## ORIGINAL RESEARCH

(MMJ

## Prevalence and clinical spectrum of hypertensive retinopathy among hypertension clinic patients at Queen Elizabeth Central Hospital in Malawi

Petros Cyrus Kayange ${ }^{1,2}$, Markus Schulze Schwering ${ }^{1,3}$, Chatonda Stephen Manda ${ }^{1,2,2}$, Isaac Singini ${ }^{4}$, Vincent Verson Phillip Moyo ${ }^{5}$, Johnstone Kumwenda ${ }^{6}$

1, Ophthalmology Unit, College of Medicine, University of Malawi, Blantyre, Malawi
. Tübingen University Eye Hospital, Tübingen, Germany
5. Trinity Mission Hospital, Nsanje, Malawi
6. Department of Medicine, College of Medicine, University of Malawi, Blantyre, Malaw

## Background

## Abstract

Prevalence and spectrum of hypertensive retinopathy in the population reflects the status of hypertension control and the associated risks for cardiovascular events. We investigated the prevalence and clinical spectrum of hypertensive retinopathy among patient attending hypertension clinic at a tertiary hospital in Malawi.
Methods
Methods
This was a cross-sectional study of systematically selected patients attending hypertension clinic at Queen Elizabeth Central Hospital Patient interviews using a structured questionnaire and review of patients' medical records (health passports) were done to obtain the
following information: demographics, duration since the diagnosis of hypertension history of stroke and blod pressure measurements The presence and severity of hypertensive retinopathy was determined by dilated fundoscopy through slit lamp biomicroscopy.
Results
We recruited 104 patients. Women outnumbered men by 3:1. Women tended to be younger compared to men (mean ages 54 and 61 years respectively). Of the surveyed patients, $80 \%$ had sub-optimal blood pressure control and $75 \%$ had evidence of hypertensive retinopathy History of stroke was associated with hypertensive retinopathy
Conclusions
Hypertensive retinopathy is very common in patients attending the hypertension clinic at Queen Elizabeth Central Hospital in Blantyre Malawi. This may be a reflection of sub-optimal blood pressure control in this patient population. There is a need to identify the actua

Key words: hypertension, hypertensive retinopathy, sub-Saharan Africa, cardiovascular, stroke

## Introduction

The World Health Organization (WHO) has warned that developing countries are sitting on a time bomb of noncommunicable diseases (NCDs), and hypertension is one of them ${ }^{1}$. Data on the prevalence and control of hypertension is sparse in sub-Saharan Africa (SSA). Where data has been obtained, hypertension seems to be very common. A STEPS nationwide survey of hypertension in Malawi commissioned by the WHO found high hypertension prevalence ( $33 \%$ ) $\mathrm{NCDs}^{2}$. Data from similar surveys elsewhere shows similar high prevalence of hypertension in African communities ${ }^{3,4}$. Awareness, detection and treatment and control of hypertension in SSA are all low ${ }^{4}$. As a result, many patients present for the first time to health care providers with evidence of end organ damage. Among those that are in evidence of end organ damage. Among those that are in
care, drug stock outs and other health structural barriers may care, drug stock outs and other health structural barriers may are high rates of cardiovascular morbidity and mortality associated with hypertension in the region ${ }^{5}$.
Hypertensive retinopathy is a spectrum of retinal signs
related pathologically to retinal microvascular damage from elevated blood pressure ${ }^{6}$. It is well documented that hypertensive retinopathy is associated with cardiovascular morbidity and mortality ${ }^{7.9}$. Therefore, prevalence and spectrum of hypertensive retinopathy in the population reflects the status of hypertension control and the associated risks for cardiovascular events. In clinical management of hypertensive patients, hypertensive retinopathy can be used in cardiovascular risk assessment and can be an indication for initiating anti-hypertensive therapy, even in persons with pre-hypertension or stage one hypertension ${ }^{10}$
We conducted a hospital based cross-sectional study in Blantyre-Malawi in order to determine the prevalence and pattern of hypertensive retinopathy among patients attending a hypertension clinic at a tertiary hospital.

## Methodology

## Setting

This cross-sectional study was conducted at Queen Elizabeth Central Hospital (QECH) in Blantyre, Malawi, between July and August 2014. QECH is the largest tertiary and teaching hospital in Malawi for nurses, clinical officers
primary, secondary and tertiary. QECH serves at all these pels. It has a lage secondary level but small tertiary leve ervice and
in Blantyre.
Hypertension clinics are conducted on Tuesday and Friday morninos. Approximately 80 patients attend each clinic The drug supply is erratic at best and the commones anti-hypertensive medications include propranolol, hydrochlorothiazide and alpha-methyldopa. Angiotensin converting enzyme (ACE) inhibitors and calcium channel inhibitors are at times available. Patients are seen once every 3 months but drug supply is usually provided for only month because of rationing

## Sampling

Our minimum sample size (97) was based on the reported prevalence of $60 \%$, with a margin of $5 \%$ and this gives the study power of above $80 \%$. We used the following sample size formula:
$\mathrm{N}=\mathrm{Z}_{\alpha / 2}{ }^{2}{ }^{*} \mathrm{p} *(1-\mathrm{p}) / \mathrm{MOE}^{2}$, where $\mathrm{Z}_{\alpha / 2}$ is the critical value of the Normal distribution at $\alpha / 2$, MOE is the margin of error, p is the sample proportion, and N is the population size. A systematic sampling method was adopted where every fourth patient on the queue at hypertension clinics at QECH was invited to participate in the study. The following was the exclusion criteria: history of diabetes mellitus, retina condition other than hypertensive retinopathy, inaccessible fundus and participant's decline to participate in the study.

## Data collection

A structured interview and review of patients' medical records was done by a research assistant (a final year medical student) by use of questionnaire in order to obtain he following information. age, sex, occupation, place of residence, duration of hypertension and blood pressure measurements as recorded in patients healk passports becorded in patients' health passprts by nursing staff an recorded in patients' health passports by nursing staff on all plood pressure measurements were recorded - most recent
Fund
Fundoscopy was done in both eyes by an ophthalmologist through slit lamp biomicroscopy with hand held Volk 90 Diopter lens in order to determine the presence and severity of hypertensive retinopathy. Pupillary dilatation with Tropicamide eye drops was done before slit lamp examination. Grading of any retinopathy was done according to Mitchell-Wong simplified classification of hypertensive retinopathy ${ }^{12}$

## Data analysis

The data was entered and cleaned using Epidata version 3.1 and exported to the Statistical Package for Social Sciences (SPSS, version 16.0). Exploratory analysis was based on findings from the eye with more severe retinopathy using variables are reported as proportions while continuous variables are reported as means and standard deviation or normally distributed variables and, where appropriate, as median (IQR) for variables with a skewed distribution. Descriptive statistics were run on all baseline covariates and inferential statistics are based on the $5 \%$ significance level and $95 \%$ confidence interval.

Ethical clearance
he study protocol was approved by the College of Medicin Research Ethics Committee (P.08/13/1441). Permission to conduct the study was sought from Queen Elizabeth Centra hospital and informed written consent was obtained from all participating patients.

## Results

We invited 108participants to participate in the study out of whom 104 were enrolled. One person declined and three persons were excluded (two patients had inaccessible fundus due to dense cataracts and one patient had Uveitis).
There were 28 men ( $26.9 \%$ ) and 76 women ( $93.1 \%$ ). The demographic and clinical chatacteristics of the stud participants are shown in Table 1
Table 1: Demographic and clinical characteristics of the study participants

|  | Males | Female | All | p value |
| :---: | :---: | :---: | :---: | :---: |
| Mean age in years (SD) | 61.3(12.2) | 53.9 (12.5) | $\begin{aligned} & 55.9 \\ & (12.8) \end{aligned}$ | 0.008 |
| Median (IQR) time since diagnosis in years | $4(2-8)$ | 3(2-9) | $\begin{array}{\|l\|l} \hline 3 \\ (2-9) \end{array}$ | - |
| Time from diagnosis of hypertension |  |  |  |  |
| < 3 years | 8 (28.6\%) | 24 (32.4\%) | $\begin{array}{\|l\|} \hline 32 \\ (31.4 \%) \\ \hline \end{array}$ |  |
| 3-4 years | 7 (25.0\%) | 17 (23.0\%) | $\begin{aligned} & \hline 24 \\ & (23.5 \%) \end{aligned}$ |  |
| $\geq 5$ years | 13 (46.4\%) | 33 (44.6\%) | $\begin{aligned} & \hline \begin{array}{l} 46 \\ (45.1 \%) \end{array} \end{aligned}$ | 0.929 |
| BP <br> classification <br> \% (n) |  |  |  |  |
| $\begin{array}{\|l\|} \hline \text { Controlled } \\ \text { (<140/90 mm Hg) } \end{array}$ | 6 (21.4\%) | 12 (16.2\%) | $\begin{array}{\|l\|} \hline 18 \\ \text { (17.6\%) } \\ \hline \end{array}$ |  |
| Grade 1 (mild) | 8 (28.6\%) | 19 (25.7\%) | $\begin{array}{\|l\|} \hline 27 \\ (26.5 \%) \\ \hline \end{array}$ |  |
| Grade 2 (moderate) | 5 (17.9\%) | 20 (27.0\%) | $\begin{array}{\|l\|} \hline 25 \\ (24.5 \%) \end{array}$ |  |
| Grade 3 (severe) | 9 (32.1) | 23 (31.1\%) | $\begin{array}{\|l\|} \hline 32 \\ (31.4 \%) \end{array}$ | 0.782 |
| On treatment of hypertension | 28 (100.0\%) | 12 (96.0\%) | $\begin{array}{\|l\|} \hline 100 \\ \text { (97.1\%) } \end{array}$ | 0.283 |
| History of stroke | 8 (28.6\%) | 7 (9.3\%) | $\begin{array}{\|l\|} \hline 15 \\ (14.6 \%) \end{array}$ | 0.014 |

Women were significantly younger ( $\mathrm{p}=0.008$ ) and were less likely to have had history of stroke than men $(\mathrm{p}=0.014)$ Almost all the study participants $100(97.1 \%)$ were on anti hypertensive medications at the time of the study. The prevalence of hypertensive retinopathy is shown in Table 2.

| Grade* | Number | Percentage | $95 \% \mathrm{Cl}$ |
| :--- | :--- | :--- | :--- |
| Any retinopathy | 78 | $75.0 \%$ | $66.7-83.3 \%$ |
| Mild retinopathy | 73 | $70.2 \%$ | $61.4-79.0 \%$ |
| Moderate retinopathy | 5 | $4.8 \%$ | $0.7-8.9 \%$ |
| "None of the participants had severe retinopathy |  |  |  |

The prevalence of retinopathy was high ( $75.0 \%, 95 \%$ CI $66.7-83.3 \%$ ) in our setting. Mild hypertensive retinopathy was the most prevalent grade. There were no patients with malignant hypertensive retinopathy.
We also studied the association between the prevalence of retinopathy and factors such as sex, old age, history of troke, severity and known duration of hypertension. We found that only history of stroke was significantly associated malysis (Table 3), analysis (Table 3)
Table 3: Factors associated with prevalence of hypertensive
retinopathy

|  |  | Retinopathy |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
| $\begin{array}{l}\text { Risk factor } \\ \text { (unit of measurement) }\end{array}$ |  | Present | Absent | N | P-Value |
| Sex: n (\%) | Male | $\begin{aligned} & \hline 19 \\ & (73.08) \end{aligned}$ | $\begin{aligned} & \hline 7 \\ & (26.92) \end{aligned}$ | 26 | 1.00 |
|  | Female | $\begin{array}{\|l\|} \hline 57 \\ (73.08) \end{array}$ | $\begin{aligned} & \hline 21 \\ & (26.92) \end{aligned}$ | 78 |  |
| Age years | $\begin{array}{\|l\|} \hline \begin{array}{l} \text { Mean } \\ \text { (SD) } \end{array} \\ \hline \end{array}$ | $\begin{array}{\|l\|} \hline 56.0 \\ (10.7) \\ \hline \end{array}$ | $\begin{aligned} & \hline 57.8 \\ & (14.2) \\ & \hline \end{aligned}$ | 103 | 0.48 |
| Time since diagnosis in years | Median (IQR) | $\begin{aligned} & \hline 4 \\ & (2,8.5) \end{aligned}$ | $\begin{aligned} & 3 \\ & (2,10) \end{aligned}$ | 102 | 0.94 |
| BP classification: n (\%) | Controlled (<140/90 mm Hg) | $\begin{array}{\|l\|} \hline 13 \\ (72.2) \end{array}$ | $\begin{aligned} & \hline 5 \\ & (27.8) \end{aligned}$ | 18 | 0.68 |
|  | Grade 1 (mild) | $\begin{array}{\|l\|} \hline 21 \\ (77.8) \\ \hline \end{array}$ | $\begin{aligned} & \hline 6 \\ & \hline(22.2) \\ & \hline \end{aligned}$ | 21 |  |
|  | Grade 2 (moderate) | $\begin{array}{\|l\|} \hline 17 \\ (68.0) \\ \hline \end{array}$ | $\begin{aligned} & \hline 8 \\ & (32.0) \\ & \hline \end{aligned}$ | 25 |  |
|  | Grade 3 (severe) | $\begin{array}{\|l\|} \hline 26 \\ (81.4) \\ \hline \end{array}$ | $\begin{array}{\|l\|} \hline 6 \\ (16.6) \\ \hline \end{array}$ | 32 |  |
| Treatment of hypertension: n (\%) | Yes | $\begin{array}{\|l\|} \hline 76 \\ (76.0) \\ \hline \end{array}$ | $\begin{aligned} & \hline 24 \\ & (24.0) \\ & \hline \end{aligned}$ | 100 | 0.16 |
|  | No | $\begin{array}{\|l\|} \hline 1 \\ (33.3) \\ \hline \end{array}$ | $\begin{array}{\|l\|} \hline 2 \\ (67.7) \\ \hline \end{array}$ | 3 |  |
| History of stroke: n(\%) | Yes | $\begin{array}{\|l\|} \hline 15 \\ (100) \\ \hline \end{array}$ | 0 | 15 | 0.019 |
|  | No | $\begin{aligned} & \hline 63 \\ & (71.6) \end{aligned}$ | $\begin{aligned} & \hline 25 \\ & (26.4) \end{aligned}$ | 88 |  |

## Discussion

In this study, we found a high prevalence of uncontrolled systemic hypertension and a high prevalence of both hypertensive retinopathy ( $75.0 \%$ ) and stroke ( $14.6 \%$ ). The prevalence of stroke was higher in men than women and his difference was statistically significant. We also found opathy Only a few of our study participants had controlled opathy. Only a surprising as previous studies in Malawi have described simi lar findings ${ }^{13,14}$. The previous studies in Malawi also found
that hypertension control is a result of many factors includ ing lack of medication and essential equipment, inadequate knowledge and skills of health workers and low clinic patien attendance rates ${ }^{13,14}$. We found a very high prevalence of mild hypertensive retinopathy ( $70.2 \%$ ) among our study pat icipants. This is similar to a study done in Ghana among civi servants where the prevalence of mild grades of retinopa thy among hypertensive persons was as high as $69.4 \%{ }^{15}$. Th high prevalence of mild hypertensive retinopathy among ou study participants is not surprising because there was high prevalence of uncontrolled hypertension. We did not find severe hypertensive retinopathy. We think this was mainly because hypertensive patients with end organ damage such as renal failure and hypertensive heart diseases are seen in renal and chest clinics respectively,
Large population based studies done outside Africa have demonstrated that hypertensive retinopathy is associated with cardiovascular morbidities such as coronary cardiac disease and left ventricular hypertrophy and stroke ${ }^{10,16}$. We found in our study participants that hypertensive retinopathy was associated with history of stroke. Previous studies have demonstrated hypertensive retinopathy as an independen risk factor for incident or subclinical stroke even in patient with controlled hypertension ${ }^{10,16}$. Future studies are needed to identify the relationship between prevalent stroke and hypertensive retinopathy in Africa in persons with controlled hypertension. It would have been interesting to study the association between hypertensive retinopathy and othe cardiovascular morbidities such as coronary cardiac disease eft ventricular hypertrophy and renal dysfunction. However, his was not possible within the confines of our curren tudy. Our study had some limitations. Some diseases such s diabetic retinopathy can present with similar feature as in hypertensive retinopathy. We excluded patients with history of diabetes meliitus. However, we did not carr out laboratory tests such as fasting blood glucose to rul out the possibility of diabetic retinopathy especially in patients with moderate hypertensive retinopathy, and this may potentially have led to overestimation of prevalence of moderate hypertensive retinopathy. Secondly, we did not measure blood pressure during the study. We recorded blood pressure that was measured routinely by the hospital clinic hurses during the clinic visit. This could have potentially led to misclassification of the severity of hypertension as the blood pressure measurements may not have been adequately standardized due to busy nature of the clinic.Nevertheless, our study had some strength. Many of the previous studies on hypertensive retinopathy used Keith, Wagener, and Barker classification or its modification in grading of hypertensive retinopathy. This classification system is associated with difficulty in distinguishing the lower grades of retinopathy in clinical practice. We used a new grading system called Mitchell-Wong Simplified classification which is reliable in classifying lower grades of hypertensive retinopathy ${ }^{9,16}$. In addition, we examined both eyes of the study participants and analyzed the eye with worse hypertensive retinopathy thereby minimizing chances of missing any retinopathy.

## Conclusion

Our study has shown high prevalence of uncontrolled hy pertension, high prevalence of hypertensive retinopathy, and that hypertensive retinopathy is associated with history of stroke. Interventions aimed at control of hypertension
among clinic patient population must be intensified. Further among clinic patient population must be intensified. Furthe
es and also oo identify the association of hypertensive retinopathy with

## Acknowledgement

We are very grateful to Dr. Boston Zimba and Ms. Catheline Lunduka for taking part in the participants' recruitment process and data entry respectively.

## References

1. World Health Organization. Global health risks: mortality and burden of disease attributable to selected major risks. World Healt disease/GlobalHealthRisks report full.pdf
2. Msyamboza KP, Ngwira B, Dzowela T, Mvula C, Kathyola D, Diseases and Their Risk Factors in Malawi: Nationwide STEPS Survey. PLoS One. 2011 May 23:6(5):e20316 doi: 10.1371/journa pone. 0020316
3. Goma FM, Nzala SH, Babaniyi O, Songolo P, Zyaambo C, Ludatsikira E , et al. Prevalence of hypertension and its correlates in Med. 2011;4(1):34. doi: 10.1186/1755-7682-4-34
4. Hendriks ME, Wit FW, Roos MT, Brewster LM, Akande TM, de Beer H, et al. Hypertension in sub-Saharan Africa: cross-sectional surveys 0.1371/journal.pone. 003263
5. Cappuccio FP, Miller MA. Cardiovascular disease and hypertension sub-Saharan Africa: burden, risk and interventions. Intern Emer ed. 2016:11(3):299-305. doi: 10.1007/s11739-016-1423-9

Wong TY, Klein R, Sharrett AR, Duncan BB, Couper DJ, Tielsch JM, et al. Retinal arteriolar narrowing and risk of coronary heart disease in men and women: the Atherosclerosis Risk in Communities Study JAMA. 2002;287(9):1153-9.
. Grosso A, Veglio F, Porta M, Grignolo FM, Wong TY. Hypertensive retinopathy revisited: some answers, more questions. Br $J$ Ophthalmol 2005;89(12):1646-54. doi: 10.1136/bjo.2005.072546
10. Baker ML, Hand PJ, Wang JJ, Wong TY. Retinal signs and stroke revisiting the link between the eye and brain. Stroke. 2008;39(4):1371 9. doi: 10.1161/STROKEAHA. 107.49609
11. Ramsay LE, Williams B, Johnston GD, MacGregor GA, Poston L, Potter JF, et al. British Hypertension Society guidelines for hypertensio management 1999: summary. BMJ. 1999;319(7210):630-5. doi: https:// doi.org/10.1136/bmj.319.7210.630
12. Wong TY, Mitchell P. Hypertensive retinopathy. N Engl J Med 2004;351(22):2310-7. doi: 10.1056/NEJMra032865
13. Manjomo RC, Mwagomba B, Ade S, Ali E, Ben-Smith A, Khoman , et al. Managing and monitoring chronic non-communicable disease 2016:6(2):60-5 doi: 10.5588/pha. 16.0003
14. Wood R, Viljoen V, Van Der Merwe L, Mash R. Quality of car Malawi. Afr J Prim Health Care Fam Med 2015;7(1):1-8. http $/$ /dx do org/10.4102/phcfm.v7i1.838
15. Addo J, Smeeth L, Leon DA. Hypertensive target organ damage Ghanaian civil servants with hypertension. PLoS One. 2009-4(8):e6672. .org/10.1371/journal.pone.0006672
16. Ong Y-T, Wong TY, Klein R, Klein BE, Mitchell P, Sharrett AR, et 1. Hypertensive Retinopathy and Risk of Stroke. Hypertension. 201 62(4):706-11 doi: 10.1161/HYPERTENSIONAHA 113.01414 approach. Elsevier Health Sciences; 2011.
7. Wong TY, Klein R, Couper DJ, Cooper LS, Shahar E, Hubbard LD, et al. Retinal microvascular abnormalities and inciden troke: the Atherosclerosis Risk in Communities Study. Lancet 2001:358(9288):1134-40. doi: 10.1161/01.STR.0000195134.04355.e5

