sets enabled us to generate regression equations for predicting normative spirometric values for use in adults across a wide age range (20 to 90 years). We were encouraged to do so by the similarities between the two studies: the sampling method was similar, and random samples of the population were obtained using

TABLE 5
Spirometry parameter values for Caucasians in the study population (COLD and LHCE cohorts combined) according to regression equations reported in other studies

Author (reference), year	Women	Men
Hankinson et al (5), 1999		
FEV ₁ , L, % predicted	98.7±13.9	99.4±13.0
FVC, L, % predicted	100.4±13.7	100.1±12.6
FEV ₁ /FVC, %	98.1±7.3	99.4±7.2
Perez-Padilla et al (8), 2006		
FEV ₁ , L, % predicted	101.6±14.0	101.1±13.0
FVC, L, % predicted	100.6±13.8	99.4±12.6
FEV ₁ /FVC, % predicted	99.7±7.4	99.3±7.1
Gutierrez et al (9), 2006		
FEV ₁ , L, % predicted	95.0±13.1	96.1±12.4
FVC, L, % predicted	100.4±13.8	96.6±12.2
FEV ₁ /FVC, % predicted	97.4±7.4	97.9±7.0
Morris et al (17), 1971		
FEV ₁ , L, % predicted	108.0±15.0	106.7±13.8
FVC, L, % predicted	104.8±14.3	102.5±13.0
FEV ₁ /FVC, % predicted	104.6±7.8	104.8±7.4
Crapo et al (3), 1981		
FEV ₁ , L, % predicted	100.3±13.8	98.4±12.9
FVC, L, % predicted	104.1±14.2	101.5±12.8
FEV ₁ /FVC, % predicted	96.9±7.0	97.2±7.2
Knudson et al (16), 1983		
FEV ₁ , L, % predicted	104.3±14.7	101.6±14.0
FVC, L, % predicted	109.0±14.8	106.0±14.5
FEV ₁ /FVC, % predicted	96.1±7.2	96.0±7.5
Quanjer et al (25), 1993		
FEV ₁ , L, % predicted	105.7±14.9	104.1±13.4
FVC, L, % predicted	114.0±16.4	110.3±14.3
FEV ₁ /FVC, % predicted	98.9±7.4	99.1±7.3

Data presented as mean ± SD. COLD Canadian Obstructive Lung Health study; FEV₁ Forced expiratory volume in 1 s; FVC Forced vital capacity; LHCE Lung Health and Canadian Environment study

either random-digit dialing or random selection from population frames (10,20). The respiratory questionnaires used in the studies contained core questions that were very similar. In the LHCE study, additional questions were included to detect asthma in subjects 20 to 44 years of age, while the COLD study, using the BOLD questionnaire, used questions aimed at detecting asthma and COPD in individuals older than 40 years of age. Both studies included stringent quality control. The LHCE cohort used the spirometry quality assurance protocols of the Lung Health study (19), while the COLD study adopted the BOLD initiative protocol for quality assurance of all spirometry tracings (13). Both of these protocols defined 'a priori' participant preparation, manoeuvre acceptability and reproducibility, technician training and performance monitoring, equipment design and calibration, and result processing to minimize short-term, intra-individual FEV₁ variability. The quality of the tests in the COLD study was excellent – 96% of the spirometry tracings fulfilled the ATS criteria for acceptability and reproducibility (1). Only data from tracings that satisfied these criteria were used for the generation of the regression equations.

In both the LHCE and the COLD studies, we used the ATS recommendation for healthy subject selection for normative reference values (4). The ATS criteria had defined 'healthy' as "never smokers, free of respiratory symptoms and disease". Standardized questions regarding cardiopulmonary symptoms and disease are contained in both the European Community Respiratory Health Survey questionnaires and the BOLD questionnaire sets used in the two studies. We attempted to

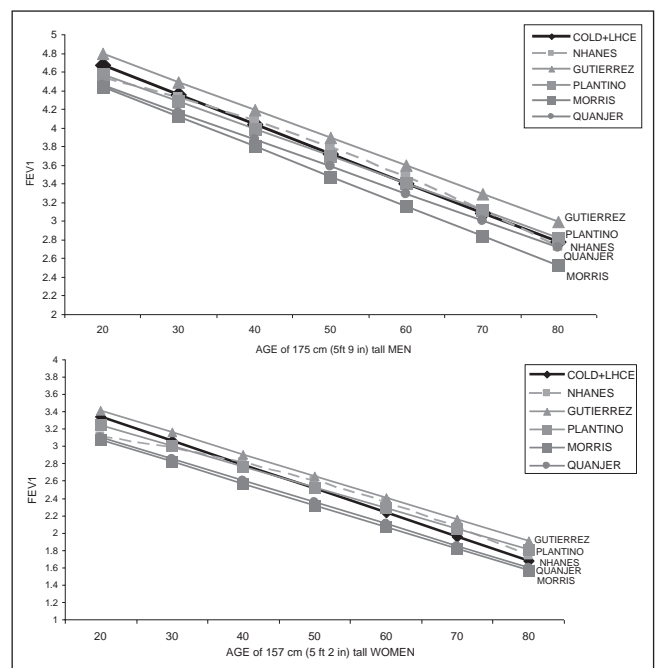


Figure 2) Predicted forced expiratory volume in 1 s (FEV₁) values for healthy men (upper panel) and for healthy women (lower panel) from combined cohorts of the COLD and LHCE studies are compared with those from other spirometry reference equations. COLD Canadian Obstructive Lung Disease study; LHCE Lung Health and Canadian Environment study; NHANES National Health and Nutrition Examination Survey

include the same list of exclusion criteria for the selection of healthy subjects from the two cohorts with small differences relevant to the two different age cohorts, such as a history of chronic bronchitis, emphysema and COPD in the older age cohort of the COLD study, and the presence of nocturnal wheezing in the younger age cohort.

A potential limitation of the present study is that the regression equations for spirometry were derived from a combined data set collated from two studies conducted 10 years apart – it is conceivable that demographic and environmental factors may have changed during the intervening time period. However, in our exploratory scrutiny of the data points, these factors did not cause real concern because the spirometric values of FEV₁ and FVC for subjects 40 to 44 years of age from the two cohorts showed good overlap, and the values with increasing age showed excellent continuity for the entire age range from 20 to 90 years. We further investigated this by generating values for FEV₁ and FVC using separate equations for the LHCE cohort and COLD cohort using standardized values for men (180 cm tall, 43 years of age) and women (160 cm tall, 43 years of age), and found that the FEV₁ differed by 190 mL for men and 100 mL for women. For FVC, the difference was 14 mL for men and 75 mL for women, which were all within the range of measurement variations (2). This further justified the merging of the two data sets.

Canada contains a large population composed of many ethnic origins (5). It would be ideal to provide lung function reference values for a variety of race/ethnic groups. In the present study, however, we were only able to generate normative spirometry values for Caucasians in Canada because the number of non-Caucasian participants who met the criteria for good respiratory health was too small (n<80) to attempt to generate separate equations according to age ranges. To generate population-based samples of major ethnic groups, including Asians and First Nations, would require additional resources and studies.

The predictive equations currently used in Canada were generated in the United States (5,15-17). We were interested in comparing these prediction equations and a set of Canadian equations derived from selected volunteers (9) with that generated from the present study.

The normative values generated from our regression equations were 8% higher for FEV₁ and 9% higher for FVC compared with those from the predictive equations of Morris et al (17) and Knudson et al (16), which are still widely used in Canada even though the data are 30 to 40 years old, and were generated using Caucasian populations of the United States.

The values predicted by our population-based, Canadian spirometric reference equations did not show perfect agreement when compared with the previously published Canadian regression equation by Gutierrez et al (9). Values for spirometric variables in healthy subjects in our study were approximately 4% to 5% lower than the predicted FEV₁ and FVC from the Gutierrez equations. A potential reason may be the difference between the healthy subjects in the two studies because healthy individuals in a convenience sample of volunteers (which was potentially biased) reported by Gutierrez et al (9) may have had 'superior' lung function than those drawn from the random representative populations in our study. It is interesting to note that the normative values generated from our regression analysis agree best with those generated by two other population-based studies – the Caucasian-American regression analysis by Hankinson et al (5), and the Latin-American regression analysis from the PLATINO study (18). These differences have clinical implications in the interpretation of lung function testing because regression equations that underestimate normative values could miss abnormalities, and equations that overestimate could overdetect abnormalities.

It would be more appropriate to use regression equations that are derived from populations most representative of the subjects to be

tested (4). Hence, the reference values in the current study represent an advancement for spirometry testing in Canada. Tables and nomograms of reference values are available for download from the CanCOLD website (www.cancold.ca). We propose that it is time for older equations, which were used as a default in the past, to be replaced by representative, population-based and Canadian-derived regression equations.

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