Table S1. ICD codes and semantic phrases considered in algorithm development

Table S1. ICD codes and semantic phrases cons	idered in algorithm development		
Inclusion Criteria for Cases	Additional Exclusion Criteria		
	for Controls		
ICD-9 Codes	ICD-9 Codes		
491.xx: Chronic bronchitis	493.xx: Asthma		
492.xx: Emphysema	516.31: Idiopathic pulmonary		
	fibrosis		
496.xx: Chronic airway obstruction, not	135.xx: Sarcoidosis		
elsewhere classified			
162.xx: Neoplasm of the trachea, bronchus,			
and lung			
ICD-10 Codes	ICD-10 Codes		
J41.xx: Simple and mucopurulent chronic	J45.xx: Asthma		
bronchitis			
J42.xx: Unspecified chronic bronchitis	J84.112: Idiopathic pulmonary		
	fibrosis		
J43.xx: Emphysema	D86.xx: Sarcoidosis		
J44.xx: Other chronic obstructive pulmonary			
disease			
C34.xx: Malignant neoplasm of bronchus and			
lung			
Problem list text			
COPD			
Chronic obstructive pulmonary disease			
Chronic obstructive lung disease			
Emphysema			
Chronic bronchitis			
Reactive airway(s) disease			
Exacerbation			
Oxygen			
O2			
Radiology report text			
Emphysema (excluding subcutaneous			
emphysema)			
Emphysematous changes			
Respiratory medications			
Accuneb			
Aclidinium			

Advair
Albuterol
Alvesco
Anoro
Arcapta
Arformoterol
Asmanex
Atrovent
Beclomethasone
Beclovent
Breo
Brovana
Budesonide
Ciclesonide
Combivent
Daliresp
Dulera
Duoneb
Entocort
Flovent
Fluticasone
Foradil
Glycopyrrolate
Indaceterol
Ipratropium
Lasma
Levalbuterol
Mometasone
Olodaterol
Perforomist
ProAir
Proventil
Pulmicort
QVar
Robinul
Roflumilast
Salbutamol

Salmeterol
Serevent
Singulair
Spiriva
Symbicort
Theo
Theochron
Theodur
Theophylline
Tiotropium
Tudorza
Umeclidinium
Uniphyl
Vanceril
Ventolin
Vilanterol
Xopenox

Table S2. Numbers of cases, controls, indeterminates, and excludes identified among adults over 45 years of age in the Synthetic Derivative (N = 1,008,552).

Case definition*	Cases, N	Control, N	Indeterminate, N	Exclude, N
3+ ICD codes	28,520	544,056	23,091	28,319
ICD code+reg-expr	12,622	544,056	38,989	28,319
PFT	1,873	11,203	NA	NA

PFT = pulmonary function test, reg ex = regular expression
\*All case definitions also include the requirement that cases have records longer than 180 days.

**Table S3**. Demographic characteristics of COPD cases identified by PFT, 3+ ICD codes, and Code + regex algorithm.

Variable	PFT cases*	3+ COPD ICD Cases	Code+Regex Cases
	N = 1,873	N = 28,520	N = 12,622
Median age (IQR)	70	69	70
	(62-76)	(60-76)	(62-77)
Sex (%)			
Female	911	13,356	6,163
	(48.6)	(46.8)	(48.8)
Male	962	15,164	6,459
	(51.4)	(53.2)	(51.2)
Unknown	0 (0.0)	0 (0.0)	0 (0.0)
Race (%)	· ·	ì	
White	1,701	24,795	11,012
	(90.8)	(86.9)	(87.2)
Black	135	2,684	1,249
	(7.2)	(9.4)	(9.9)
Other	15	119	46
	(0.8)	(0.4)	(0.4)
Unknown	22	922	315
	(1.2)	(3.2)	(2.5)
Smoking status	(1.2)	(512)	(=10)
Ever	1,464	15,645	7,548
	(78.2)	(54.9)	(59.8)
Never	402	2,900	1,034
	(21.5)	(10.2)	(8.2)
Missing	7	9,975	4,040
	(0.4)	(35.0)	(32.0)
Median record length (years) (IQR)	10.3	7.3	8.3
	(5.1-16.1)	(3.2-13.0)	(4.0-13.8)
Number of respiratory meds** (median, IQR)	188	107	170
	(72-385)	(32-277)	(56-387)
Top 5 phecodes by frequency (%)			

1	Other symptoms of respiratory system (88.7)	Chronic airway obstruction (100)	Chronic airway obstruction (100)
2	Chronic airway obstruction (78.0)	Hypertension (77.1)	Other symptoms of respiratory system (86.1)
3	Shortness of breath (71.8)	Other symptoms of respiratory system (75.4)	Hypertension (80.3)
4	Hypertension (71.8)	Essential hypertension (75.4)	Essential hypertension (78.9)
5	Essential hypertension (70.7)	Tobacco use disorder (63.0)	Shortness of breath (73.5)

<sup>\*</sup>Pulmonary function testing (PFT) cases are defined as individuals with a post-bronchodilator FEV<sub>1</sub>/FVC ratio < 0.7.
\*\*Number of respiratory medications includes only mentions of medications on distinct days.

Table S4. Association of known clinical risk factors with COPD in each algorithm set and in the PFT set.

	3+ code only		Code + regex		PF	Ts
Risk factor	OR	95% CI	OR	95% CI	OR	95% CI
Age	1.04	1.04-1.04	1.05	1.05-1.05	1.04	1.03-1.04
Sex	1.40	1.37-1.43	1.27	1.23-1.31	1.22	1.11-1.34
Ever smoking	9.22	8.87-9.57	12.36	11.61-13.16	3.36	3.00-3.75

OR: odds ratio; 95% CI: 95% confidence interval

**Table S5**. Comparison of demographics and clinical features for cases (post-bronchodilator FEV<sub>1</sub>/FVC < 0.7) and controls identified by PFTs.

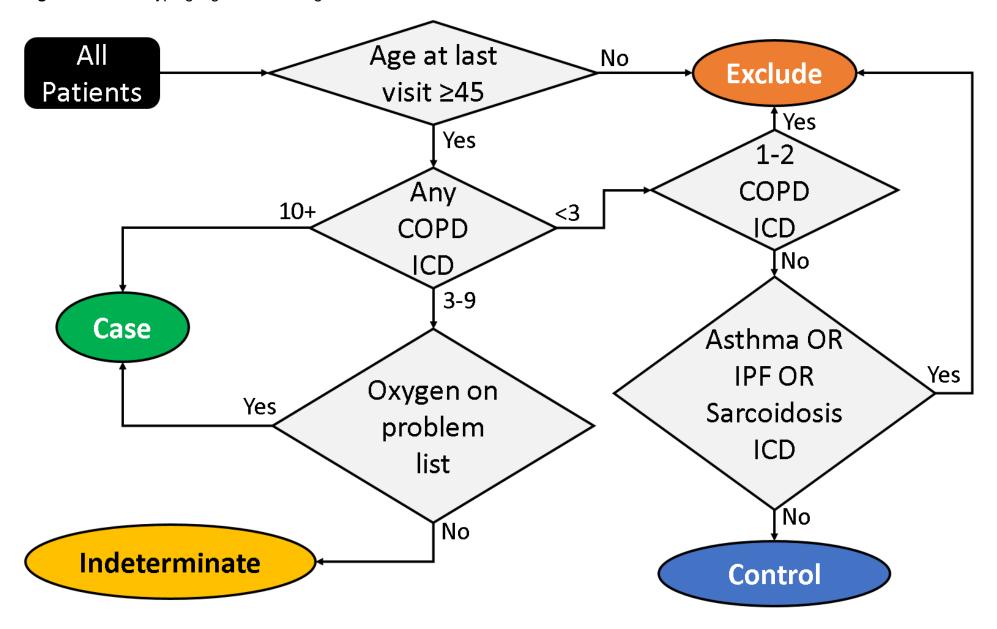
Characteristic	Cases*	Controls*	All PFT*
	N = 1,873	N = 11,203	N = 13,076
Median age (IQR)	70	65	65
	(62-76)	(57-72)	(57-73)
Sex (%)			
Female	911	6,023	6,934
	(48.6)	(53.8)	(53.0)
Male	962	5,179	6,141
	(51.4)	(46.2)	(47.0)
Unknown	0	1	1
	(0.0)	(0.0)	(0.0)
Race (%)	,		,
White	1,701	9,717	11,418
	(90.8)	(86.7)	(87.3)
Black	135	1,201	1,336
	(7.2)	(10.7)	(10.2)
Other	15	162	177
	(0.8)	(1.4)	(1.4)
Unknown	22	123	145
	(1.2)	(1.1)	(1.1)
Smoking status	,		,
Ever	1,464	5,779	7,243
	(78.2)	(51.6)	(55.4)
Never	402	5,359	5,761
	(21.5)	(47.8)	(44.1)
Missing	7	65	72
3	(0.4)	(0.6)	(0.5)
Median record length	10.3	9.5	9.6
(years) (IQR)	(5.1-16.1)	(4.5-15.5)	(4.6-15.6)
Number of respiratory	188	168	171
meds** (median, IQR)	(72-385)	(66-372)	(67-374)
Top 5 phecodes by	,	, ,	`
frequency (%)			
1	Other symptoms of	Other symptoms of respiratory	Other symptoms of respiratory
	respiratory system (88.7)	system (84.5)	system (85.1)

2	Chronic airway obstruction (78.0)	Hypertension (66.7)	Hypertension (67.1)
3	Shortness of breath (71.8)	Essential hypertension (65.6)	Essential hypertension (66.3)
4	Hypertension (71.8)	Shortness of breath (64.1)	Shortness of breath (65.2)
5	Essential hypertension (70.7)	Diseases of esophagus (54.9)	Disorders of lipoid metabolism (55.2)

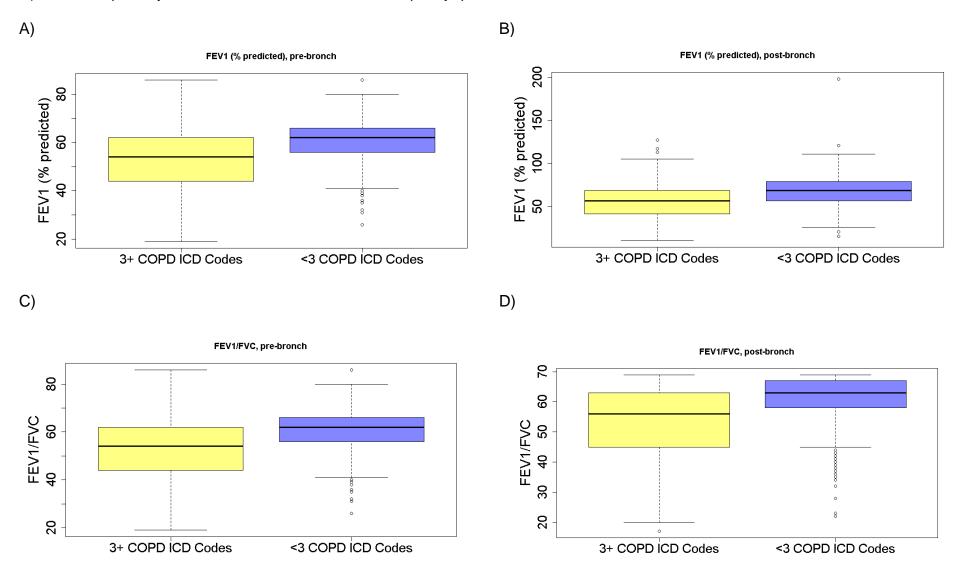
<sup>\*</sup>Cases, controls, and all PFT data sets were restricted to individuals with record lengths greater than 180 days to be comparable to those identified by the phenotyping algorithm.

\*\*Number of respiratory medications includes only mentions of medications on distinct days.

Figure S1. Phenotyping algorithm with highest PPV in the chart review validation set.



**Figure S2.** Comparison of PFT-defined cases with 3+ ICD codes for COPD and those with fewer than three ICD codes for COPD among individuals with record lengths greater than 180 days. A) Forced expiratory volume in 1 second, pre-bronchodilator. B) Forced expiratory volume in 1 second, post-bronchodilator. C) Forced expiratory volume in 1 second/forced vital capacity, pre-bronchodilator. D) Forced expiratory volume in 1 second/forced vital capacity, post-bronchodilator.



**Figure S3**. Odds ratios from genetic associations of COPD cases (3+ code) and controls in BioVU compared with associations of select variants identified in a prior GWAS of COPD by Wain et al. (32)

CHR	BP	SNP	
1	17306675	rs2284746	<del></del>
1	40035686	rs17513135	
1	92068967	rs1192404	<del></del>
1	150586971	rs6681426	<del></del>
1	218860068	rs993925	<del></del>
1	219963088	rs4328080	
2	18309132	rs62126408	<del>-</del>
2	218683154	rs2571445	<del></del>
2	229502503	rs10498230	<del></del>
2	239316560	rs61332075	<del></del>
2	239877148	rs12477314	<del>-</del>
3	67452043	rs1490265	<del>-</del>
3	127991527	rs2811415	
4	89870964	rs2045517	<b>-+</b>
4	106688904	rs10516526	<del></del>
4	106819053	rs34712979	-
5	52195033	rs1551943	<del></del>
5	131788334	rs7713065	
5	147856392	rs7715901	-
5	156810072	rs10515750	<del></del>
6	27459923	rs34864796	<del></del>
6	109270656	rs2768551	<del></del>
6	142838173	rs148274477	
9	4124377	rs7872188	<del>   </del>
9	98204792	rs16909859	-
9	139102831	rs10858246	<del>**</del>
9	139257411	rs10870202	<del>                                      </del>
10	69957350	rs7095607	<del></del>
10	78312002	rs2637254	
11	45250732	rs2863171	<del>-   -</del>
12	28689514	rs2348418	
12	57527283	rs11172113	
12	65824670	rs1494502	
12	95554771	rs113745635	
12	115201436	rs10850377	
12	115500691	rs35506	
15 40	41977690	rs72724130	
16 47	78187138	rs1079572	SD SD
17	36886828	rs11658500	Wain et al
21	35690499	rs2834440 rs11704827	
22	18450287	1511/0482/	
			0.8 1 1.2 1.4 1.6
			Odds Ratio