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The Burden of Cardiovascular Disease Attributable to Hypertension in Nigeria: A Modelling Study Using Summary-Level Data

ORIGINAL RESEARCH

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ABSTRACT

Background: Globally, cardiovascular disease (CVD) remains the leading cause of mortality and disability, with hypertension being the single most important modifiable risk factor. Hypertension is responsible for about 18% of global deaths from CVD, of which African regions are disproportionately affected, especially sub-Saharan Africa. This study assessed the burden of major CVD subtypes attributable to hypertension in Nigeria.

Methods: The population attributable fractions (PAF) for myocardial infarction, all strokes, ischaemic stroke and intracerebral haemorrhagic stroke attributable to hypertension in Nigeria were calculated using published results from the INTERHEART and INTERSTROKE studies and prevalence estimates of hypertension in Nigeria. PAF estimates were obtained for age, sex, and geopolitical zones.

Results: Overall, hypertension contributed to 13.2% of all myocardial infarctions and 24.6% of all strokes, including 21.6% of all ischaemic strokes and 33.1% of all intracerebral haemorrhagic strokes. Among men aged \leq 55 years, the PAF for myocardial infarction ranged from 11.7% (North-West) to 14.6% (South-East), while in older men, it spanned 9.2% (North-West) to 11.9% (South-East). Among women aged \leq 65 years, PAF varied from 18.6% (South-South) to 20.8% (South-East and North-Central), and among women aged >65 years, it ranged from 10.4% (South-South) to 12.7% (South-East).

Conclusion: Hypertension is a key contributor to the burden of CVD in Nigeria. Understanding the burden of hypertension in the Nigerian population overall and key subgroups is crucial to developing and implementing contextualised health policies to reduce the burden of CVD. Public health interventions and policies centred on hypertension will play a critical role in potentially alleviating the burden of cardiovascular diseases (CVD) in Nigeria.

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INTRODUCTION

Globally, cardiovascular disease (CVD) remains the leading cause of death, accounting for 32% of all global deaths [1]. Hypertension is the key modifiable risk factor for CVD. An estimated 1.3 billion people around the world have hypertension, with two-thirds of these persons living in low- and middle-income countries (LMICs) [2]. In Sub-Saharan Africa (SSA), hypertension is a major public health problem, with approximately more than 75 million people living with hypertension [3]. The burden of hypertension and consequent morbidity and mortality is expected to continue to increase, in part due to weak health systems and the poor availability of effective preventive strategies [4].

Nigeria, like most other SSA countries, is currently experiencing an epidemiological transition characterised by increasing urbanisation and changes in lifestyle factors, resulting in an increased burden of CVD [5, 6]. The WHO Global Action Plan on the prevention and control of non-communicable diseases (NCDs) as well as the National Multi-sectoral Action Plan for NCDs in Nigeria both put major emphasis on the burden of hypertension and CVD [7–10]. Understanding the burden of hypertension in the Nigerian population is urgently needed to raise awareness and develop policies to reduce the burden of hypertension-related CVD in Nigeria.

In this study, we aimed to obtain a representative estimate of the burden of CVDs attributable to hypertension in Nigeria overall and in key subgroups to generate data that will drive policies on preventing CVD in Nigeria [11].

METHODS

STUDY DESIGN

This was a modelling study using summary-level data. Employing this approach from systematic reviews and global studies like INTERHEART and INTERSTROKE, the study estimated the population attributable risk of hypertension across Nigeria's geopolitical zones. The design, integrating retrospective analysis and modelling, overcomes traditional observational study limitations, providing comprehensive insights into CVD attributable to hypertension. Modelling studies efficiently leverage existing data to assess population-level impacts, which is crucial for Nigeria's diverse regions. The approach accommodates temporal trends, estimating the population attributable fraction for 1995 and 2020 and offering a robust, resource-efficient method to understand hypertension's evolving impact on cardiovascular health in Nigeria.

STUDY SETTING

Nigeria is located on the western coast of Africa [12]. With around 217 million inhabitants, it is the most populous country in Africa and the most populous Black nation on earth. The country has 36 states and its Federal Capital Territory, Abuja. The 36 states are grouped into six geopolitical zones: the North Central, North East, North West, South West, South East and South South (Figure 1) [13].

PREVALENCE OF HYPERTENSION

We obtained the most recent prevalence estimates of hypertension across the six geopolitical zones in Nigeria from a previous systematic review with meta-analyses [14]. In the review, hypertension was defined as an SBP of 140 mmHg or more and a DBP of 90 mmHg or more, a self-reported history of hypertension. The review included 52 studies from the six geopolitical zones in Nigeria, including 21 studies each from urban and rural settings and 13 studies from mixed settings (rural-urban), totaling 78,949 individuals. The prevalence of hypertension was estimated for 1995 and 2020, by age and by sex.

RISK OF CHD AND STROKE ASSOCIATED WITH HYPERTENSION

The odds ratios (OR) for myocardial infarction, overall stroke, ischaemic stroke, and intracerebral haemorrhagic stroke associated with hypertension in both men and women were obtained from INTERHEART and INTERSTROKE studies [15, 16]. INTERHEART was a case-control study that included 15,152 cases with acute myocardial infarction and 14,820 controls from across 262 centres in 52

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regions worldwide; a total of 578 cases and 789 controls were from Africa [17]. INTERSTROKE was a case-control study that included 663 and 2237 cases of intracerebral haemorrhagic stroke and ischaemic stroke, respectively, and 3,000 controls from 84 centres in 22 countries. A total of 141 and 214 cases of intracerebral hemorrhagic stroke and ischemic stroke were respectively reported for Africa (Nigeria, South Africa, Sudan, and Uganda were among the African countries included in the study). A self-reported history of hypertension was used to define hypertension in both studies.

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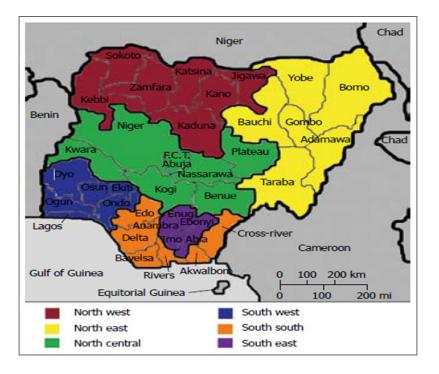


Figure 1 Map of Nigeria showing the six geopolitical zones, 36 states, and Federal Capital Territory [6].

STATISTICAL ANALYSIS

The population attributable fraction (PAF%) was estimated using the Levin formula shown below [18]:

$$PAF(\%) = \frac{P_e (RR - 1)}{[P_e (RR - 1) + 1]} \times 100\%$$

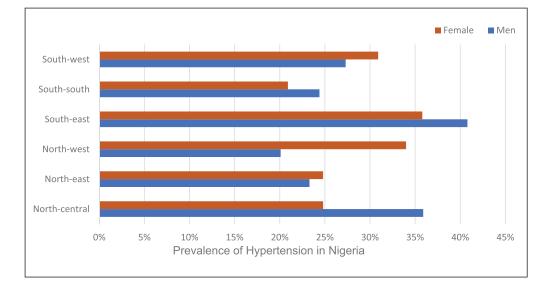
Where; P_e is the probability of exposure to the risk factor, that is, the prevalence of hypertension in Nigeria and RR is the relative risk. Estimation of relative risks from the ORs was done using the equation below, and subsequently, the PAF for the burden of CVDs in Nigeria was obtained.

$$RR = \frac{OR}{\left(1 - P_e\right) + \left(P_e * OR\right)}$$

We estimated the above PAF% for the burden of CVDs overall and across subgroups defined by age, sex, and geopolitical zone. Sex-specific PAFs% were calculated for all the outcomes. For myocardial infarction, age- and region-specific PAFs% were computed based on the information available [14]. PAF% for all the outcomes for both 1995 and 2020 were also computed. A student t-test was used to evaluate the significance of the differences in PAF across the region. The test was conducted at a two-sided 5% level of significance with no adjustment for multiple testing. Statistical analysis was carried out using R version 4.1.0 (2021-05-18).

RESULTS

The reported prevalence of hypertension in Nigeria was 30.9% and ranged across the regions from 24.7% (North West) to 33.3% (South East). Men in the South East had the highest pooled prevalence of hypertension (40.8%) compared to women (35.1%), and this was also among women in the South East. The lowest prevalence of hypertension was in the North West for men (20.1%) and in the South South for women (20.9%) (Figure 2).



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Figure 2 Prevalence of hypertension in the six geopolitical zones in Nigeria [14].

SEX-SPECIFIC PAFS FOR ALL OUTCOMES

Table 1 shows the sex-specific PAF of myocardial infarction, all stroke, ischaemic stroke and intracerebral haemorrhagic stroke in the six geopolitical zones of Nigeria. Overall, 13.2% of the burden of myocardial infarction and 24.6% of the burden of all strokes could be attributed to hypertension, with slight variation between the regions. Hypertension was estimated to have contributed about 21.6% and 33.1% of the burden of ischaemic stroke and intracerebral haemorrhagic stroke, respectively.

GEOPOLITICAL ZONES	BOTH SEXES	MEN	WOMEN		
	PAF (%)	PAF (%)	PAF (%)	ABSOLUTE DIFFERENCE IN PAF%	P-VALUE
MYOCARDIAL INFARCTIO	N				
North Central	13.3	13.6	12.2	1.5	0.78
North East	12.1	11.8	12.2	0.3	_
North West	13.3	11.0	13.5	2.5	-
South East	13.4	13.81	13.6	0.2	-
South South	12.7	12.1	11.2	0.9	-
South West	13.1	12.6	13.2	0.5	_
Overall	13.2				_
ALL STROKE					
North Central	24.6	24.6	23.9	0.7	0.56
North East	23.9	23.6	23.9	0.3	_
North West	24.6	22.7	24.7	2.0	-
South East	24.7	24.3	24.6	0.4	_
South South	24.3	23.8	23.0	0.8	_
South West	24.5	24.3	24.6	0.3	_
Overall	24.6				_
ISCHAEMIC STROKE					
North Central	21.7	21.8	20.7	1.1	0.66
North East	20.7	20.3	20.7	0.4	-
North West	21.7	19.4	21.8	2.4	-
South East	21.7	21.6	21.8	0.2	-
South South	21.2	20.6	19.6	1.0	_
South West	21.5	21.1	21.6	0.4	_
Overall	21.6				_

Table 1Sex-Specific PAF (%)of myocardial infarction, allstroke, ischaemic stroke andintracerebral haemorrgagicstroke in the six geopoliticalzones in Nigeria.

GEOPOLITICAL ZONES BOTH SEXES MEN WOMEN PAF (%) PAF (%) PAF (%) ABSOLUTE DIFFERENCE P-VALUE IN PAF% PAF (%) PAF (%) ABSOLUTE DIFFERENCE P-VALUE

INTRACEREBRAL HAE	MORRGAGIC ST	ROKE			
North Central	33.0	32.4	33.5	1.2	0.63
North East	33.5	33.5	33.5	0.0	
North West	33.0	33.2	32.7	0.5	
South East	32.8	31.3	32.4	1.1	
South South	33.4	33.5	33.3	0.2	
South West	33.2	33.4	33.1	0.3	
Overall	33.1				

AGE-AND REGION-SPECIFIC PAF OF MYOCARDIAL INFARCTION

Across regions, hypertension conferred 15.0–16.3% of the burden of myocardial infarction among younger individuals and 15.1–19.3% of the burden of myocardial infarction among older individuals. In men aged \leq 55 years, the PAF for myocardial infarction ranged from 11.7% (North West) to 14.6% (South East). Among men aged \geq 55 years, the range of PAF was 9.2% (North West) to 11.9% (South East). In women aged \leq 65 years, the PAF ranged from 18.6% (South South) to 20.8% (South East and North Central). Among women aged \geq 65 years, the PAF ranged from 10.4% (South South) to 12.7% (South East) (Table 2).

REGION	вотн			MEN			WOMEN		
	SEXES Young	Old	Absolute Difference	≤55 years	>55 years	Absolute Difference	≤65 years	>65 years	Absolute Difference
North Central	16.2	18.8	2.6	14.4	11.6	2.8	19.6	11.3	8.3
North East	15.0	15.1	0.1	12.6	9.9	2.7	19.6	11.3	8.3
North West	16.2	18.7	2.5	11.7	9.2	2.5	20.8	12.6	8.2
South East	16.3	19.3	3.0	14.6	11.9	2.7	20.8	12.7	8.1
South South	15.6	16.6	1.0	12.8	10.2	2.6	18.6	10.4	8.2
South West	16.0	17.9	1.9	13.4	10.7	2.7	20.6	12.3	8.3
Urban	16.3	19.5	3.2	13.4	10.7	2.7	20.8	12.6	8.2
Rural	15.2	15.5	0.3	13.2	10.5	2.7	19.2	10.9	8.3
Mixed	16.4	19.5	3.1	14.4	11.6	2.8	20.8	12.6	8.3
p-value			0.005			<0.0001			<0.0001

Table 2Age and sex-specificPAF% of hypertension forthe burden of myocardialInfarction.

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AGE-SPECIFIC PAF OF ALL OUTCOMES IN 1995 AND 2020

For myocardial infarction, the PAF increased from 6.2% in 1995 to 13.3% in 2020. For all strokes, the PAF increased from 15.4% in 1995 to 24.6% in 2020. For ischaemic stroke, the PAF increased from 12.4% in 1995 to 21.7% in 2020. For intracerebral haemorrhagic stroke, the PAF increased from 27.4% in 1995 to 32.9% in 2020. The observed differences in PAF values were found to be statistically significant, as evidenced by the p-values of <0.0001 for the comparisons except for intracerebral haemorrhagic stroke, where p > 0.05 (Figures 3 and 4; Table 3).

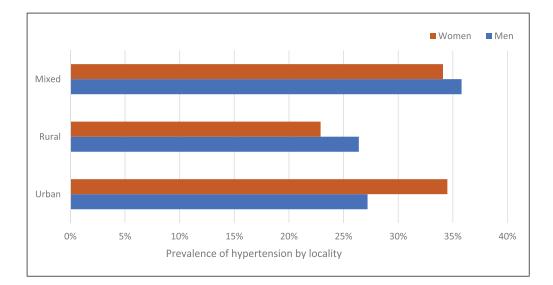




Figure 3 Prevalence of hypertension in the rural, urban and mixed settings in Nigeria [14].

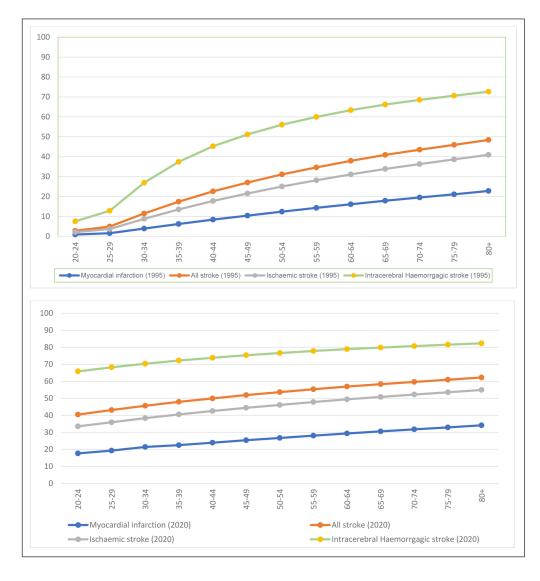


Figure 4 Age-specific population attributable risk of hypertension for the burden of myocardial infarction, all strokes and major stroke types in 1995 (top) and 2020 (bottom).

DISCUSSION

This study assessed the impact of hypertension on the prevalence of CVDs in Nigeria. The findings highlight the escalating burden of hypertension, which has emerged as the primary driver of CVDs. Hypertension plays a significant role in cardiovascular health, contributing to 13.2% of myocardial infarction burden and 24.6% of all stroke cases. It is associated with around 21.6% of ischaemic stroke cases, while having an even greater influence, at 33.1%, on the burden of intracerebral haemorrhagic strokes. The substantial increase in the PAF for myocardial infarction from 6.2% in 1995 to 13.3% in 2020 indicates a significant shift in the contribution of hypertension

	MYOCARD14	MYOCARDIAL INFARCTION	N	ALL STROKE			ISCHAEMIC STROKE	TROKE		INTRACERI	EBRAL HAEMO	INTRACEREBRAL HAEMORRGAGIC STROKE
	1995	2020		1995	2020		1995	2020		1995	2020	
AGE CATEGORY	PAF (%)	PAF(%)	ABSOLUTE DIFFERENCE	PAF(%)	PAF(%)	ABSOLUTE DIFFERENCE	PAF(%)	PAF (%)	ABSOLUTE DIFFERENCE	PAF(%)	PAF (%)	ABSOLUTE DIFFERENCE
20-24	7.0	11.9	16.7	16.8	23.6	6.8	13.7	20.4	6.7	28.8	33.5	4.7
25-29	10.3	12.4	17.7	21.9	24.1	2.2	18.6	21.0	2.4	32.8	33.5	0.7
30-34	13.8	12.9	17.5	23.7	24.5	0.8	21.2	21.4	0.2	30.2	33.3	3.1
35-39	9.7	13.3	16.2	15.5	24.6	9.1	14.1	21.6	7.5	18.8	33.0	14.2
40-44	7.0	13.5	15.5	16.9	24.6	7.7	13.8	21.8	8.0	28.9	32.6	3.7
45-49	8.3	13.7	14.9	19.1	24.5	5.4	15.8	21.8	6.0	30.8	32.1	1.2
50-54	9.5	13.8	14.3	20.8	24.3	3.5	17.4	21.7	4.3	32.1	31.5	0.6
55-59	10.4	13.8	13.7	22.0	24.0	2.0	18.7	21.5	2.8	32.9	30.8	2.1
60-64	11.3	13.8	13.2	23.0	23.6	0.6	19.7	21.2	1.5	33.3	30.0	3.3
65–69	12.0	13.6	12.6	23.7	23.1	0.6	20.5	20.8	0.3	33.5	29.2	4.3
70-74	12.5	13.4	12.2	24.2	22.6	1.6	21.0	20.3	0.7	33.5	28.3	5.2
75-79	13.0	13.2	11.8	24.5	21.9	2.6	21.4	19.8	1.6	33.3	27.4	5.9
80+	13.3	12.8	11.3	24.6	21.1	3.5	21.7	19.1	2.6	32.9	26.2	6.7
All (20+)	6.2	13.3	15.5	15.4	24.6	9.2	12.4	21.7	9.3	27.4	32.9	5.5
P-value			0.0001			0.008			0.002			0.777

Table 3 PAF of hypertension for the burden of myocardial infarction, all strokes and major stroke types (age 220 years), during 1995 and 2020.

to this specific condition over time. Similarly, the escalating PAF values for all stroke cases, rising from 12.4% to 21.7%, suggest a growing impact of hypertension on the overall stroke burden. Of notable significance is the trend observed in ischaemic stroke, where the PAF surged from 15.4% to 24.6% between 1995 and 2020. This highlights a substantial increase in the proportion of ischaemic stroke cases attributed to hypertension. Additionally, the remarkable rise in the PAF for intracerebral haemorrhagic stroke, soaring from 27.4% to 32.9%, underscores a significant augmentation in the proportion of cases linked to hypertension for this specific stroke subtype. These findings collectively underscore the evolving impact of hypertension as a key contributor to the burden of myocardial infarction and various stroke types.

The observed variations in PAFs of hypertension across the geopolitical zones in Nigeria have notable implications. The South East zone displayed the most significant relative difference in the PAF for myocardial infarction at 13.4%, highlighting the potential influence of regional factors on cardiovascular health. Likewise, the PAF values for all strokes, ischaemic stroke and intracerebral haemorrhagic stroke were notably high in the South East (24.7%), 'North Central, North West, South East, South South, South West' (21.7% each), North West (33.4%), and North East (33.5%). These findings emphasise the varying impact of hypertension on different types of strokes and cardiovascular events in different geographical regions of Nigeria, suggesting the need for tailored interventions and strategies to address the disparities and mitigate the associated health risks. This finding aligns with previous research, as hypertension has been consistently identified as one of the most prevalent risk factors for CVDs and plays a substantial role in the escalating burden of these diseases [3, 19, 20]. We observed significant variations in the burden of CVDs based on age and gender.

Notably, the PAF estimates differed among various age groups, with higher attributable fractions seen in older age groups compared to younger ones, given the increasing prevalence of hypertension with age. These findings align with earlier studies in Africa, where hypertension has consistently been ranked as the major risk factor for stroke events, particularly in relation to age. For instance, in South Africa, hypertension was identified as a major contributor to the burden of stroke, with a higher PAF reported compared to Nigeria (38.0% vs. 24.6%) [21]. In a study carried out in the Ashanti Region of Ghana, lower attributable burdens of both myocardial infarction (8.8% vs. 13.2%) and stroke (17.3% vs. 24.6%) related to hypertension were observed in Ghana compared to Nigeria.

The results of this study emphasise the significant variation in the burden of CVDs attributable to hypertension across different age and sex groups. Hypertension contributes more significantly to CVDs in men, particularly in older age groups. However, implementing population-wide measures and controls would be equally beneficial for both younger and older populations, as well as for both men and women. The study suggests that hypertension, as a prevalent risk factor, may contribute significantly to the burden of CVDs across Nigeria's six geopolitical zones. This observation underscores the importance of global efforts to prioritise the control and prevention of elevated blood pressure. This aligns with recent initiatives by the Nigerian Federal Ministry of Health, focusing on measures outlined in the National Multi-sectoral Action Plan to combat the growing cardiovascular epidemic and implementing WHO-recommended interventions ("best buys") for the prevention and control of noncommunicable diseases, including CVD. The findings support the notion that a significant burden of cardiovascular morbidity and mortality in the Nigerian population can be preventable through timely identification and prioritisation of public health interventions, including effective population-based prevention strategies centred on CVDs in resource-constrained settings, such as population-wide sodium-reduction strategies.

Aligned with Nigeria's National Multi-Sectoral Action Plan for the Prevention and Control of Noncommunicable Diseases (NCDs), the country is experiencing an epidemiological transition, leading to a dual burden of NCDs. Our study corroborates this trend, revealing a notable increase in the burden of CVD between 1995 and 2020. As seen in other nations, hypertension remains a dominant risk factor driving the surge in CVDs in Nigeria. However, it is crucial to emphasise that CVDs attributable to hypertension are largely preventable, and timely interventions can significantly improve outcomes. Our findings are in line with previous research in other LMICs, which also highlighted the rising prevalence of hypertension as a key driver of the increasing burden of CVDs in the regions [22–24]. For instance, both our study and the SIREN Stroke study showed the significant impact of hypertension on the occurrence of stroke. In our study, we observed that

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hypertension plays a crucial role in stroke incidence, with a PAF of 24.6% compared to 88.7% in the SIREN Stroke study. Furthermore, the SIREN Stroke study found a higher PAF for ischaemic strokes (77.5%) compared to haemorrhagic strokes (52.5%). In contrast, our study found a lower PAF for ischaemic strokes (21.6%) and a higher PAF for haemorrhagic strokes (33.1%). These discrepancies may reflect differences in the study populations, methodologies or other factors influencing the prevalence and impact of hypertension on stroke risk. Most notably, the SIREN Stroke study was conducted among young West Africans, and the odds ratio of stroke associated with hypertension was considerably higher than in INTERSTROKE. Overall, these findings emphasise the complex relationship between hypertension and stroke risk, highlighting the need for tailored prevention and management strategies based on population-specific characteristics and risk profiles [25].

This study has several strengths, including its extensive nationwide scope, covering all six geopolitical zones in Nigeria and ensuring comprehensive coverage. Additionally, the inclusion of studies with large population sizes enhances the statistical power and reliability of our findings. However, it is essential to acknowledge potential limitations in our study. One limitation of the modelling study design employed in this research is its reliance on existing data. The accuracy and applicability of the estimates are contingent upon the quality and representativeness of the data sources. Despite employing standardised data collection approaches for both cases and controls, inherent biases such as selection and recall bias in a case-control study could have influenced the results. However, these biases are likely to have minimal impact on the observed time trends. Another limitation is the inability to assess the PAF for fatal myocardial infarction or stroke, as participants in the INTERHEART and INTERSTROKE studies were those who survived their initial event.

CONCLUSION

The population-attributable risk fractions for various CVDs in Nigeria were found to be substantial. Notably, the burden of hypertension-attributable CVDs has significantly increased from 1995 to 2020, with notable differences observed across age groups, with older individuals bearing the highest burden. Hypertension has been identified as a primary driver of CVDs in Nigeria, presenting a significant opportunity for prevention. Public health interventions and policies focusing on hypertension are crucial to potentially reducing the burden of CVD in the country. Building upon existing interventions like the National Multi-Sectoral Action Plan, it is essential to implement additional population-based prevention strategies to address the control of CVDs effectively. In line with WHO recommendations, enhancing health promotion interventions and awareness campaigns regarding hypertension risk factors becomes imperative, with the goal of reducing hypertension prevalence by 30%.

APPENDIX

LEVEL OF RURALITY	BOTH SEXES	MEN	WOMEN	ABSOLUTE DIFFERENCE IN PAF%	P-VALUE
MYOCARDIAL INFARCTION	PAF (%)	PAF (%)	PAF (%)		
Urban	13.5	12.6	13.5	0.9	0.999
Rural	12.3	12.5	11.7	0.7	
Mixed	13.5	13.6	13.5	0.1	
ALL STROKE					
Urban	24.7	24.3	24.6	0.4	0.816
Rural	24.0	24.2	23.5	0.67	
Mixed	24.7	24.6	24.7	0.04	
ISCHAEMIC STROKE					
Urban	21.7	21.1	21.8	0.6	0.958
Rural	20.8	21.0	20.2	0.8	_
Mixed	21.7	21.8	21.8	0.0	-

Table S3Sex-specific PAF (%)of myocardial infarction, allstroke, ischaemic stroke andintracerebral haemorrgagicstroke by level of rurality.

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LEVEL OF RURALITY	BOTH SEXES	MEN	WOMEN	ABSOLUTE DIFFERENCE IN PAF%	P-VALUE
INTRACEREBRAL HAEMO	RRHAGIC STROKE				
Urban	32.7	33.4	32.6	0.8	0.731
Rural	33.5	33.5	33.5	0.0	
Mixed	32.7	32.4	32.7	0.3	-

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COMPETING INTERESTS

The authors have no competing interests to declare.

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