Chronic obstructive pulmonary disease in sub-Saharan Africa

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SUMMARY

BACKGROUND: Chronic obstructive pulmonary disease (COPD) is the third leading cause of death worldwide and an important cause of death in sub-Saharan Africa (SSA). We conducted a systematic review and meta-analysis on the prevalence of and risk factors for COPD in SSA.

METHODS: We conducted a protocol-driven systematic literature search in MEDLINE, EMBASE, CINAHL and Global Health, supplemented by a manual search of the abstracts from thoracic conference proceedings from 2017 to 2020. We did a meta-analysis of COPD prevalence and its association with current smoking. RESULTS: We identified 831 titles, of which 27 were eligible for inclusion in the review and meta-analysis. The

population prevalence of COPD ranged from 1.7% to 24.8% (pooled prevalence: 8%, 95% CI 6–11). An increased prevalence of COPD was associated with increasing age, smoking and biomass smoke exposure. The pooled odds ratio for the effect of current smoking (vs. never smoked) on COPD was 2.20 (95% CI 1.62–2.99). CONCLUSION: COPD causes morbidity and mortality in adults in SSA. Smoking is an important risk factor for COPD in SSA, and this exposure needs to be reduced through the combined efforts of clinicians, researchers and policymakers to address this debilitating and preventable lung disease.

KEY WORDS: COPD; risk factors; sub-Saharan Africa; systematic review; meta-analysis

COPD is a progressively debilitating disease, characterised by persistent respiratory symptoms and airflow limitation due to airway abnormalities caused by significant exposure to noxious stimuli, air pollution and poor pre- and post-natal lung growth. Once regarded as a disease of high-income countries, COPD is now recognised as a major health problem in low- and middle-income countries (LMICs). It is currently the third single cause of mortality worldwide: 3.8 million deaths due to COPD were reported in 2019, with approximately 90% of these occurring in LMICs. He WHO estimates that about 251 million people have COPD worldwide and that this burden is increasing. 13–15

Smoking is the main risk factor for COPD.¹⁶ In LMICs, the use of biomass fuel has been put forward as an important risk factor, but the current evidence to support this assertion is weak.^{17–19} Other risk factors include occupational dust exposure and previous serious lung infections such as TB.^{2,7,20}

COPD is often unrecognised by patients and physicians and therefore underdiagnosed and under-

treated, especially in many SSA settings, where attention remains more focused on communicable diseases. ^{21–24} The Gambia, for example, currently lacks population COPD prevalence estimates, the closest study being the Nigerian Burden of Obstructive Lung Disease (BOLD) study, which reported a 7% prevalence among adults in 2015. ²⁵

We therefore conducted a systematic review and meta-analysis to assess the current prevalence of COPD in SSA and its risk factors. This review is intended to assist clinicians with diagnostic issues and draw the attention of policymakers to strategic interventions on risk factors to reduce the burden of COPD. The current review builds on and updates an earlier systematic review we carried out in 2012 by Finney et al.²⁶

METHODOLOGY

We followed the steps outlined in the PRISMA guidelines.²⁷ The search strategy and systematic review protocol was registered with PROSPERO

http://www.crd.york.ac.uk/PROSPERO (ID: CRD42019138198).

Data sources and search strategy

We searched MEDLINE, EMBASE, CINAHL and Global Health in a comprehensive manner using a protocol-driven search strategy (see Supplementary Data 1). We also searched clinical trial registers, reference lists from published reviews and included publications and abstracts from major thoracic medicine conference proceedings of the American Thoracic Society (ATS), the European Respiratory Society (ERS) and proceedings of the Pan African Thoracic Society (PATS) and the International Union Against Tuberculosis and Lung Diseases (The Union) from 2017 to 2020. We searched in English language and no translations were required.

Study selection and data extraction

Studies were included if they met the following criteria: 1) focus on COPD, its prevalence and risk factors, 2) physician-diagnosed COPD exacerbations, 3) all COPD severities, 4) study population included people from SSA aged ≥18 years, 5) use of spirometry for COPD diagnosis, and 6) any COPD outcome—hospitalisation, discharge or death. Studies on pneumonia in chronic airway diseases, nonhuman subjects, presumptive COPD diagnoses and pharmaceutical agents were excluded. Meeting all of criteria (1), (4) and (5) were critical for inclusion.

Studies were initially assessed for inclusion based on their titles and abstracts. Full texts were obtained for studies potentially meeting the inclusion criteria. Titles, abstracts and full texts were screened for inclusion by two independent reviewers (BA and GA) separately, with each reviewer maintaining a separate record of the data extracted. Data were extracted using a standardised data extraction form that was piloted before the study (see Supplementary Data 2). A third author (CJ) reviewed the output and adjudicated in case of disagreement until a consensus was reached. We retained studies that provided numerical estimates of COPD prevalence in SSA and had clearly defined research methodologies, especially regarding the spirometric diagnosis of COPD. The case definition for COPD had to comply with one of the following: 1) Global Initiative on Obstructive Lung Disease (GOLD) criteria,²⁸ or ATS/ ERS criteria²⁹ (see Supplementary Data 3).

Quality assessment

The methodological quality of the observational studies was assessed using the Newcastle-Ottawa scale.³⁰ A score out of a maximum of six was used for all the selected cross-sectional studies.

Data analysis

Data were entered into an MS Excel® (Microsoft,

Redmond, WA, USA) spreadsheet and then read into RStudio® (Rstudio, Boston, MA, USA).³¹ We used the *metaphor* and *metaviz* packages in R Studio® to establish COPD prevalence and the effect of current smoking on COPD in SSA. We aggregated the population-based studies separately from the occupational-based studies. A random-effects model was used to pool individual studies included in the meta-analysis and to estimate the pooled effect of current smoking (vs. never smoked) on COPD prevalence. We decided not to assess for biomass exposure, as the relevant data were missing in most studies.

Assessment of heterogeneity

Study heterogeneity was assessed using Cochrane's Q, which was quantified with the inconsistency test (I^2) using a random-effects model in the Cochrane Q statistic. The forest plots were also visually inspected for closeness of point estimates and overlapping confidence intervals (CIs). The accompanying Baujat plot showed the contribution of each study to the observed heterogeneity. Finally, meta-regression was done for identified factors in order to explore the reason for a high heterogeneity when found.

Publication bias

The risk of publication bias was tested by constructing a funnel plot for a random effects-model.³³ The extracted study information was the modelling input and a pictorial depiction of how the studies fall within the 'funnel' was the output.

RESULTS

Study selection

The search yielded a total of 831 titles: MEDLINE (368 titles), EMBASE (222 titles), CINAHL (72 titles), Global Health (136 titles) and 33 titles came for handsearching the book of abstracts for ATS, ERS, PATS and The Union conferences. After the removal of duplicates, 825 titles remained. These were screened for relevance, study location, outcomes of interest and a focus on COPD leaving only 86 abstracts. After excluding mortality studies, out of scope studies, review articles and others not in line with the set protocol, we obtained full text versions of 47 papers for further consideration. Of these, 20 publications were excluded (studies not focused on COPD in SSA, and some were review articles). Twenty-seven studies were then included in the systematic review narrative and the metaanalysis (Figure 1). These comprised 23 population-based studies and four occupational-based studies.

Study characteristics

All studies included were cross-sectional studies and

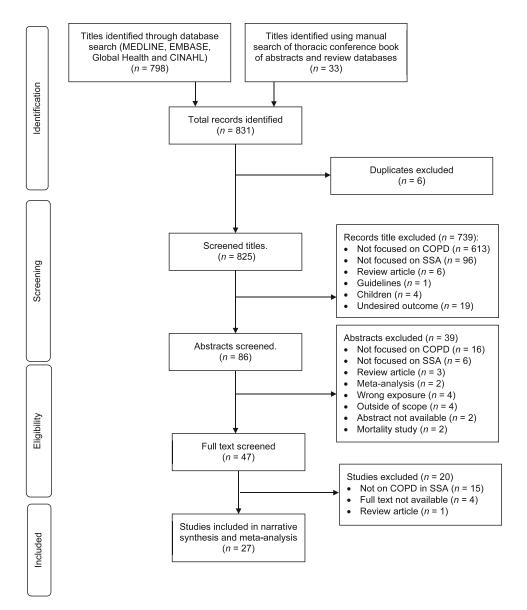


Figure 1 PRISMA flow diagram depicting the selection process. COPD = chronic obstructive pulmonary disease; SSA = sub-Saharan Africa; PRISMA = Preferred Reporting Items for Systematic reviews and Meta-Analyses.

were conducted between 1977 and 2019 (Table 1).^{20,25,34–52} In total, they represent 17,566 adults, with mean ages ranging between 38 and 80 years. Males and females were usually equally represented in the studies, except two that had industrial settings (i.e., cement and shoe factories), where only males were employed.^{50,51}

In all 27 studies, COPD diagnosis was based on spirometry,^{20,25,34–52} mainly based on the GOLD criteria (20 studies), while 6 studies used the ERS/ATS criteria, and 1 study used both criteria. The last study⁴⁸ was analysed alongside the GOLD criteria studies for the sake of computational convenience (Supplementary Data 3).

The prevalence of COPD ranged between 1.7% and 24.8% across the population-based studies

(Table 1); the pooled prevalence was estimated at 8% (95% CI 6–11) (Figure 2). Prevalence across the occupational-based studies was 1.9% to 28.2% (Table 1), with a pooled prevalence of 9% (95% CI 3–23) (Supplementary Data 4).

The respondents in the studies reported a history of exposure to smoking, biomass fuel and occupational dust; the prevalence of the exposures were documented (Table 2, Supplementary Data 5, Supplementary Data 6). A prior history of TB was also another highlighted risk factor (Table 2). In the studies where information regarding sex and smoking status was provided, significantly more men than women were current smokers (Table 2).

All but 14 of the population-based studies showed that the odds ratio (OR) of developing COPD among

Table 1 Overall characteristics of the selected studies on COPD confirmed using spirometry in sub-Saharan Africa

Author	Country	Study period	Participants n	Population	Age mean (range) years	COPD definition used	COPD prevalence %	Method quality score
Population-based st	udies							
Burney ²⁰	Cape Town, South Africa	2005	840	Urban	M: 53 F: 54	PBD FEV ₁ /FVC <lln< td=""><td>18.9</td><td>5/6</td></lln<>	18.9	5/6
Buist ³⁴	Cape Town, South Africa	2007	896	Urban	M: 52.7 F: 54.2 (40->70)	PBD FEV ₁ /FVC <0.7 FEV ₁ /FVC<0.7 and FEV ₁ ≤ 80% predicted	24.8	6/6
Burney ²⁰	Gezira, Sudan	2016	575	Rural	M: 55 F: 52	PBD FEV ₁ /FVC <lln< td=""><td>5.6</td><td>5/6</td></lln<>	5.6	5/6
Burney ²⁰	Khartoum, Sudan	2014	516	Urban	M: 55 F: 51	PBD FEV ₁ /FVC <lln< td=""><td>10.5</td><td>5/6</td></lln<>	10.5	5/6
Fullerton ³⁵	Malawi	2011	372	Mixed	41.53	Pre-BD FEV ₁ /FVC <0.7	16	4/6
Garthuru ³⁶	Nigeria	2002	410	Urban	47.8 (30–69)	Pre-BD FEV₁/FVC <0.7	9.3	4/6
Burney ²⁰	Benin	2013	545	Urban	M: 53 F: 50	PBD FEV ₁ /FVC <lln< td=""><td>7.7</td><td>5/6</td></lln<>	7.7	5/6
Magitta ³⁷	Tanzania	2016	496	Rural	51.8 (41.2-62.4)	PBD FEV ₁ /FVC < 70%	17.5	5/6
Burney ²⁰	Blantyre, Malawi	2016	401	Urban	M: 53 F: 51	PBD FEV ₁ /FVC <lln< td=""><td>8.2</td><td>5/6</td></lln<>	8.2	5/6
Burney ²⁰	Chikhwawa, Malawi	2016	432	Rural	M: 54 F: 52	PBD FEV ₁ /FVC <lln< td=""><td>14</td><td>5/6</td></lln<>	14	5/6
Musafari ³⁸	Rwanda	2009	1824	Urban and rural	38.3 (15–80)	Pre-BD FEV ₁ /FVC <lln< td=""><td>4.5</td><td>5/6</td></lln<>	4.5	5/6
Ngahane ³⁹	Cameroon	2016	337	Rural	46 (37–59)	PBD FEV ₁ /FVC <lln< td=""><td>18.4</td><td>4/6</td></lln<>	18.4	4/6
Nightingale ⁴⁰	Malawi	2016	1481	Rural	43.8 (36.0–61.6)	FEV ₁ /FVC <0.7	8.7*	6/6
North ⁴¹	Uganda	2019	565	Rural	39 ± 17	PBD FEV ₁ /FVC < LLN	2.0	6/6
Obaseki ²⁵	lle-lfe, Nigeria	2005	1169	Urban	≥40	PBD FEV ₁ /FVC < LLN	7.7	6/6
Ozoh ⁴²	Lagos, Nigeria	2012	412	Urban	53.7 (42.5–64.9)	PBD FEV ₁ /FVC < 0.7	5.3	4/6
Pefura-Yone ⁴³	Cameroon	2014	1287	Urban	34.4 (21.6–47.2)	PBD FEV ₁ /FVC < LLN and FEV ₁ /FVC <0.7	2.4	5/6
Siddharthan ⁴⁴	Uganda	2016	837	Rural	49.1	PBD FEV ₁ /FVC <i>Z</i> -score ≤−1.64	6.1	5/6
Siddharthan ⁴⁴	Uganda	2016	665	Urban	44.1	PBD FEV ₁ /FVC <i>Z</i> -score ≤−1.64	1.7	5/6
Van Gemert ⁴⁵	Uganda	2012	588	Rural	45.0 (31.3–58.7)	$FEV_1/FVC < LLN$ and $FVC < 80\%$ as cut-off	16.2	6/6
Wicht ⁴⁶ *	South Africa	1977	509	Urban	40.2 (median 20-79)	FEV ₁ /FVC <0.7	9.3	4/6
Wolderamanuel ⁴⁷	Ethiopia	2019	734	Rural	39.15 ± 9.36)	PBD FEV₁/FVC <0.7	17.8	5/6
Zoller ⁴⁸	Tanzania	2016	598		46.0 (37–57)	$FEV_1/FVC < 5^{th}$ percentile, FEV_1 < 0.7	4 (ATS), 5 (GOLD)	6/6
Occupational studie	S							
Girdler-Brown ^{49†} Mbelambela ⁵⁰	South Africa Democratic Republic Congo	2008 2016	779 379	Goldmines Cement factories	47.8 (30–69) Exposed to cement dust: 48 (37.6–58.4) Non-exposed: 51.8 (51.7–51.9)	$\begin{array}{l} {\sf FEV_1/FVC} < & 0.7 \\ {\sf PBD} \ {\sf FEV_1/FVC} < & {\sf LLN} \\ {\sf FEV_1/FVC} < & 70\% \end{array}$	9.3 28.2 (Exposed) 9.6 (non- Exposed)	4/6 4/6
Oleru & Onyekwe ⁵¹	Nigeria	1992	134	Shoe factory	33.1	FEV ₁ /FVC \leq 0.7 and FVC \geq 0.8 of	6.8	4/6
Rusibamayila ⁵²	Tanzania	2017	112	Goldmine	37.4 (31.0–43.8)	predicted FVC FEV ₁ /FVC < 0.7	1.9	5/6

^{*} Signifies prevalence of obstructive airway disease.

current smokers vs. non-smokers was at least 1.15.³⁴ Eight studies reported very high respondent exposure to biomass sources. In those with over 99% biomass-exposed population, COPD prevalence ranged between 17% and 18.5%.

Of the 15 studies that documented occupational dust exposure, five reported high proportion of its respondents being exposed; all had urban settings, and three were conducted in industrial settings: a shoe factory and gold mines (Tables 1 and 2).

Assessment of methodological quality

The funnel plot for random-effects model for all the population-based studies demonstrated good symmetry. Seventeen out of 23 studies fell within the funnel domain, with three more studies close to the funnel (Figure 3), which indicates a low risk of publication bias. In terms of their Newcastle-Ottawa quality scores, 66% of the studies had a score of at least 5 out of a total of 6, with all studies having a score of at least 4 out of 6 (Table 1).

[†] Underground and open pits.

COPD = chronic obstructive pulmonary disease; M = male; F = female; PBD = post bronchodilator; $FEV_1 = forced$ expiratory volume in 1 sec; FVC = forced vital capacity; LLN = lower limit of normal.

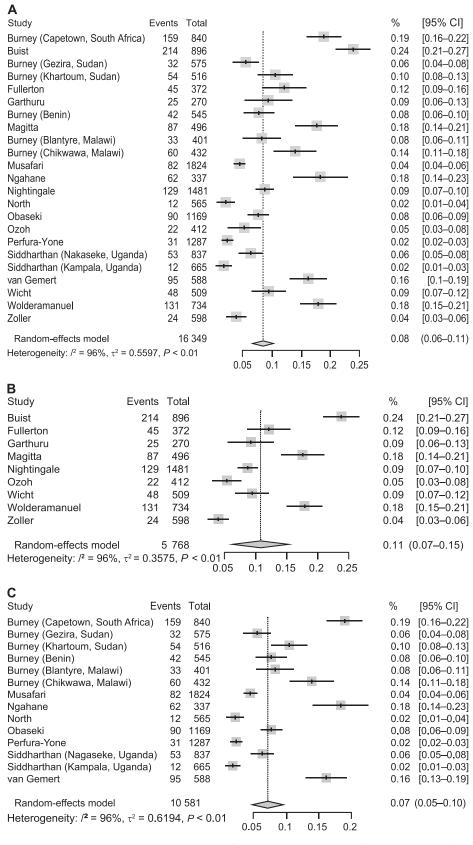


Figure 2 Forest plots showing the study-specific and pooled prevalence of (top to bottom): **A)** all 23 population-based studies, **B)** studies using a fixed ratio to define COPD, and **C)** studies using LLN to define COPD. CI = confidence interval; COPD = chronic obstructive pulmonary disease; LLN = lower limit of normal.

 Table 2
 Exposure to different risk factors for COPD in sub-Saharan Africa in the selected studies*

		Sm		Biomass		Occupational		
Study, year	Sex n (%)	Current %	Ever %	Never %	exposure %	History of TB	dust exposure	
Population-based								
Burney, 2016 ²⁰	M: 311 (37.0)	10.0	84.4	76.0	_	19	26.0	
Barriey, 2010	F: 529 (63.0)	6.4	57.9	16.0		12	87.0	
Buist, 2007 ³⁴	M: 335 (37.0)	56.9	83.0	17.0	_	19.2	61.9	
Baist, 2007	F: 561 (63.0)	40.6	59.0	41.0	_	11.9	38.8	
Burney, 2016 ²⁰	M: 298 (51.8)	8.0	47.8	68	_	0	18.0	
	F: 277 (48.2)	0.2	1.4	56			58.0	
Burney, 2014 ²⁰	M: 306 (59.3)	6.4	38.4	44	_	1	90.0	
,	F: 210 (40.7)	0.8	2.9	14			71.0	
Fullerton, 2011 ³⁵	M: 126 (37.9)	7.5	21.4	78.6	100	5.1	_	
26	F: 206 (62.1)							
Garthuru, 2002 ³⁶	M: 235 (57.2)	9.4	32.2	67.8	_	_	_	
20	F: 175 (42.8)	0.0	0.6	99.4				
Burney, 2016 ²⁰	M: 237 (43.5)	0.4	4.6	78	_	0	65	
	F: 308 (56.5)	0	0	100			96	
Magitta, 2016 ³⁷	M: 263 (53.0)	5.4	19.8	74.8	99.5	10.0	_	
201620	F: 233 (47.0)	7.5	20.6	20.0		0	75.0	
Burney, 2016 ²⁰	M: 160 (40.0) F: 241 (60.0)	7.5 0.2	30.6 2.5	38.0 51.0	_	9	75.0	
Burney, 2016 ²⁰	, ,	10.9	48.6	58.0	_	7	62.0 36.0	
Burriey, 2016	M: 221 (51.2) F: 211 (48.8)	2.1	11.3	63.0	_	/	84.0	
Musafari, 2009 ³⁸	M: 878 (48.1)	20.9	19.8	59.2	5.3		20.5	
Iviusarari, 2009	F: 946 (51.9)	12.6	11.8	75.5	5.5		20.5	
Ngahane, 2016 ³⁹	M: 168 (49.1)	9.5	—	75.5	100.0		_	
rigariarie, 2010	F: 169 (50.9)	5.5			100.0			
Nightingale, 2013 ⁴⁰	M: 637 (43.0) F: 844 (57.0)	22.2		77.8	99.2	3.2	_	
North, 2019 ⁴¹	M: 217 (38.0)	10.0	19.0	71.0	_	_	_	
Obaseki, 2016 ²⁵	F: 348 (62.0) M: 609 (40.0)	2.3	8.4	89	67.9	0.5	35.3	
Obaseki, 2010	F: 915 (60.0)	2.3	0.4	09	07.9	0.5	33.3	
Ozoh, 2012 ⁴²	M: 172 (41.7)	1.5	13.8	84.7	23.1	1.5	24.5	
02011, 2012	F: 240 (58.3)	1.5	15.0	04.7	23.1	1.5	24.5	
Pefura-Yone, 2014 ⁴³	M: 619 (48.1)	9.3	6.8	83.9	47.6	1.6	_	
	F: 668 (51.9)							
Siddhartan, 2016 ⁴⁴	M: 380 (45.4)	7.9	_	_	99.6	_	_	
	F: 457 (62.0)							
Siddhartan, 2016 ⁴⁴	M: 318 (47.8)	9.8	_	_	93.6	_	_	
	F: 347 (52.2)							
van Gemert, 2015 ⁴⁵	M: 291 (49.0)	20.7	14.8	64.5	92.9	_	_	
	F: 297 (51.0)				(indoor)			
Wicht, 1977 ⁴⁶	M: 272 (49.9)	_	_	_	_	_	_	
W-1-1	F: 273 (50.1)	0.0	2.7	00.3			00.0	
Wolderamauel, 2019 ⁴⁷	M: 421 (57.4)	9.0	2.7	88.3	_	_	90.0	
Zoller, 2016 ⁴⁸	F: 213 (42.6) M: 310 (52.0)	28.1		71.9	85.5	21.9		
Zoller, Zo ro	F: 288 (48.0)	20.1	_	71.5	(indoor)	21.9	_	
Ossupational studios	1. 200 (10.0)				(IIIdooI)			
Occupational studies Gridler-Brown, 2008 ⁴⁹	M: 624 (100)	35.0	61.0	39.0	32.2		100.0	
Mbelambela, 2016 ⁵⁰	M: 379 (100.0)	Exposed to	35 (15.5)	132 (59.2)	32.2		58.8	
Midelallibela, 2010	WI. 379 (100.0)	cement dust: 56 (25.1)	55 (15.5)	132 (33.2)	_	_	36.6	
	F: 0 (0.0)	Non-exposed:	25 (16.0)	96 (61.3)				
Oleru & Onyekwere,	M: 134 (100)	35 (22.8) 10.4	_	89	_	_	100.0	
1992 ⁵¹	M: 107 (00 E)						100.0	
Rusibamayila, 2017 ⁵²	M: 107 (99.5) F: 5 (4.5)	_	_	_	_	_	100.0	

^{*} Cells with single proportions represents a combined value for both males and females unless otherwise stated. COPD = chronic obstructive pulmonary disease; M = male; F = female.

Synthesis of results

The pooled prevalence of COPD in sub-Saharan Africa from the 23 population-based studies included was 8% (95% CI 6–11; Figure 2). The accompanying Bajaut plot and funnel plots are displayed in Figures 3 and 4, respectively. The pooled OR of current

smoking on COPD prevalence is 2.20 (95% CI 1.62–2.99; Figure 5), suggesting that current smoking is a risk for COPD and not a protective factor.

Heterogeneity assessment

There was considerable heterogeneity between stud-

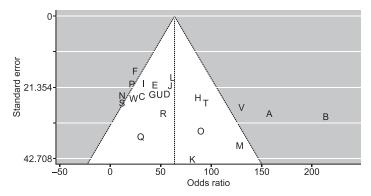


Figure 3 Funnel plot showing the risk of publication bias in the meta-analysis.

ies (Q statistic: 521.13; degrees of freedom: 11; P < 0.0001; $I^2 = 98\%$). We eyeballed our data to identify elements that can explain this significant heterogeneity. The definition of COPD (fixed ratio [0.7] vs. lower limit of normal [<5th percentile]), the population studied (rural vs. urban/semi-urban), overall criteria used in study design (GOLD/ERS-ATS), and probably, pre- vs. post-bronchodilator spirometry were possible reasons. We explored these using a sequential meta-regression of all studies on each item, hoping to observe a decrease in the unaccounted variability or I^2 value. The I^2 value did not change much (Supplementary Data 7). Despite this limitation, we proceeded with the meta-analysis to bring together all the data available on COPD prevalence in SSA.

DISCUSSION

The findings in this systematic review showed that COPD prevalence estimates in sub-Saharan African populations vary widely between studies and settings, from as low as 1.7% (rural Uganda) to as high as 24.8% (urban South Africa), with an average of 8%. African occupational studies had an average COPD prevalence of 9% (range 1.9–28.2). Overall, urban study sites tended to have higher prevalence than rural sites, and COPD prevalence increased with increasing age. As expected, current smoking was the main risk factor for COPD, with a pooled OR of 2.20. Due to the omission of risk factor reporting by many studies, we could not assess the impact of biomass fuel use on COPD prevalence. Our study is the first to utilise a meta-analysis of proportions, alongside a protocol driven systematic analysis, to describe COPD data from SSA.

There have been four previous systematic reviews of COPD in SSA. Three focused on the COPD prevalence in the continent, while one focused on spirometry availability.^{22–24,26} Mehrotra et al. (2009) reported a wide range of COPD prevalence between different SSA countries and the lack of spirometers and quality spirometry services in much of the subcontinent.²³ van Gemert et al. (2010) focused on the

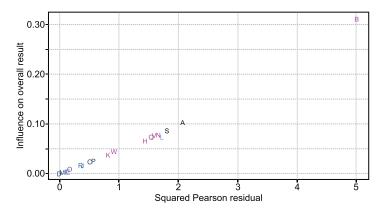


Figure 4 Baujat plot showing the contribution of individual studies to the heterogeneity of the meta-analysis shown in Figure 2. A = Burney et al. (Cape Town, South Africa); B = Buist et al.; C = Burney et al. (Gezira, Sudan); D = Burney et al. (Khartoum, Sudan); E = Fullerton et al.; F = Garthuru et al.; G = Burney et al. (Benin); H = Magitta et al.; I = Burney et al. (Blantyre, Malawi); J = Burney et al. (Chikwawa, Malawi); K = Musafari et al.; L = Ngahane et al.; M = Nightingale et al.; N = North et al.; O = Obaseki et al.; P = Ozoh et al.; Q = Perfura-Yone et al.; R = Siddharthan et al. (Kampala, Uganda [urban]); S = Siddharthan et al. (Nagaseke, Uganda [rural]); T = van Gemert et al.; U = Wicht et al.; V = Wolderamanuel et al.; W = Zoller et al.

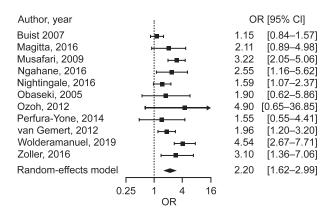


Figure 5 Forest plot displaying the meta-analysis of the effect of current smoking on COPD among respondents from studies on COPD in sub-Saharan Africa that reported current smoking. OR = odds ratio; CI = confidence interval; COPD = chronic obstructive pulmonary disease.

risk factors for asthma and COPD: current or ever smoking, prior TB, occupational exposures, indoor and outdoor air pollution, biomass fuel use, to mention a few.²² The systematic reviews by Adeloye et al. (2012)²⁴ and Finney et al. (2013)²⁶ highlighted the shortage of spirometry and lack of a noncommunicable disease healthcare strategy in most African countries as barriers to diagnosis.

Similar to the findings in the systematic reviews by Adeloye and van Gemert, we highlight a wide range in COPD prevalence estimates in SSA.^{22,24} This may be explained by the variations in the study population (community, primary care clinics, factories, etc.), differences in study settings (urban vs. rural), COPD diagnostic criteria and the effect exerted by other health determinants.

In keeping with findings from Europe, America, Asia and Australasia, we report a rising prevalence of COPD with increasing age.34,53 The mean age of participants with COPD in the Burden of Obstructive Lung Disease (BOLD) study, involving 12 cities worldwide was 53 years.^{20,34} This can be explained by several factors. In recent times, Africa has experienced a demographic shift with increasing life expectancy across many SSA nations. This has been attributed to increased standard of living and improved health-seeking behaviour.⁵⁴ The effect of this is an increase in the number of elderly people, and thus, an increase in the number of cases of COPD. Furthermore, urbanisation and westernisation of African communities have also led to increased prevalence of non-communicable diseases.⁵

Smoking is an established risk factor for developing COPD.^{55–60} There was a consensus between our review and others regarding this risk factor.^{22,24,26} Our meta-analysis showed that studies where two-fifths or more of its participants smoked tobacco, the COPD prevalence was at least 20%. Smoking rates in LMICs have increased as econo-

mies develop,^{1,60} putting the African continent on the verge of a smoking epidemic.^{24,60} Current smoking was highlighted due to availability of data on it in all the selected studies. Selective reporting in many of the retained studies made it difficult for us to conclude on other known risk factors such as biomass exposure, occupational dust exposure and a prior history of TB.

The inclusion of a meta-analysis of the prevalence of COPD in SSA and that of the effect of current smoking on COPD prevalence is a major strength of this systematic review. However, in the light of the unexplained high heterogeneity, this needs to be interpreted carefully. Second, the assessment of risk factors for COPD was another strength. Third, 13 countries were represented in this review compared to four in the previous one by Finney et al.,26 which makes our results more generalisable. Fourth, to ensure that we have good confidence in our conclusions, only studies with spirometry-based diagnoses were included. Finally, all the 27 spirometry-based studies used currently acceptable case definition criteria in diagnosing the participants: either ERS-ATS or the GOLD criteria. This is also an improvement over the systematic review conducted by Finney et al., where only six spirometry-based studies used up-to-date criteria, with eight other studies using spirometry without strictly following the standard guidelines.26

There was a high level of heterogeneity in the studies selected but we decided to execute the metaanalysis to bring together all the data available on COPD prevalence in SSA following an unsuccessful exploration of this heterogeneity using meta-regression analysis. We acknowledge this is a study limitation. Also, in a bid to capture as much spirometry-diagnosed COPD cases in our systematic review, we included three population-based studies that used pre-bronchodilatory FEV₁/FVC (forced expiratory volume in 1 sec/forced vital capacity) ratio in defining COPD.35,36,38 This may have inadvertently led to the inclusion of patients with bronchial asthma, along with COPD. The absence of complete information about other risk factors besides smoking is another limitation, especially in SSA settings, where other risk factors in addition to smoking exist. This could have been an opportunity for collecting evidence, given that current evidence supporting biomass fuel as a risk factor for developing obstructive lung disease is weak. 17-19,35

The need to identify the true burden of COPD in SSA cannot be overemphasised. This will provide stakeholders with a clearer perspective on the magnitude of this neglected health problem, which should enable them to respond appropriately. Accurate quantification of the burden of COPD using spirometry is needed now more than ever. Spirometry is currently unavailable in many parts of SSA, thus

hampering objective diagnosis as a prelude to effective therapy.^{23,61,62} Mehrotra et al. reported an availability of spirometry service and spirometer for COPD diagnosis among less than 20% of physicians contacted in 34 African countries.²³ Where these were available, spirometry results were of variable quality, based on the skills of the operator.^{40,61} In addition to providing access to spirometers, health workers using spirometry need proper training and re-training. Health policy makers need to ensure high-quality spirometry and spirometers in the existing public health service delivery channels to improve the diagnosis and management of chronic lung diseases.

In order to improve the awareness, diagnosis and management of COPD in SSA, clinical and operational research is crucial. Regarding COPD research in SSA, a lot still needs to be done to quantify the problems, proffer sustainable solutions and drive policy change successfully.61,63 Awareness campaigns on the dangers of tobacco use, tobacco use control and anti-tobacco legislation are additional vital steps to be taken for any government interested in curtailing the increase in COPD prevalence. Clinicians should also inquire about the willingness to quit smoking among smokers and be able to provide basic tobacco cessation services.²² Unfortunately, many governments in SSA have not yet given COPD the attention it requires. Concerted efforts are thus needed to ensure that this untreatable, yet potentially preventable disease is put on the health agenda of LMICs where the impact is most felt.

In a bid to control COPD, tobacco use control, anti-tobacco awareness campaigns and anti-tobacco legislations must be taken very seriously and pushed aggressively by all concerned. While the following were not shown in this review, biomass exposure, occupational dust exposure and traffic-related exposure should also be reduced in every country. Establishing and enforcing these and other similar initiatives will likely deliver lung health benefits in good time, thus bringing the menace of COPD in SSA under good control.

In conclusion, COPD causes a substantial burden of morbidity and mortality in SSA. The ongoing demographic shift in SSA suggests that this burden will increase in line with increases in life expectancy. Smoking and biomass fuel exposures are important risk factors for COPD in SSA and these exposures need to be substantially reduced through the combined efforts of clinicians, researchers and policymakers.

Acknowledgements

The authors thank P De Marie Katoto, M Lesosky, E Mukonda, W Kagima and A Derbyshire for their helpful comments during this review; the National Institute for Health Research (NIHR)

Global Health Research Unit on Lung Health and TB in Africa at the Liverpool School of Tropical Medicine, "IMPALA", for making this work possible; IMPALA (project reference 16/136/35) was funded by the NIHR using UK aid from the UK Government to support global health research; and the Aldama Foundation (London, UK) and MRC Doctoral Training Program (DTP) for sponsoring the studentship of the Principal investigator.

The views expressed in this publication are those of the author(s) and do not necessarily reflect those of the NIHR or the UK Department of Health and Social Care.

Conflicts of interest: none declared.

References

- 1 Haplin DMG, et al. It is time for the world to take COPD seriously: a statement from the GOLD board of directors Eur Respir J 2019; 54: 1900914.
- 2 Vesto J, et al. Global strategy for the diagnosis, management and prevention of Chronic Obstructive Pulmonary Disease. GOLD Executive Summary. Am J Respir Crit Care Med 2013; 187 (4): 347–365.
- 3 Meghji J, et al. Improving lung health in low-income and middle-income countries: from challenges to solutions. Lancet 2021; 397(10277): 928–940.
- 4 Halpin DMG, et al. The GOLD Summit on chronic obstructive pulmonary disease in low- and middle-income countries. Int J Tuberc Lung Dis 2019; 23(11): 1131–1141.
- 5 Dalal S, et al. Non-communicable diseases in sub-Saharan Africa: what we know now. Int J Epidemiol 2011; 40(4): 885–901
- 6 International Finance Cooperation. The business of health in Africa: partnering with the private sector to improve lives. Washington DC, USA: World Bank, 2007.
- 7 Mannino M, Buist AS. Global burden of COPD: risk factors, prevalence, and future trends. Lancet 2007; 370: 765–773
- 8 World Health Organization. World Health Statistics. Geneva, Switzerland: WHO, 2020. https://apps.who.int/iris/bitstream/ handle/10665/332070/9789240005105-eng.pdf. Accessed June 2021
- 9 Forum of International Respiratory Societies. Respiratory diseases in the world: reality of today - opportunities for tomorrow. Lausanne, Switzerland: FIRS, 2013. https://www. thoracic.org/about/global-public-health/firs/resources/firsreport-for-web.pdf Accessed May 2019.
- 10 Global Initiative for Chronic Obstructive Lung Disease. Global strategy for the diagnosis, management, and prevention of COPD. Fontana, WI, USA: GOLD, 2010. www.goldcopd.org/ guidelines-global-strategy-for-diagnosis-management.html Accessed May 2019.
- 11 World Health Organization. Geneva, Switzerland: WHO, 2015. http://www.who.int/mediacentre/factsheets/fs315/en/. Accessed June 2019.
- 12 Quedri SA, Hurst JR. The unmet burden of COPD. Global Health Epidemio Genom 2018. 3: e4.
- 13 GBD 2013 Mortality and Causes of Death Collaborators. Global, regional, and national age-sex specific all-cause and cause-specific mortality for 240 causes of death, 1990–2013: a systematic analysis for the Global Burden of Disease Study 2013. Lancet 2015; 385: 117–171.
- 14 World Health Organization. Fact sheet. Chronic obstructive pulmonary disease. Geneva, Switzerland: WHO, 2017. http://who.int/news-room/factsheets/detail/chroni-obstructive-pulmonary-disease-(copd)#:~:text=The%20Global%20Burden%20of%20Disease,in%20low%20and%20middleincome%20countries Accessed June 2021
- 15 Beaglehole R, et al. Priority actions for the non-communicable disease crisis. Lancet 2011; 377: 1438–1447.

- 16 Gupta RP, et al. Summarising published results from spirometric surveys of COPD: the problem of inconsistent definitions. Int J Tuberc Lung Dis 2014; 18: 998–1003.
- 17 André FS, et al. Airflow obstruction and use of solid fuels for cooking or heating. BOLD (Burden of Obstructive Lung Disease) results. Am J Respir Crit Care Med 197(5): 595–610.
- 18 Siddharthan T, et al. Association between household air pollution exposure and chronic obstructive pulmonary disease outcomes in 13 low- and middle-income country settings. Am J Respir Crit Care Med 2018; 197(5): 611–620.
- 19 Burney P, Amaral AFS. Air pollution and chronic airway disease: is the evidence always clear? Lancet 2019; 394(10215): 2198–2200.
- 20 Burney P, et al. Prevalence and population attributable risk for chronic airflow obstruction in a large multinational study. Am J Respir Crit Care Med 2020; 203(11): 1353–1365.
- 21 van Schayck CP, Chavannes NH. Detection of asthma and chronic obstructive pulmonary disease in primary care. Eur Respir J Suppl 2003; 39: 16s–22s
- 22 van Gemert F, et al. The impact of asthma and COPD in sub-Saharan Africa. Primary Care Respir J 2011; 20(3): 240–248.
- 23 Mehrotra A, Oluwole, AM, Gordon SB. The burden of COPD in Africa: a literature review and prospective survey of the availability of spirometry for COPD diagnosis in Africa. Trop Med Int Health 2009; 14: 840–848.
- 24 Adeloye D, et al. An estimate of the prevalence of COPD in Africa: a systematic analysis. COPD 2015; 12: 71–81.
- 25 Obaseki DO, et al. Chronic airflow obstruction in a Black African population: results of BOLD Study, Ile-Ife, Nigeria. J COPD 2016; 13: (1): 42–49.
- 26 Finney LJ, et al. Chronic obstructive pulmonary disease in sub-Saharan Africa: a systematic review. Int J Tuberc Lung Dis 2013; 17(5): 583–589.
- 27 Moher D, et al. The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. Ann Intern Med 2009; 151(4): 264–269.
- 28 Global Initiative for Chronic Obstructive Lung Disease. Global strategy for the diagnosis, management, and prevention of COPD, December 2011 (updated 2015). Fontana, WI, USA: GOLD, 2015. http://www.goldcopd.org/ Accessed June 2019.
- 29 European Respiratory Society/American Thoracic Society. COPD Guidelines. Geneva, Switzerland: ERS/ATS, 2005. www.ersnet.org
- 30 Wells GA, et al. The Newcastle-Ottawa Scale (NOS) for assessing the quality of non-randomised studies in meta-analyses. Ottawa, ON, Canada: University of Ottawa, 2015. http://www.medicine.mcgill.ca/rtamblyn/Readings%5CThe%20 Newcastle%20%20Scale%20r%20assessing%20the%20 quality%20of%20nonrandomised%20studies%20in%20 metaanalyses.pdf Accessed May 2019.
- 31 RStudio Team. RStudio: integrated development for R. Boston, MA, USA: RStudio, 2016. http://www.rstudio.com/
- 32 Higgins JPT, et al. Measuring inconsistency in Metanalysis. BMJ 2003; 327(7414): 557–560.
- 33 Deeks JJ, Macaskill P, Irwig L. The performance of tests of publication bias and other sample size effects in systematic reviews of diagnostic test accuracy was assessed. J Clin Epidemiol 2005; 58: 882–893.
- 34 Buist AS, et al. International variation in the prevalence of COPD (The BOLD Study): a population-based prevalence study. Lancet 2007; 370: 741–750.
- 35 Fullerton DG, et al. Wood smoke exposure, poverty and impaired lung function in Malawian adults. Int J Tuberc Lung Dis 2011; 15: 391–398.
- 36 Gathuru IM, et al. Differences in rates of obstructive lung disease between Africans and African Americans. Ethn Dis 2002; 12 (Suppl 3): 107–113.

- 37 Magitta NF, et al. Prevalence, risk factors and clinical correlates of COPD in a rural setting in Tanzania. Eur Respir J 2018; 51: 1700182.
- 38 Musafari S, et al. Prevalence of atopy, asthma, and COPD in an urban and a rural area of an African country. Respir Med 2011: 105; 1596–1605.
- 39 Ngahane BM, et al. Prevalence and determinants of chronic obstructive pulmonary disease in a rural area in Cameroon. Am J Respir Crit Care Med 2017; 195: A1763.
- 40 Nightingale R, et al. Noncommunicable respiratory disease in Malawi. Am J Respir Crit Care Med 2019; 199(5): 613–621.
- 41 North CM, et al. Prevalence and correlates of chronic obstructive pulmonary disease and chronic respiratory symptoms in rural southwestern Uganda: a cross-sectional, population-based study. J Glob Health 2019; 9(1): 010434.
- 42 Ozoh O, et al. The prevalence and determinants of COPD in an urban community in Lagos. Eur Respir J 2013; 42(57): 933.
- 43 Pefura-Yone EW, et al. Prevalence of obstructive lung disease in an African country using definitions from different international guidelines: a community-based cross-sectional survey BMC Res Notes 2016. 9: 124.
- 44 Siddharthan T, et al. Prevalence of chronic respiratory disease in urban and rural Uganda. Bull World Health Organ 2019; 97: 318–327.
- 45 van Gemert F, et al. Prevalence of chronic obstructive pulmonary disease and associated risk factors in Uganda (FRESH AIR Uganda): a prospective cross-sectional observational study. Lancet Glob Health 2015; 3: e44–51.
- 46 Wicht C L, de Kock M A, van Wyk Kotze T J. An epidemiological study of the diffuse obstructive pulmonary syndrome. S Afr Med J 1977; 52 (Suppl): 1–15.
- 47 Woldeamanuel GG, Mingude AB, Geta TG. Prevalence of chronic obstructive pulmonary disease (COPD) and its associated factors among adults in Abeshge District, Ethiopia: a cross sectional study. BMC Pulm Med 2019; 19(1): 181.
- 48 Zoller T, et al Chronic airflow obstruction in Tanzania a cross-sectional study. BMC Pulm Med 2018; 18: 11.
- 49 Girdler-Brown BV, et al. The burden of silicosis, pulmonary tuberculosis and COPD among former Basotho goldminers. Am J Ind Med 2008; 51: 640–647.
- 50 Mbelambela EP, et al. Prevalence of chronic obstructive pulmonary disease (COPD) among Congolese cement workers exposed to cement dust, in Kongo Central Province. Environ Sci Pollut Res Int 2018; 25(35): 35074–35083.
- 51 Oleru UG, Onyekwere C. Exposures to polyvinyl chloride, methyl ketone and other chemicals: The pulmonary and nonpulmonary effect. Int Arch Occup Environ Health 1992; 63: 503–507.
- 52 Rusibamayila M, Meshi E, Mamuya S. Respiratory impairment and personal respirable dust exposure among the underground and open cast gold miners in Tanzania. Ann Global Health 2018; 84(3): 419–428.
- 53 Chan KY, et al. How big is the 'next big thing'? Estimating the burden of noncommunicable diseases in low- and middle-income countries. J Glob Health 2012; 2(2): 20101.
- 54 Mufuda J, et al. Emerging non-communicable disease epidemic in Africa: Preventive measures from the WHO Regional office for Africa. Ethn Dis Spring 2006; 16(2): 521–526.
- 55 Ayo Yusuf OA, Reddy PS, van den Borne BW. Association of snuff use with chronic bronchitis among South African women: implications for tobacco harm reduction. Tob Control 2008; 17(2): 99–104.
- 56 Desalu OO. Prevalence of chronic bronchitis and tobacco smoking in some rural communities in Ekiti state, Nigeria. Niger Postgrad Med J 2011; 18(2): 91–97.
- 57 Ehrlich RI, et al. Predictors of chronic bronchitis in South African adults. Int J Tuberc Lung Dis Dis 2004; 8(3): 369–376.

- 58 Chatila WM, et al. Smoking patterns in African Americans and whites with advanced COPD. Chest 2004; 125(1): 15–21.
- 59 Erhabor GE, Kolawole OA. Chronic obstructive pulmonary disease: a ten-year review of clinical features in O.A.U.T.H.C., Ile-Ife. Niger J Med 2002; 11(3): 101–104.
- 60 Salvi SS, Barnes P. Chronic obstructive pulmonary disease in non-smokers. Lancet 2009; 374: 733–743.
- 61 Zubar L. Spirometry: coming of age in Africa. Breathe 2016; 12: 205–208.
- 62 Abdool-Gaffar MS, et al. Guideline for the management of chronic obstructive pulmonary disease, 2011 update. S Afr Med J 2011: 101()1): 63–73.
- 63 Plum C, et al. Availability of diagnostic services and essential medicines for non-communicable respiratory disease in African countries. Int J Tuberc Lung Dis 2021; 25(2): 120–125.

__ R É S U M É

CONTEXTE: La bronchopneumopathie chronique obstructive (COPD) représente la troisième cause de décès dans le monde et une cause importante de décès en Afrique subsaharienne (SSA). Nous avons réalisé une revue systématique et une méta-analyse de la prévalence et des facteurs de risque de COPD en SSA.

MÉTHODES: Nous avons réalisé une analyse systématique de la littérature déterminée par un protocole, en utilisant les bases de données MEDLINE, EMBASE, CINAHL et Global Health, complétée d'une recherche manuelle des résumés des conférences sur les maladies respiratoires de 2017 à 2020. Nous avons réalisé une méta-analyse de la prévalence de la COPD et de son association avec le tabagisme.

RÉSULTATS: Nous avons identifié 831 titres, dont 27

étaient éligibles à l'inclusion dans la revue et la métaanalyse. La prévalence de la COPD dans la population générale variait de 1,7% à 24,8% (prévalence groupée : 8%, IC 95% 6–11). La prévalence de la COPD augmentait avec l'âge, le tabagisme et l'exposition aux fumées dégagées lors de la combustion de la biomasse. L'odds ratio groupé pour l'effet du tabagisme actuel (vs. personnes n'ayant jamais fumé) sur la COPD était de 2,20 (IC 95% 1,62–2,99).

CONCLUSION: La COPD est à l'origine d'une morbidité et d'une mortalité chez les adultes en SSA. Le tabagisme est un facteur de risque important de COPD en SSA. Cette exposition doit être réduite grâce aux efforts conjoints des cliniciens, chercheurs et responsables politiques afin de s'attaquer à cette maladie pulmonaire évitable et invalidante.