Association between health insurance access and chronic obstructive pulmonary disease (COPD) among adults with asthma in the United States

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Methods (XX words)

Study populations

This study includes respondents from non-institutionalized United States (US) civilian adults aged 20-80 years with asthma, in all 50 states and Washington D.C. The data for the study was extracted from the National Health and Nutrition Examination Survey (NHANES). NHANES collects data through interviews and physical examinations to study the health and nutritional status of adults and children in the US. NHANES has a publicly available database released by the National Center for Health Statistics (NCHS) [1,2].

To increase the sample size two data cycles were combined for this analysis. NHANES releases public data in two-year cycles, but the 2019-2020 cycle was not completed due to the coronavirus (COVID-19) pandemic. Therefore data collected from 2019 to March 2020 (pre-pandemic) were combined with NHANES 2017-2018 to get a nationally representative sample [2]. The 2017-2020 pre-pandemic and 2015-2016 cycles were used for this study for a total of 5.2 years.

There are 9,971 records (respondents) in the 2015-2016 cycle and 15,560 in the 2017-March 2020 cycle. Therefore the full dataset contains 25,531 records. From the full data 21,766 non-asthma patient records were excluded. Further 1,593 respondents younger than 20 years old and older than 80 years, were removed. Missing values for the exposure and outcome were defined as exclusion criteria. The final sample included respondents 2,154 (see Figure 1).

Study variables

The outcome of the study is if the participants have been diagnosed with Chronic obstructive pulmonary disease (COPD). For this information questions "Have you ever been told by a doctor or other health professional that you had chronic obstructive pulmonary disease or COPD, emphysema, or chronic bronchitis?" in the 2017-March 2020 cycle and the individual questions for emphysema, chronic bronchitis, and COPD in the 2015-2016 cycles were used. Participants who responded "Yes" to either one of the 3 diseases in the 2015-2016 data were coded as "Yes". Respondents who responded "No" to all diseases were coded as "No".

The primary exposure of interest is the respondents access to health insurance. This was obtained through the question "Are you covered by health insurance or some other kind of health care plan?".

Confounders used in this analysis were selected based on the literature. These include the number of times the respondent received healthcare over past year (healthcare use), lung cancer, smoking status, number of smokers inside the home, and diabetes. *Healthcare use* was taken from the question "During the past 12"

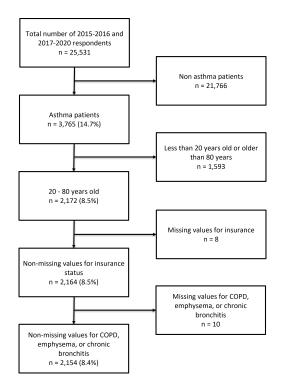


Figure 1: Flowchart of the study population and sample size for the analysis.

months, how many times have you seen a doctor or other health care professional about your health at a doctor's office, a clinic or some other place?".

Lung cancer was defined from the set of questions for "What kind of cancer?". If the respondents selected lung cancer for any one of the corresponding variables they were considered to be diagnosed with lung cancer. If participants said "No" to the question "Have you ever been told by a doctor or other health professional that you had cancer or a malignancy of any kind?", they are defined as not having lung cancer.

Smoking status was defined from questions "Do you now smoke cigarettes?" and "Have you smoked at least 100 cigarettes in your entire life?". If the respondents smoke every day or some days they are considered "current smokers". If the respondents are not smoking now but have smoked more than 100 cigarettes, they are considered as "former smokers". If they haven't smoked 100 cigarettes throughout life, they are considered "never smoked". Number of smokers inside the home was directly obtained through the question "Not counting decks, porches, or detached garages, how many people who live here smoke cigarettes, cigars, little cigars, pipes, water pipes, hookah, or any other tobacco product inside this home?".

Sex and age, in years, at the time of screening were used as other covariates in this analysis. For all above mentioned variables if the respondents answered "Refused" or "Don't know", the response was considered missing.

Statistical analyses

NHANES uses a four-stage clustered sampling design and provides sampling weights for each respondent after accounting for the said sampling design and oversampling criteria [1,2]. To account for this complex sampling design, survey weights were used. Since multiple cycles were combined sample weights should be adjusted accordingly. Guidelines specify multiplying 2015-2016 weights by $\frac{2}{5.2}$ and 2017-March 2020 weights

by $\frac{3.2}{5.2}$. All variables used in this study were collected through the household questionnaire. Therefore interview weights were used in this analysis [2].

Descriptive statistics were obtained with design adjusted medians, 1st and 3rd quartiles for numerical variables and design adjusted percentages for categorical variables. Odds ratios (OR) were obtained using a logistic regression model while adjusting for variables healthcare use, lung cancer, smoking status, number of smokers inside the home, and diabetes.

Missing values were imputed using multiple imputation (MI) based on Fully Conditional Specification (FCS) [3]. FCS was used in this data assuming that data are missing at random (MAR) [4]. Predictive mean matching was used when imputing missing numeric variables, logistic regression imputation for dichotomous variables and polytomous regression imputation for categorical data with more than 2 levels.

All analysis were conducted using R version 4.3.1 [5]. Data manipulation was done using the dplyr package [6] and complex survey design features were incorporated using the survey package [7]. Multiple imputation was performed using the mice package [3].

Results (XX words)

Variables used used in this study, grouped by levels of the outcome, are summarized in Table 1 accounting for the complex survey design. Respondents with diagnosed COPD are older (with a median age of 57) than respondents without COPD (median age 39). There are approximately 50% males and females without COPD while more females in the sample are diagnosed with COPD. The median number of healthcare visits in the past year is similar between the two response groups. The proportion of lung cancer among respondents in this sample is close to zero (2% in the COPD group and almost 0% in the non-COPD group). Conforming to what was observed in the literature, there are more current and former smokers with COPD and more respondents who have never smoked in the non-COPD group. There are more respondents with diabetes in the COPD group (24%) compared to the non-COPD group (10%).

Majority of the respondents didn't answer the number smokers inside the house, with 52% missing in the COPD group and 72% in the non-COPD group.

Table 1: Characteristics of the participants by COPD status. All proportions adjusted for the survey design.

Characteristic	Overall $(n=2,154)^1$	Has COPD $(n=508)^1$	Doesn't have COPD (n=1,646) ¹
Has insurance			
Yes	88%	91%	87%
No	12%	9%	13%
Age (years)	$43.0\ (29.0,\ 58.3)$	57.0 (44.9, 65.0)	39.0 (27.0, 55.0)
Sex		,	
Male	42%	33%	45%
Female	58%	67%	55%
No. of healthcare visits	$2.0\ (1.0,\ 4.0)$	$3.0\ (2.0,\ 6.0)$	$2.0\ (1.0,\ 4.0)$
Lung cancer			
Yes	0%	2%	0%
No	99%	97%	100%
Missing	0%	1%	0%
Smoking status			
Current smokers	21%	37%	17%
Former smokers	27%	31%	26%
Never smoked	51%	33%	56%
Missing	0%	0%	0%

No. of people who smoke in	side		
0	20%	24%	19%
1	7%	13%	6%
2	4%	8%	3%
3	1%	2%	1%
Missing	68%	53%	72%
Has diabetes			
Yes	13%	24%	10%
No	85%	72%	88%
Borderline	2%	3%	2%
Missing	0%	0%	0%

¹%; Median (IQR)

Unadjusted OR for the outcome (COPD) and exposure (insurance status) was calculated for the complete data, containing 738 records. The OR was 1.55 with a confidence interval of (1.12, 2.15). Therefore, the odds of being diagnosed with COPD is 1.6 times higher in US adults without health insurance, compared to US adults with health insurance.

OR from the logistic regression model for the complete data are shown in Table 2. Lung cancer is included in the model, despite having proportions with the disease close to zero, to avoid misspecification of the model. But the OR for lung cancer produced by the regression model is ignored. After adjusting for the confounders and other variables, the OR (1.08) shows that there is no significant difference (p-value = 0.80) in the odds of being diagnosed with COPD between US adults with or without health insurance.

Table 2: Adjusted odds ratios. All proportions adjusted for the survey design.

Characteristic	\mathbf{OR}^1	95% CI 1	p-value
Has insurance			
Yes	_	_	
No	1.08	0.57, 2.04	0.8
No. of healthcare visits	0.77	0.69, 0.86	< 0.001
Smoking status			
Current smokers	_	_	
Former smokers	2.95	1.32, 6.60	0.010
Never smoked	3.95	1.98, 7.88	< 0.001
No. of people who smoke inside	0.86	0.60, 1.24	0.4
Has diabetes			
Yes	_	_	
No	2.41	1.25, 4.62	0.010
Borderline	2.96	0.79, 11.0	0.10
Lung cancer			
Yes	_	_	
No	2,459,942	1,075,010, 5,629,078	< 0.001
Age (years)	0.95	0.93, 0.97	< 0.001
Sex			
Male	_	_	
Female	0.55	0.28, 1.06	0.072

¹OR = Odds Ratio, CI = Confidence Interval

Results in progress

The reported OR uses the complete data with no missing values. This approach might not be suitable since there is a variable with high missing values. Therefore, multiple imputation would be used in the future submission to impute missing data. A sensitivity analysis will be carried out with the "duration since last healthcare visit" replacing the number of visits, already included in the model.

PICOT

Description

- P Non-institutionalized United States civilian adults aged 20-80 years with asthma, in all 50 states and Washington D.C.
- I Individuals without access to health insurance
- C Individuals with access to health insurance
- O Chronic obstructive pulmonary disease (COPD)
- T NHANES 2015-March 2020 (pre-pandemic)

Changes since the SAP

- The population was changed with people 80 years and older removed. This was done because NHANES reports 80 and above as 80.
- The variable "close relatives with asthma" was removed following reviewer feedback.
- "Frequency of healthcare visits" (HUQ051) was considered in the base case instead of "duration since last healthcare visit", following reviewer feedback. The duration will still be used in the sensitivity analysis.
- SMQ040 ("Do you now smoke cigarettes?") was combined with SMQ020 ("Smoked at least 100 cigarettes in life") to define smoking status. The new definition is mentioned in [ssec-variables?].

References

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Appendix

Table 4: Characteristics of the participants by COPD status ignoring complex sampling des

Characteristic	Overall, $N = 2.154^{1}$	Has COPD, emphysema, ChB, $N = 508^{1}$	Doesn't have CO
Has insurance	<u> </u>		
Yes	86%	89%	
No	14%	11%	
Age (years)	47.0 (32.0, 61.0)	58.0 (46.0, 66.0)	
Sex	,	,	
Male	42%	40%	
Female	58%	60%	
No. of healthcare visits	$2.0\ (2.0,\ 4.0)$	$3.0\ (2.0,\ 6.0)$	
Lung cancer	` '		
Yes	1%	3%	
No	99%	97%	
Missing	0%	1%	
Smoking status			
Current smokers	23%	37%	
Former smokers	24%	31%	
Never smoked	53%	32%	
Missing	0%	0%	
No. of people who smoke insi	de		
0	19%	22%	
1	9%	16%	
2	5%	7%	
3	1%	2%	
Missing	66%	53%	
Has diabetes			
Yes	18%	30%	
No	80%	66%	
Borderline	3%	4%	
Missing	0%	0%	

 $^{^{1}\%}$; Median (IQR)

Table 5: Adjusted odds ratios ignoring complex sampling design.

Characteristic	\mathbf{OR}^{1}	95% CI ¹	p-value
Has insurance			
Yes			
No	0.97	0.58, 1.63	> 0.9
No. of healthcare visits	0.83	0.77, 0.90	< 0.001
Smoking status			
Current smokers			

Former smokers	2.46	1.46, 4.21	< 0.001
Never smoked	4.36	2.68, 7.29	< 0.001
No. of people who smoke inside	0.96	0.78, 1.20	0.7
Has diabetes			
Yes	_	_	
No	1.49	0.92, 2.41	0.11
Borderline	1.53	0.51, 5.03	0.5
Lung cancer			
Yes			
No	8,926,307	0.00, NA	> 0.9
Age (years)	0.95	0.94, 0.96	< 0.001
Sex			
Male		_	
Female	0.78	0.54, 1.14	0.2

 $^{^{1}\}mathrm{OR}=\mathrm{Odds}$ Ratio, CI = Confidence Interval