

#### ORIGINAL RESEARCH

# Prevalence and Associated Factors of Chronic Obstructive Pulmonary Disease Among Adults in Neno District, Malawi: A Cross-Sectional **Analytical Study**

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**Introduction:** Chronic obstructive pulmonary disease (COPD) continues to pose a global public health challenge. However, literature is scarce on the burden of COPD in Malawi. We assessed the prevalence and risk factors for COPD among adults in Neno, Malawi. Methodology: We conducted a population-based analytical cross-sectional study in Neno District between December 2021 and November 2022. Using a multi-stage sampling technique, we included 525 adults aged≥40 years. All participants underwent spirometry according to the American Thoracic Society (ATS) guidelines and were interviewed using the IMPALA questionnaire. For this study, we utilized the definition of COPD as a post-bronchodilator FEV1/FVC <0.70. We collected data using Kobo collect, exported to Microsoft Excel, and analysed using R software. We used descriptive statistics and logistic regression analysis; a p-value of <0.05 was considered statistically significant.

Results: Out of 525 participants, 510 participants were included in the final analysis. Fifty-eight percent of the participants were females (n=296), and 62.2% (n=317) were between 40 and 49 years with a median (IQR) age of 46 (40-86). For patient characteristics, 15.1% (n=77) were current smokers, and 4.1% (n=21) had a history of pulmonary tuberculosis (PTB). Cough was the most commonly reported respiratory symptom (n=249, 48.8%). The prevalence of COPD was 10.0% (n=51) and higher (15.0%) among males compared to females (6.4%). Factors significantly associated with COPD were age 60 years and above (adjusted odds ratio [aOR] = 3.27, 95% CI: 1.48–7.34, p<0.004), ever smoked (aOR = 6.17, 95% CI: 1.89–18.7, p<0.002), current smoker (aOR = 17.6, 95% CI: 8.47–38.4, p<0.001), and previous PTB (aOR = 4.42, 95% CI: 1.16–15.5, p<0.023).

Conclusion: The cross-sectional prevalence of COPD in rural Malawi is high, especially among males. Factors significantly associated were older age (60 years and above), cigarette smoking, and previous PTB. Longitudinal studies are needed to better understand disease etiology and progression in this setting.

Keywords: chronic obstructive pulmonary disease, prevalence, spirometry, associated factors, Malawi

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### Introduction

Chronic obstructive pulmonary disease (COPD), which is among the chronic respiratory diseases (CRDs), is an increasing public health concern in Africa. COPD affects 12% of the global population or 300 million people and it is the 12th most prevalent cause of years of life lost globally and the fourth leading cause of death. The current 2023 Global Initiative for Chronic Obstructive Lung Disease (GOLD) defines COPD as a heterogeneous lung condition characterized by chronic respiratory symptoms (dyspnea, cough, expectoration and/or exacerbations) due to abnormalities of the airways (bronchitis, bronchiolitis) and/or alveoli (emphysema) that cause persistent, often progressive, airflow obstruction. In low-income and middle-income countries (LMICs), non-communicable respiratory diseases are increasing among all age groups. However, COPD has not been extensively studied and described in Malawi and other African countries.

Cigarette smoking is the most important risk factor for COPD. In addition, exposure to biomass smoke, tuberculosis, severe childhood respiratory infections, malnutrition, poverty, occupational exposures, and chemical agents are associated with the increase in COPD burden in Africa. Furthermore, occupational exposures, chemical agents, and fumes have been revealed to be unrecognized risk factors for the development of COPD. Malnutrition and poverty are reported as other contributing risk factors for COPD in LMICs. However, these factors are not adequately described, and more studies are needed, especially in rural areas.

There is a wide variation in the prevalence, morbidity, and mortality of COPD between and within countries. <sup>12</sup> The results of the Burden of Obstructive Pulmonary Disease (BOLD) study conducted in Johannesburg, South Africa, in 2007 revealed a COPD prevalence of 22.2% in men and 16.7% in women had a prevalence of COPD. <sup>13</sup> Between 2008–09, a survey conducted in urban and rural Rwanda found a COPD prevalence of 9.6% in people older than 45 years. <sup>14</sup> Another study in rural Uganda in 2012 reported that the prevalence of COPD using The Global Initiative for Chronic Obstructive Lung Disease (GOLD) criterion in people older than 30 years was 16.2%. <sup>9</sup>

Malawi had an estimated population of approximatey 22,245,431 in 2023, and more than 98% use biomass fuel as their primary source of household energy<sup>15,16</sup> Diagnosing and treating non-communicable diseases (NCDs) at the district level is a challenge in Malawi due to limited health education on NCDs in the community, the number and expertise of healthcare staff, diagnostic equipment, and treatment. In 2016, a study conducted in an urban setting of the country involving individuals aged 18 years and older found a COPD prevalence of 4.3%. More recently, a prospective cohort study conducted in Chikwawa, Malawi, found a 11.2% COPD prevalence. Nevertheless, there remains a dearth of literature concerning COPD prevalence, particularly in rural areas in Malawi and other regions of sub-Saharan Africa. We conducted this study to assess the prevalence of COPD and the associated risk factors among adults in the Neno District, Malawi.

#### **Materials and Methods**

## Study Design and Setting

We conducted a population-based analytical cross-sectional study in Neno District, Malawi. Neno has an estimated population of approximately 150,211 in 2023. <sup>18</sup> The district has 12 primary health centers and two secondary hospitals (Lisungwi Community and Neno District Hospitals). Most of the Neno District population are subsistence farmers who live on less than US\$ 1.90 daily and have little access to electricity. <sup>19</sup> Neno District Health Office with support from Partners In Health (PIH) Malawi implements an integrated chronic care program that includes treatment of CRDs including COPD.

# Study Population

We included all adults aged ≥ 40 years who were Neno District residents for greater than 6 months. Other studies done in Tanzania, Russia, and Iran used ≥35 years as a cutoff point, <sup>12,20,21</sup> while a BOLD study in 2007 used ≥40 years. <sup>13</sup> In our study, the age cut-off of 40 years was chosen because COPD prevalence, morbidity, and mortality increase with age. Lung function, which reaches its peak level in young adults, starts to decline in the third and fourth decades of life. <sup>22</sup> It has been found that tobacco smoking, which is the primary risk factor for COPD, commonly begins in adolescence with often 20–25 years of exposure to tobacco smoke to induce characteristic pathophysiologic changes of COPD in human lungs. <sup>23</sup> Exclusion criteria included all adults who had a current history of mental illness, those with confirmed active pulmonary tuberculosis or symptoms suggestive of active

tuberculosis, pregnant women, and anyone with contraindication to performing spirometry, eg, history of any cerebral, ophthalmic, thoracic, or abdominal surgery in the last 1 month, or severe heart disease, recent myocardial infarction (in the last 1 month), uncontrolled angina, current respiratory infection, recent history of pneumothorax, emboli or aneurysms.

## Sample Size and Sampling Method

The required sample size for this study was calculated using a statistical formula for single population proportion<sup>24,25</sup> by considering a 95% confidence interval, 4% level of precision, a design effect of 2, 11.2% prevalence of COPD<sup>17</sup> and 10% non-response rate. The final sample size was 525 participants.

We used a multi-stage sampling technique. Neno District is administratively divided into four Traditional Authorities (TAs), namely: Chekucheku with three Group Village Heads (GVHs), Dambe with five GVHs, Mlauli with eight GVHs, and Symon with four GVHs. The GVHs formed our primary sampling unit, and we purposefully selected 15 GVHs depending on the location that they were not adjacent. The selected GVHs had a total of 87 villages, which were secondary sampling units. We purposefully selected 20 villages depending on the location of the villages such that they were not adjacent to each other. The estimated number of households per village in Neno was around 90, which totalled 1800 households. We used systematic random sampling where we skipped every three or four houses depending on the settlement pattern to arrive at the house of interview. Finally, an individual in the selected households was our final sampling unit to reach the sample size of 525 participants. A simple random technique was used to select one individual aged 40 years and above in a household with more than one eligible study participant.

#### Data Collection

We conducted face-to-face interviews using a validated International Multidisciplinary Programme to Address Lung Health and TB in Africa (IMPALA) questionnaire designed by the IMPALA network for use in sub-Saharan Africa. This is a questionnaire that was developed by lung health experts for assessment of respiratory disease and risk factors in sub-Saharan Africa. The questionnaire included socio-demographic characteristics, respiratory symptoms, exposure to biomass and wood smoke, history of PTB, and history of cigarette smoking. We hired and trained two data collectors to ensure high-quality data and collected the data between December 2021 and November 2022 using Kobo Collect. Informed consent was obtained from all participants before the interview. After the interview, the study participants' body weight, height, and body mass index (BMI) were measured. The normal BMI value that was considered was 18.5–24.9

All participants underwent spirometry using a 3-L daily calibrated and Easy On-PC Spirometer (and Medical Device Depot, Ellicott, MD, USA) after a thorough explanation of the procedure by the research assistant in Chichewa, one of the official vernacular languages spoken in Neno district. Before performing each measurement, all participants were rested for at least 5 min. The test procedure was then clearly explained to all subjects. Spirometry was performed according to the American Thoracic Society (ATS) guidelines.<sup>27</sup> Each participant used their own mouth piece to prevent cross-infection. Each spirometry manoeuvre was repeated at least three times to meet reproducibility criteria, and the highest results from the best traces were reported. We regularly checked and calibrated the spirometers prior to each measurement.

All participants with a pre-bronchodilator FEV1/FVC < 0.7 took inhaled salbutamol (400 mcg) administered via a metered-dose inhaler, and then, post-bronchodilator spirometry was performed after 20 min. Those with irreversible obstruction (FEV1/FVC <70% postbronchodilator) were classified as having COPD, while those with reversible obstruction and had symptoms suggestive of asthma were classified as asthmatic. The spirometry measurements that we used for analysis were forced expiratory volume in 1 s (FEV1), forced vital capacity (FVC) and FEV1/FVC.

In this study, we defined COPD Diagnosis based on the GOLD definition, which states that a post-bronchodilator FEV1/FVC of less than 0.70 confirms the presence of persistent airflow limitation and identifies the presence of COPD in patients with appropriate symptoms and predisposing risks. We used Quanjer (GLI), 2012 reference values. We continually checked for data completeness and consistency. We transferred spirometry data into a password-protected computer. Each test result was graded and scored as A, B, C, or D, where grade A had >3 acceptable tests with repeatability within 0.150L, B had  $\geq$ 2 acceptable tests with repeatability within 0.150L, C had  $\geq$ 2 acceptable tests with repeatability within 0.250L. We excluded low-quality spirometry results from the analysis. Three-spirometry measurements, namely FEV1, FVC, and FEV1/FVC, were considered for analysis.

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## Data Analysis

We exported data from Kobo Collect to Microsoft Excel for data cleaning and analyzed using R 4.2.2 Software. We used descriptive statistics including median and interquartile range (IQR) for continuous variables and counts and percentages for categorical variables. We used the chi-square test for proportions, Fisher's exact, and Wilcoxon rank sum tests to test for differences between sex and categorical and continuous variables. We fitted a binary logistic regression to assess the association between COPD and independent variables, including age, sex, smoking, BMI, cooking area, house ventilation status, exposure to biomass, and history of PTB. House ventilation status was defined if the house had windows allowing airflow between outside and inside. Cooking area was defined as a space for cooking for the household outside the house, inside the house or in a separate shelter. We fitted multivariable logistic regression for all variables significantly associated with COPD at the bivariate level. We considered a p-value of less than 0.05 as statistically significant.

#### Results

Of 525 participants, 510 had acceptable spirometry reports (grades A, B, or C) and were included in the final analysis. The median (IQR) age and BMI of the participants were 46 (42–57) years and 24.7kg/m<sup>2</sup>(22.2–27.6), respectively. Fifty-eight percent of participants were females (n=296). Most of the participants (62.2%, n=317) were between 40 and 49 years and 49.0% (n=250) had normal BMI with a significant difference between females and males (p<0.001). Farmers accounted for nearly half of the study (44.5%, n=227). We found that only 3.0% (n=9) of all women and 6.1% (n=13) of all men were former smokers. More men, 24.8% (n=53), were current smokers as compared to women 8.1% (n=24) with p<0.001. Similarly, of the total participants (27.8%, n=142) who drank alcohol, more men (38.8%) than women (19.9%) consumed (p<0.001). Cough was the main respiratory symptom (48.8%, n=249). Of the participants, 3.9% (n=20) had asthma and 4.1% (n=21) reported previous PTB with a statistical difference among men and women (p<0.012) (Table 1).

Table I Sociodemographic, Behavioural, and Clinical Characteristics of the Study Participants

Characteristic	Females, n = 296 <sup>a</sup>	Males, n = 214 <sup>a</sup>	Overall, N = 510 <sup>a</sup>	p-value <sup>b</sup>
Age				>0.9
40–49	182 (61.5%)	135 (63.1%)	317 (62.2%)	
50–59	52 (17.6%)	37 (17.3%)	89 (17.5%)	
≥60	62 (20.9%)	42 (19.6%)	104 (20.4%)	
Body mass index (kg/m²)				<0.001
< 18.5	6 (2.0%)	10 (4.7%)	16 (3.1%)	
18.5–24.9	110 (37.2%)	140 (65.4%)	250 (49.0%)	
≥ 25	180 (60.8%)	64 (29.9%)	244 (47.8%)	
Occupation				<0.001
Business	25 (8.4%)	30 (14.0%)	55 (10.8%)	
Employed	26 (8.8%)	44 (20.6%)	70 (13.7%)	
Farmer	144 (48.6%)	83 (38.8%)	227 (44.5%)	
Not employed	101 (34.1%)	57 (26.6%)	158 (31.0%)	
Smoking status				<0.001

(Continued)

Table I (Continued).

Characteristic	Females, n = 296 <sup>a</sup>	Males, n = 214 <sup>a</sup>	Overall, N = 510 <sup>a</sup>	p-value <sup>b</sup>
Never smoker	263 (88.9%)	148 (69.2%)	411 (80.6%)	
Former smoker	9 (3.0%)	13 (6.1%)	22 (4.3%)	
Current smoker	24 (8.1%)	53 (24.8%)	77 (15.1%)	
Years smoked for both current and former smokers (median, IQR)	24 (15–37)	24 (15–34)	24 (15–35)	0.8
Total cigarette per day for both current and former smokers (median, IQR)	2 (2–3)	4 (3–5)	3 (2–5)	<0.001
Current consumer of alcohol				<0.001
Yes	59 (19.9%)	83 (38.8%)	142 (27.8%)	
Biomass fuel exposure				0.14
Yes	294 (99.3%)	212 (99%)	506 (99.2%)	
Fuel used				0.14
Dried wood	165 (55.7%)	137 (64.0%)	302 (59.2%)	
Charcoal	116 (39.2%)	64 (29.9%)	180 (35.3%)	
Crop residue/plant products	13 (4.4%)	11 (5.1%)	24 (4.7%)	
Electricity	2 (0.7%)	2 (0.9%)	4 (0.8%)	
Cooking area				0.6
Outside on Open Air	74 (25.0%)	46 (21.5%)	120 (23.5%)	
On Veranda	13 (4.4%)	12 (5.6%)	25 (4.9%)	
In the main house, in a separate room (used as a kitchen)	42 (14.2%)	24 (11.2%)	66 (12.9%)	
In the main house in the room used for living/sleeping	5 (1.7%)	4 (1.9%)	9 (1.8%)	
Outside of the main house, in a separate building	162 (54.7%)	128 (59.8%)	290 (56.9%)	
House ventilation status				0.7
Not ventilated	160 (54.1%)	119 (55.6%)	279 (54.7%)	
Ventilated	136 (45.9%)	95 (44.4%)	231 (45.3%)	
Cough				>0.9
Yes	144 (48.6%)	105 (49.1%)	249 (48.8%)	
Phlegm				0.3
Yes	46 (15.5%)	41 (19.2%)	87 (17.1%)	
Wheezing				0.010
Yes	17 (5.7%)	26 (12.1%)	43 (8.4%)	
Asthma				0.8
Yes	11 (3.7%)	9 (4.2%)	20 (3.9%)	
Previous TB				0.6
Yes	11 (3.7%)	10 (4.7%)	21 (4.1%)	

**Notes:** \*P-values in bold are statistically significant. <sup>a</sup>n (%). <sup>b</sup>Pearson's Chi-squared test; Fisher's exact test.

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Based on post-bronchodilator assessment, the prevalence of COPD was 10.0% (n=51). The prevalence was higher in men, 15.0% (n=32), compared to women, 6.4% (n=19), and the difference was statistically significant (p=0.002). According to GOLD criteria, 51.0% (26/51) of COPD patients had moderate COPD. Severe COPD was not found among any participants. Using the mMRC dyspnoea scale, we found that more than half of patients with COPD had no dyspnoea (Table 2).

Table 2 COPD Severity and MRC Dyspnoea Score

Characteristic	Female n = 19 <sup>a</sup>	Male n = 32 <sup>a</sup>	Overall N = 51 <sup>a</sup>	p-value <sup>b</sup>
COPD Severity				0.055
Mild (FEV <sub>1</sub> ≥80% predicted)	6 (31.6%)	19 (59.4%)	25 (49.0%)	
Moderate (FEV <sub>1</sub> 50–79%)	13 (68.4%)	13 (40.6%)	26 (51.0%)	
Severe (FEV <sub>1</sub> 30–49%)	0 (0)	0 (0)	0 (0)	
MRC_dyspnoea_score				
0	16 (84.2%)	21 (65.6%)	37 (72.5%)	
1	3 (15.8%)	6 (18.8%)	9 (17.6%)	
2	0 (0.0%)	4 (12.5%)	4 (7.8%)	
3	0 (0.0%)	I (3.1%)	I (2.0%)	
4	0 (0.0%)	0 (0.0%)	0 (0.0%)	
5	0 (0.0%)	0 (0.0%)	0 (0.0%)	

Notes: an (%). Pearson's Chi-squared test.

We found no significant differences between males and females with regard to cough and phlegm (all p > 0.05), however, there was statistical difference in regard to wheezing (p < 0.010) (Figure 1).

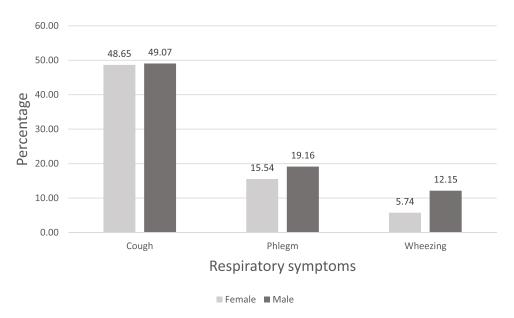


Figure I Respiratory symptoms distribution by sex.

We found that the prevalence of respiratory symptoms (cough, phlegm, and wheeze) was significantly higher in participants with COPD as compared to those without (all p<0.001) (Figure 2).

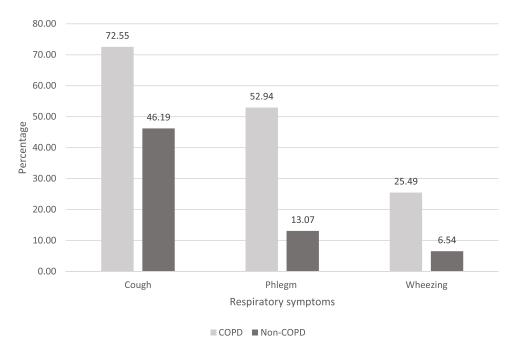


Figure 2 Prevalence of respiratory symptoms in participants with and without COPD.

### **COPD** Associated Factors

We found an association between the occurrence of COPD and age, sex, smoking status, and previous pulmonary TB. No association was observed between COPD, body mass index, cooking area, and kitchen ventilation status (p>0.05). The odds of developing COPD among people aged 60 and above were 3.19 times higher than those aged 40–49 (p<0.001, Table 3). The odds of COPD were significantly higher among males than females (OR=2.56, 95% CI [1.42–4.74], p=0.002). In comparison to those who never smoked, the odds of developing COPD were higher among those who ever smoked and are currently smoking (OR=10.6, 95% CI [3.41–30.5], p=<0.001 and OR=19.1, 95% CI [9.66–39.6], p=<0.001, respectively). For participants who had previous PTB, the odds of developing COPD increased by 3.95 times compared to those who have never suffered from TB (OR=3.95, 95% CI [1.35–10.2], p<0.007). We found that 99.2% of the participants were exposed to biomass fuel, of which 59.2% used dry wood, 35.3% used charcoal, 4.7% used crop residue/plant products, and 0.8% used electricity as the main source of cooking energy and heating (Table 1). We did not have sufficient power to assess the association between biomass exposure and COPD since 99.2% of the participants were exposed to biomass fuel.

Table 3 Factors Associated with COPD

Variable	Bivariate logistic regression			Multivariate	ate Logistic Regression		
	OR	95% CI	p-value	aOR	95% CI	p-value	
BMI							
<18.5	Reference						
18.5–24.9	0.61	0.18-2.78	0.5				
≥25	0.32	0.09-1.51	0.10				
Cooking area							
Outside	Reference						

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Table 3 (Continued).

Variable	Bivariate lo	Bivariate logistic regression			Multivariate Logistic Regression		
	OR	95% CI	p-value	aOR	95% CI	p-value	
Same building	1.45	0.51-3.86	0.5				
Separate building	1.51	0.76-3.22	0.3				
House ventilation status							
Not ventilated	Reference						
Ventilated	0.91	0.50-1.62	0.7				
Years Smoked	1.02	0.99-1.05	0.2				
Asthma							
No	Reference						
Yes	0.46	0.03-2.31	0.5				
Gender							
Female	Reference						
Male	2.56	1.42-4.74	0.002	1.37	0.67-2.80	0.4	
Age (in years)							
40-49	Reference						
50-59	1.51	0.64-3.31	0.300	1.80	0.69-4.47	0.2	
60 and above	3.19	1.65–6.14	<0.001	3.27	1.48-7.34	0.004	
Smoking status							
Never smoked	Reference						
Ever smoked	10.6	3.41-30.5	<0.001	6.17	1.89-18.7	0.002	
Current Smoker	19.1	9.66–39.6	<0.001	17.6	8.47–38.4	<0.001	
Previous TB							
No	Reference						
Yes	3.95	1.35-10.2	0.007	4.42	1.16–15.5	0.023	

Note: \*P-values in bold are statistically significant.

On multivariate logistic regression analysis, the odds of COPD were significantly higher among participants aged 60 years and above (aOR=3.27, 95% CI [1.48–7.34], p=0.004). The odds of COPD were significantly higher among eversmoked and currently smoking (aOR=6.17, 95% CI [1.89–18.7], p=0.002 and aOR=17.6, 95% CI [8.47–38.4], p<0.001, respectively). The adjusted odds of developing COPD were 4.42 times compared to those with no previous TB (aOR = 4.42, 95% CI [1.16–15.5], p<0.023), as shown in Table 3. No association was observed between COPD and exposure to biomass, cooking area, and kitchen ventilation status (p>0.05).

## Discussion

Despite several risk factors for COPD, such as biomass fuel exposure and TB infection in sub-Saharan Africa, there are few epidemiological and community-based spirometry studies to identify patients with COPD. To develop future preventative and management measures, community-based research with case finding methodologies and identification of prevalence is required to ascertain the epidemiology of COPD. 30 This cross-sectional study is aimed at determining the prevalence and associated factors among adults in Neno District, Malawi. This study provides knowledge on COPD epidemiology. We found a 10.0% COPD prevalence among adults aged 40 and above. Older age, smoking, and previous PTB were associated with COPD, with cough being the most reported respiratory symptom.

The COPD prevalence of 10.0% (6.4% in females and 15.0% in males) was based on the post-bronchodilator FEV1/ FVC < 70 criteria. This criterion was chosen despite shortfalls in the under-diagnosis of COPD among those under 40 years and over-diagnosis among those over 60 years. 31,32 The COPD prevalence we found is similar to the systematic review by Awokola et al, who found that COPD prevalence in sub-Saharan Africa ranges from 1.7% to 24.8%.<sup>33</sup> It is also similar to the overall COPD prevalence in high-income countries (HICs), which was found to be at 10.1% even though countries in LMICs have reported high COPD prevalence compared to HICs.<sup>34</sup> We found a relatively higher prevalence

than another study done in a rural district of Malawi, Chikwawa, which saw an 8.7% prevalence.<sup>35</sup> However, our prevalence was significantly different as compared to other studies done in other African countries such as Ethiopia, which found a prevalence of 17.80%, Tanzania at 17.5%, Uganda at 16.2%, South Africa at 8.5%, Nigeria at 9.2%, and Uganda at 2%.<sup>9,30,36–39</sup> The variation in the prevalence of COPD across countries could be due to differences in study population, study design, and criteria used for COPD diagnosis.<sup>33</sup> Even though there are variations in the prevalence of COPD in Africa, the use of spirometry in confirming COPD is very rare in sub-Saharan Africa, and this could imply that COPD might be underreported.<sup>40</sup> Therefore, there is a need for health facilities to have spirometry services so that patients with COPD are appropriately diagnosed.

We found that more males had COPD compared to females. Similar findings have also been reported elsewhere. <sup>34,36,41,42</sup> Our study has shown that participants 60 years and above had increased odds of developing COPD. This finding is consistent with other studies. <sup>43–45</sup> The increased exposure to risk factors and the physiological decline in respiratory function that occurs with aging may be responsible for the link between COPD and old age. <sup>45–48</sup> We also found that participants with previous pulmonary tuberculosis were four times higher at risk of developing COPD than those without. A systematic review and meta-analysis by Fan et al <sup>49</sup> and other studies have reported similar findings that a history of PTB is a risk factor for COPD. <sup>50–53</sup> Even though the exact mechanism of developing COPD in post-tuberculosis patients is not clear, a mechanism that includes bronchiectasis, bronchiolar narrowing, bronchiolitis obliterans, and accelerated emphysematous changes was proposed in a systematic review by Allwood et al. <sup>54</sup> This positive association between previous PTB history and COPD calls for integrating COPD and PTB management guidelines. It should be placed on the public health policy agenda to improve early COPD detection and treatment among people with a history of PTB.

Our study found that former and current smokers had increased odds of developing COPD compared to non-smokers, as supported by the literature and a known risk factor. 9,30,34,36,37,41 Consistent with other studies, 9,55,56 men smoked more and for longer than women in our study. However, we found a relatively low prevalence (15.1%) compared to a study from Tanzania, which found a prevalence of smoking at 25.2% (32), and also in comparison with HIC, such as Canada, reporting a prevalence of 40%. Even though smoking prevalence trends in Africa are low compared to other regions, smoking poses a greater risk for respiratory conditions such as COPD. Malawi is one of the countries whose economy depends on tobacco (55); there is a need for increased investments in smoking cessation programs, cigarette price regulation, and tobacco control laws. 59–61

We found a prevalence of asthma of 3.9%. This prevalence is similar to a cohort study of Malawian adults in rural Chikwawa conducted by Njoroge et al 2021 who found a 2.6% prevalence of asthma in 2014 and 3.7% in 2019.<sup>62</sup> Despite being a longitudinal study, it is similar to our study firstly because it was also a community-based study in a rural setting and secondly, they used spirometry to measure lung function. Similarly, a study conducted in a suburban Nigerian town found a 3.2% prevalence, but overall there is a paucity of research on asthma prevalence in sub-Saharan Africa. Whereas, asthma has been documented as a risk factor for the development of COPD, <sup>6,63</sup> we did not find any association between asthma and COPD in this study with only one participant having asthma and COPD overlap (ACO). Patients with ACO are being recognised as having a unique subset of obstructive lung disease.<sup>64</sup> A study conducted in 2017 in two districts of Malawi found one person with ACO.<sup>65</sup> Morgan et al 2019 found ACO prevalence of 3.8% in LMICs.<sup>66</sup> A systematic review and meta-analysis on global prevalence of ACO in general population found 2.0% (95% CI: 1.4–2.6%), 26.5% (95% CI: 19.5–33.6%) among patients with asthma, and 29.6% (95% CI: 19.3–39.9%) among patients with COPD.<sup>67</sup> However, we did not find this high prevalence in COPD patients in Neno District with further study warranted.

In our study, most participants (99.2%) reported having biomass fuel as their main fuel source. On the one hand, several epidemiological studies have found associations between biomass smoke exposure and COPD with odds ratios of 2 or more.<sup>68–71</sup> On the other hand, other studies, including the BOLD study and others done in Malawi, found no association between airflow obstruction and exposure to biomass.<sup>4,35,72</sup> For this study, the sample size was not powered to determine the differences between the cases and the comparison group due to the high use of biomass fuel. Further research, preferably through a prospective intervention, is required. Previous research demonstrates that biomass smokerelated COPD has different pathologies compared to cigarette-related COPD, suggesting different phenotypes from the two risk factors.<sup>73</sup> Studies have indicated biomass fuel exposure is associated with less emphysema but more air trapping than tobacco smoke exposure, implying hyperinflation with biomass smoke-related COPD.<sup>74</sup> This indicates a higher

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Residual Volume (RV), Total Lung Capacity (TLC), and RV:TLC ratio<sup>75</sup> in these patients. In our study, we could not differentiate between phenotypes as we were not able to report TLC and RV. However, we believe that there would be more biomass phenotype due to the biomass exposure in almost all participants.

Our study had several limitations. Firstly, since this was a cross-sectional study, it could not establish a cause-effect relationship. Secondly, our study was conducted in a particular setting of Malawi. As such, the results might not be generalised to the entire country. However, over 80% of Malawians live in rural areas like Neno District and use biomass fuel. This enhances generalizability for similar settings. Thirdly, this study was done during the COVID-19 pandemic with a high stigma associated with respiratory diseases and vaccination. This may have led to some potential participants hiding from the study team despite a high number of consenting for participation in the study, leading to a biased sample. However, the findings are consistent with studies outside the COVID-19 pandemic<sup>9,30,35-39,43</sup> except for no diagnosis of severe COPD, suggesting that any bias did not significantly alter the results. Fourthly, we used a fixed FEV1: FVC ratio to define COPD, which could result in over-diagnosis among older people and under-diagnosis among young adults.

Furthermore, we know that tuberculosis is underdiagnosed in Malawi, and it is possible that underlying TB could either exacerbate COPD or be misdiagnosed as COPD in this study.<sup>77</sup> Finally, we did not find a study participant with severe COPD, which may be due to the low prevalence of smoking or sampling bias. Further studies in Malawi and Neno District are required for broader sampling techniques and comparisons of diverse populations.

Despite these limitations, our study has strengths. This was a community survey with a large sample size that had adequate power to study COPD prevalence and associated factors with using community spirometry for diagnosis. It provides important data for risk factors, the diagnosis, and management of COPD in Malawi.

#### Conclusion

We found a high prevalence of COPD in rural Malawian communities with increasing age, smoking, and previous PTB risk factors. Additional longitudinal studies are needed to better understand the disease's progression and other risk factors, such as biomass exposure. Early detection and management of COPD should be considered a public health priority. We urge the Ministry of Health and other stakeholders to promote and enforce tobacco control laws, smoking cessation initiatives, and follow-up of post-TB patients to detect and improve COPD management.

#### **Abbreviations**

COPD, chronic obstructive pulmonary disease; ATS, American Thoracic Society; PTB, pulmonary tuberculosis; CRDs, chronic respiratory diseases; LMICs, low-income and middle-income countries; BOLD, The Burden of Obstructive Pulmonary Disease; GOLD, The Global Initiative for Chronic Obstructive Lung Disease; NCDs, Non-communicable diseases; FEV1, forced expiratory volume in 1 s; FVC, forced vital capacity; PIH/APZU, Partners In Health/Abwenzi Pa za Umoyo; T/As, Traditional Authorities; GVHs, Group Village Heads; BMI, Body mass index; Interquartile range (IQR); KUHeS, Kamuzu University of Health Sciences.

## **Data Sharing Statement**

The data presented in this study are available on request from the corresponding author.

## Ethical Approval and Informed Consent

The protocol was approved by the Neno District Health Research Committee and College of Medicine Research and Ethics Committee (COMREC) (P.08/20/3110). Written informed consent was obtained from study participants. The study was conducted in accordance with the Declaration of Helsinki.

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#### **Author Contributions**

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising, or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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