ORIGINAL ARTICLE ASSOCIATION OF HYPERTENSIVE RETINOPATHY WITH ANGIOGRAPHIC SEVERITY OF CORONARY ARTERY DISEASE DETERMINED BY SYNTAX SCORE

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Background: Hypertension is a leading cause of morbidity among developing and developed countries. Hypertensive Retinopathy is a micro vascular complication of long standing hypertension while CAD is a macro vascular complication. The main objective of the study was to determine the association between worsening grade of hypertensive retinopathy with angiographic severity of coronary artery disease (CAD) measured by Syntax Score. Methods: This was a cross sectional study which was conducted after approval from IRB. All patients with history of hypertension, who underwent coronary angiography, were included in the study. After a detailed history and physical exam, all included patients were subjected to fundoscopy. Patients were categorized into 4 groups according to Keith et al classification of hypertensive retinopathy: No HR, Mild HR, Moderate HR and Severe HR. Patients were also categorized into three groups on the basis of angiographic severity of CAD by syntax score (SS): Mild CAD (SS<22), Moderate CAD (SS: 22-32) and Severe CAD (SS>32). Data was analysed in SPSS Version 20.0. Categorical and continuous variables were described as frequencies/percentages and Mean±SD respectively. Results: A total of 370 patients were included in the study out of which 205 were males with a mean age of 55.3±10.07 years. Mean duration of hypertension was 8.1±2.7 years with a mean SBP of 130.1±37.2 mmHg and mean DBP of 90.3±17.3 mmHg. Patients with no HR, mild HR, moderate HR and severe HR had a mean SS of 11.7±4.5, 17.1±3.9, 26.3±5.1 and 37.9±5.1 respectively. Significant association was found between HR and severity of CAD with a chi square value of 285.53 (p<0.001). PORs for worsening grade of HR with severity of CAD increased from 0.341 (p < 0.001) for mild HR to 2.33 (p < 0.001) times for severe HR. Conclusion: A higher grade of hypertensive retinopathy is significantly associated to a higher angiographic severity of CAD by syntax score.

Keywords: Coronary Artery Disease; Syntax Score; Hypertensive Retinopathy

Citation: Habib SA, Jibran MS, Khan SB, Gul AM. Association of hypertensive retinopathy with angiographic severity of coronary artery disease determined by syntax score. J Ayub Med Coll Abbottabad 2019;31(2):189–91.

INTRODUCTION

Cardiovascular disease is the leading cause of morbidity and mortality with a prevalence of 422 million and 17.9 million deaths worldwide.¹ In addition to the conventional risk factors used for risk stratification, monitoring and early treatment of patients with hypertension, hypertensive pathophysiological changes in the retina have been proposed for identifying at-risk individuals for cardiovascular disease.^{2–5} In fact, some studies have shown that HR is an independent risk factor for incident coronary artery disease in patients with hypertension.⁶ Since micro- and macro-vascular diseases share the same pathophysiological pathways, studies have shown that signs of retinopathy are independently associated with coronary artery calcification.

The purpose of this study was to investigate the relationship between worsening grades of hypertensive retinopathy and angiographic severity of CAD assessed by Syntax Score. We found that there was little data available on the subject and whether higher grades of HR were associated with higher grades of angiographic severity of CAD.

MATERIAL AND METHODS

This cross-sectional study was conducted from 1st July, 2016 to 31st of January, 2018 in cardiology department, LRH after approval from IRB. All patients with history of hypertension who underwent coronary angiography for indications set forth by international guidelines were included in the study. Patients were excluded if they had history of CLD, CKD, cardiomyopathy, congenital heart disease, malignancy, cataracts, diabetic retinopathy, panretinitis retinitis or pigmentosa. Written informed consent was taken from all the included patients for participation in study. After a detailed history and physical exam, all included patients were subjected to fundoscopy with Reisters ophthalmoscope by two residents of cardiology unit under the supervision of a consultant ophthalmologist of our hospital. Angiographic severity of CAD was assessed in cardiac cath lab after angiography. Angiography was performed by an experienced interventional cardiologist under Semins Artis 2005

machine. Angiographic severity of CAD was determined by using the Syntax Scoring System.⁸ Patients were categorized into 4 groups on the basis of grade of hypertensive retinopathy using the Keith et al classification, i.e., no HR (mild generalized arteriolar narrowing), mild HR (focal arteriolar narrowing and arteriovenous nipping), moderate HR (signs of grade 2 retinopathy plus retinal haemorrhages, exudates and cotton wool spots) and severe HR (signs of grade 3 plus papilledema). We also classified the patients on the basis of severity of CAD by syntax score into 3 groups, i.e., mild (SS<22), moderate (SS: 22-32) and severe (SS≥33) CAD. We also calculated the PORs for severity of CAD with worsening grade of HR. All data was collected on a predesigned pro forma. Categorical and continuous variables were described as frequencies/percentages and Mean±SD respectively. Data was analysed in SPSS Version 20.0.

RESULTS

A total of 370 patients were included in the study out of which 205 were males with a mean age of 55.3 ± 10.07 years. Mean duration of hypertension was 8.1 ± 2.7 years with a mean SBP of 130.1 ± 37.2 mmHg and mean DBP of 90.3 ± 17.3 mmHg. Other baseline variables are given in table-1. Patients with no HR, mild HR, moderate HR and severe HR had a mean SS of 11.7 ± 4.5 , 17.1 ± 3.9 , 26.3 ± 5.1 and 37.9 ± 5.1 respectively as shown in Table II. PORs for worsening grade of HR with severity of CAD increased from 0.341 (p<0.001) for mild HR to 2.33 (p<0.001) times for severe HR as shown in table-3.

| Table-1. Dasenne Characteristics | | |
|----------------------------------|-------------|--|
| Age (in years) | 55.3±10.07 | |
| 30–60 years | 260 (70.3%) | |
| >60 years | 110 (29.7%) | |
| Sex | | |
| Male | 205 (55.4%) | |
| Female | 165 (44.6%) | |
| History of Smoking | 85 (23%) | |
| Family History of CAD | 55 (14.8%) | |
| Duration of HTN (in years) | 8.1±2.7 | |
| CCBs | 49.2% | |
| Beta Blockers | 7% | |
| ACEIs/ARBs | 33% | |
| Combination Therapy | 10% | |
| SBP | 130.1±37.2 | |
| DBP | 90.3±17.3 | |
| | | |

Table-1: Baseline Characteristics

 Table-2: Association of HR grade with severity of CAD

| Hypertensive | CAD on Vessel Basis | | | Basis | Syntax | Severity of | |
|---|---------------------|-----|-----|-------|----------|--------------|--|
| Retinopathy | No | SVD | DVD | TVD | Score | Syntax Score | |
| No | 15 | 65 | 20 | 5 | 11.7±4.5 | Mild (<22) | |
| Mild | 15 | 60 | 30 | 0 | 17.1±3.9 | Mild (<22) | |
| Moderate | 15 | 10 | 60 | 20 | 26.3±5.1 | Mod (22-32) | |
| Severe | 0 | 0 | 5 | 50 | 37.9±5.1 | Severe (≥33) | |
| Total | 45 | 135 | 115 | 75 | | | |
| Chi square Value 285. 53 (p<0.001) | | | | | | | |
| Likelihood Ratio 279.42 (p<0.001) | | | | | | | |
| Linear-by-linear association 129.92 (p<0.001) | | | | | | | |

| Table-3: PORs for severity of CAD with worsening | |
|--|--|
| grade of HR | |

| Grade of HR | POR | Exp: (POR) | Adjusted PR | CI | <i>p</i> -value |
|----------------|-------|---------------|----------------|-----------|-----------------|
| Mild | 0.341 | 0.96 | 0.32 | 0.31-1.21 | 0.001 |
| Moderate | 0.627 | 1.86 | 1.16 | 1.21-2.52 | 0.001 |
| Severe | 0.927 | 2.52 | 2.33 | 1.81-3.52 | 0.001 |

DISCUSSION

Hypertension is a major public health concern worldwide especially in the low-income developing countries where the proportion of hypertensive individuals is increasing over the past 2 decades and the majority of hypertensive individuals in these countries are either undiagnosed or have poorly controlled blood pressure.⁹ It affects all the major organs of the body especially the cardiovascular system.^{10–12} When it damages the retinal microvascular circulation, it results in hypertensive retinopathy.¹³ The association between HR and CAD has been well established in numerous studies elsewhere. But data on severity of CAD correlation with worsening grades of HR is fairly limited.

The ARIC study found a 3-year incident CHD risk in women and a 3-year incident CVA risk in patients with micro-vascular changes in the retina irrespective of other baseline factors like DM, smoking or blood pressure.^{14,15}

We found in our study that there was an obvious association between grade of HR and severity of CAD. In our study, we found that patients with mild HR either had no CAD or mild CAD whereas patients with severe HR were more likely to have severe CAD on angiography which is a gold standard for assessing coronary artery disease. We used the Syntax Score to assess the severity of CAD which is a recent advancement in interventional cardiology that takes into account not only the number of vessels involved in CAD but also the complexity of the lesion, level of calcification and the anatomical location of the lesion thereby giving a more appropriate assessment of CAD.

CONCLUSION

A higher grade of hypertensive retinopathy is significantly associated with a worsening severity of CAD on coronary angiography assessed by Syntax Score.

AUTHORS' CONTRIBUTION

SAH & MSJ: Conceived the idea, planned the study, collected data, did the statistical analysis and drafted the manuscript. SBK & AMG: Critically supervised the whole process and did critical review of drafted manuscript

REFERENCES

- Roth GA, Johnson C, Abajobir A, Abd-Allah F, Abera SF, Abyu G, *et al.* Global, regional, and national burden of cardiovascular diseases for 10 causes, 1990 to 2015. J Am Coll Cardiol 2017;70(1):1–25.
- Wong TY, Klein R, Klein BE, Tielsch JM, Hubbard L, Nieto FJ. Retinal microvascular abnormalities and their relationship with hypertension, cardiovascular disease, and mortality. Surv Ophthalmol 2001;46(1):59–80.
- Kim GH, Youn HJ, Kang S, Choi YS, Moon JI. Relation between grade II hypertensive retinopathy and coronary artery disease in treated essential hypertensives. Clin Exp Hypertens 2010;32(7):469–73.
- Wong TY, Kamineni A, Klein R, Sharrett AR, Klein BE, Siscovick DS, *et al.* Quantitative retinal venular caliber and risk of cardiovascular disease in older persons: the cardiovascular health study. Arch Intern Med 2006;166(21):2388–94.
- Wang JJ, Liew G, Wong TY, Smith W, Klein R, Leeder SR, et al. Retinal vascular calibre and the risk of coronary heart disease-related death. Heart 2006;92(11):1583–7.
- Duncan BB, Wong TY, Tyroler HA, Davis CE, Fuchs FD. Hypertensive retinopathy and incident coronary heart disease in high risk men. Br J Ophthalmol 2002;86(9):1002–6.
- Wong TY, Cheung N, Islam FA, Klein R, Criqui MH, Cotch MF, *et al.* Relation of retinopathy to coronary artery calcification: the multi-ethnic study of atherosclerosis. Am J Epidemiol 2007;167(1):51–8.
- Sianos G, Morel MA, Kappetein AP, Morice MC, Colombo A, Dawkins K, et al. The SYNTAX Score: an angiographic tool

grading the complexity of coronary artery disease. EuroIntervention 2005;1(2):219–27.

- Mills KT, Bundy JD, Kelly TN, Reed J, Kearney PM, Reynolds K, *et al.* Global Disparities of Hypertension Prevalence and Control: A Systematic Analysis of Populationbased Studies From 90 Countries. Circulation 2016;134(6):441–50.
- Kearney PM, Whelton M, Reynolds K, Muntner P, Whelton PK, He J. Global burden of hypertension: analysis of worldwide data. Lancet 2005;365(9455):217–23.
- 11. Lawes CM, Vander Hoorn S, Rodgers A. Global burden of blood-pressure-related disease, 2001. Lancet 2008;371(9623):1513–8.
- Forouzanfar MH, Alexander L, Anderson HR, Bachman VF, Biryukov S, Brauer M, et al. Global, regional, and national comparative risk assessment of 79 behavioural, environmental and occupational, and metabolic risks or clusters of risks in 188 countries, 1990-2013: a systematic analysis for the Global Burden of Disease Study 2013. Lancet 2015;386(10010):2287–323.
- 13. Wong TY, Mitchell P. Hypertensive retinopathy. N Engl J Med 2004;351(22):2310–7.
- 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.
- Wong TY, Klein R, Couper DJ, Cooper LS, Shahar E, Hubbard LD, *et al.* Retinal microvascular abnormalities and incident stroke: the Atherosclerosis Risk in Communities Study. Lancet 2001;358(9288):1134–40.

| Submitted: 14 August, 2018 | Revised: 20 January, 2019 | Accepted: 3 February, 2019 |
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