



IJMPR



Copyright@IJMPR

Study of QT Prolongation in ECG in Type 2 Diabetes patients and its Correlation to HbA1c Levels

Dr. Midhun S¹ & Dr.Suneetha DK²

¹MBBS, MD Internal Medicine, Mysore Medical College

²MBBS, MD Medicine, Mysore Medical College

ABSTRACT

Diabetes is one of the common diseases of 21st century. Long standing Diabetes is one of the risk factor for development of all microvascular complications of diabetes. HbA1c variability is an independent risk factor for the same. Cardiovascular autonomic neuropathy is present in 16-40% of diabetic population. High QTc indicates a dysfunction of the autonomic system of heart in patients with DM. In our study 75 adults with duration of diabetes longer than 5 years were randomly selected, their HbA1c measured and correlated with QTc interval. Analysis showed that ECG changes (blocks or ischemia) were present in 22.6 % of subjects. And the mean QTc interval was 397ms. Correlation crosstabs between QTc and HbA1c showed strong positive correlation (pearson correlation =0.677, p value =0.001) between HbA1c and QTc. The study demonstrates that there is strong positive correlation between HbA1c and QTc interval suggesting high incidence of microvascular complications in patients with high HbA1c.

Key Words: Type 2 Diabetes, QTc prolongation, Cardiac autonomic neuropathy, Retinopathy, HbA1c, microvascular complication, Sudden cardiac death.



*Corresponding Author

Dr. Suneetha DK

MBBS, MD Medicine, Mysore Medical College

INTRODUCTION

- Diabetes is one of the most prevalent diseases in the world. It has been projected that worldwide approximately 570 million people would be afflicted with it by 2030.[1]
- Autonomic dysfunction in diabetes is common. Abnormal cardiovascular test suggesting cardiovascular autonomic neuropathy is present in 16-40% of diabetic population.[2,3,]
- Cardiac autonomic neuropathy is a serious complication with poor prognosis in type 2 diabetes.[4]
- Autonomic dysfunction is often asymptomatic. Hence diagnosing asymptomatic cardiac autonomic dysfunction, a precursor of symptomatic cardiac autonomic neuropathy is important.[5,6]
- CAN usually occurs in those with a long duration of diabetes, but can manifest earlier, even before the diagnosis of diabetes.[7]
- Among its clinical manifestations, resting tachycardia, orthostatic hypotension, light-headedness, visual impairment, syncope, and exercise intolerance are the most common.[8,9]
- Modifications in the equilibrium between the Sympathetic nervous system and Parasympathetic nervous system cause autonomic dysfunction.[10]
- Cardiovascular autonomic dysfunction is potentially arrhythmogenic and may predispose to atrial and ventricular arrhythmias and sudden cardiac death. [11,12]
- HbA1c variability is an independent risk factor for the development of various diabetic complications including microvascular, macrovascular and all-cause mortality in diabetes patients. This study aims to understand the correlation between HbA1c and QTc interval in type 2 diabetic patients.[13,14,15]

Methodology

This was a cross sectional observational study conducted in K R Hospital, Mysuru. This study was conducted between July 2021 and August 2022 K R hospital in Mysuru, Karnataka.

Sample size calculated using the formula $n = 4pq/d^2$ with 95% confidence interval and 5% level of significance of (Standard Deviation)=35.95 and absolute allowable error of 7%.The sample size is 75.

Inclusion criterion:

Subjects with duration of diabetes longer than 5 years were randomly selected.

Exclusion criterion :

1. Age >60
2. Documented heart disease.
3. Hypertension.
4. Uremia.
5. Dyselectrolytemia.
6. Hypo/hyperthyroidism.
7. Chronic lung disorders.
8. Fever and any other infection.

Intervention : ECG of subjects were recorded and corrected QTc interval calculated and correlated with HbA1c.

Ethical approval

Ethical approval was obtained from Mysore medical college and research institute Ethics Committee and the ethical protocols of the declaration of Helsinki (1967) including the ethical principles of informed consent, voluntary participation and withdrawal, privacy and confidentiality, were followed.

Data analysis and statistics

Data obtained from the study has been entered in excel sheets and analyzed using SPSS(Statistical package for social sciences) software version 20, and has been presented as descriptive statistics in the form of frequency, tables, figures and graphs.

- Descriptive statistics of the explanatory and outcome variables were calculated by mean, Standard deviation for quantitative variables, frequency and proportions for qualitative variables.
- Inferential statistics like-
 - 1) Chi-square test was applied for qualitative variables.
 - 2) Independent sample t test will be applied to compare the quantitative variables between the groups. The level of significance is set at 5%. A 'p' value of <0.05 is considered statistically significant.

RESULTS

49.3 % of study participants were males and rest females. The mean age was 47.9 years. 34.7% of participants had history of insulin use. 41.3% of subjects reported neuropathy symptoms. The mean duration of diabetes was 9.1 years. Further analysis showed that ECG changes (blocks or ischemia) were present in 22.6 % of subjects. The mean QTc interval was 397ms. Correlation crosstabs between QTc and HbA1c showed strong positive correlation between HbA1c and QTc.

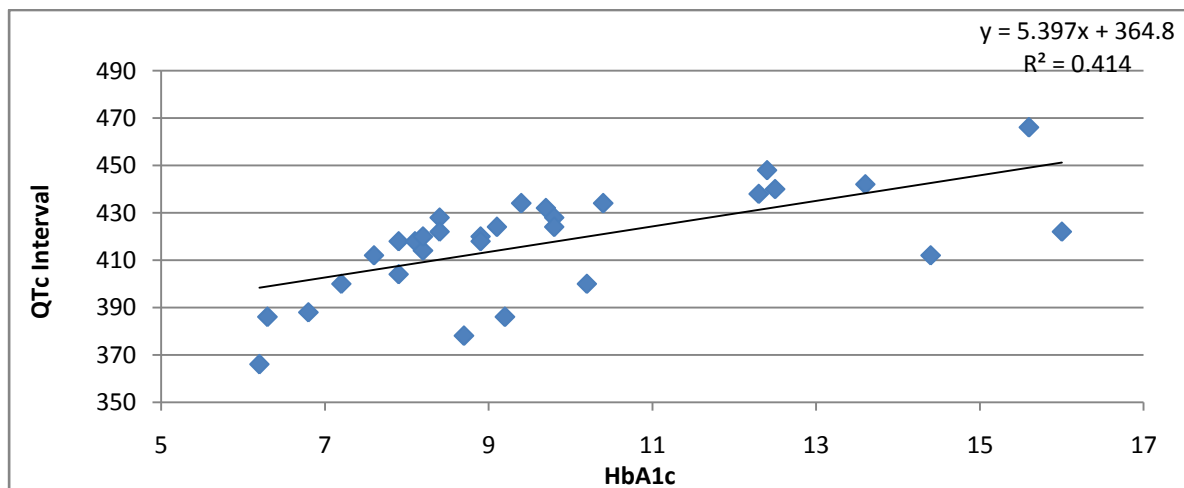


Figure 1 – Correlation between QTc and HbA1c

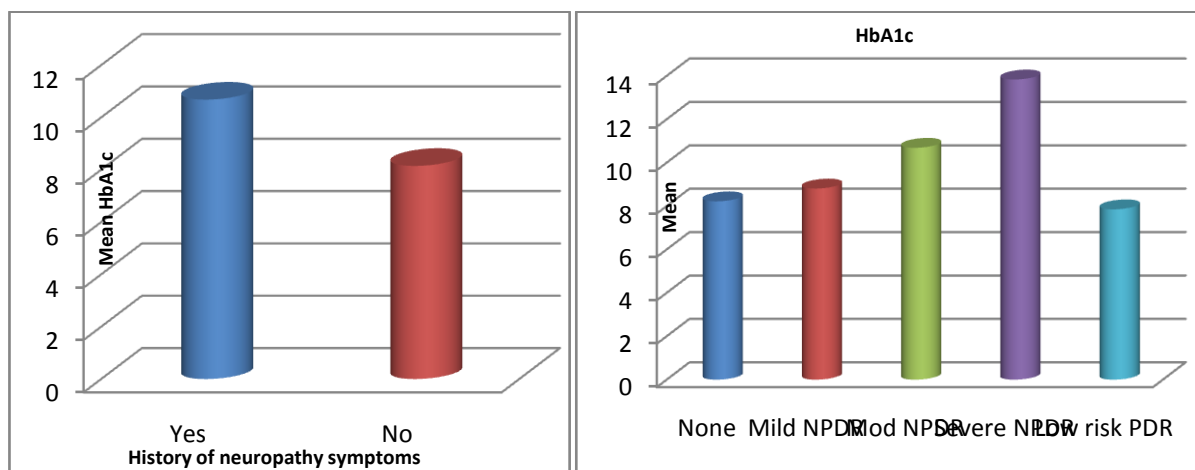


Figure 2 – Correlation between HbA1c and microvascular complications

DISCUSSION

The present study concludes that there is a strong positive between HbA1c, QTc interval suggesting high incidence of microvascular complications in patients with high HbA1c. This is also supported by the sub-study done as part of the above study which showed positive correlation between HbA1c and microvascular complications. QTc interval is a marker of cardiac autonomic neuropathy. So strict monitoring of sugars and adequate diabetic control is vital for preventing complications such as neuropathy, retinopathy, nephropathy.[16,17,18,19]

The measurement of high levels of HbA1c could be a strategic biomarker to detect diabetic complications. Indeed, intensive glycaemic control and lower levels of HbA1c are followed by a reduction in diabetic complications: in HbA1c, <7% is associated a 60% reduction in the incidence of neuropathy[20,21].

Stringent glycaemic control remains crucial for the prevention of long-term microvascular complications. Additionally, in people with diabetes, early intensive glucose control therapy not only lowers blood glucose levels but also reduces blood pressure, cholesterol, and its atherogenic subfractions. [22,23,24]

Self-monitoring of blood glucose level and enhancing medication adherence can be instrumental in meaningful improvements in HbA1c control. Additionally, the quality of diabetes management in India varies considerably with the awareness level, attitude, and perception of physicians on diabetes care[25,26]

CONCLUSION

As duration of diabetes increases odds of having any microvascular complication increases[27]. The study demonstrates that there is strong positive correlation between QTc interval and HbA1c suggesting high incidence of microvascular complications in patients with high QTc. Hence QTc can be used as a surrogate marker for diabetic control and degree of microvascular complications[28,29].

Limitations:

- This is a cross sectional study, however follow up of patients would have added more information.
- The sample size could have been a larger population.

Funding: None declared

Conflict of interest: None declared.

Ethical Approval: The study was approved by the Institutional Ethics Committee

REFERENCES

1. WHO "The Global Diabetes Compact" available at www.who.int/news/item/17-11-2020-the-global-diabetes-compact
2. National diabetes fact sheet. Available at <http://www.diabetes.org/diabetes-statistics>.
3. International Diabetes Federation Diabetes Atlas. Available at www.idf.org/diabetesatlas.
4. Rolim LC, SaÃ JR, Chacra AR, Dib SA(2008). Diabetic cardiovascular autonomic neuropathy: risk factors, clinical impact and early diagnosis. *Arq Bras Cardiol*; 90: e24-e31 [PMID: 18516377 DOI: 10.1590/s0066-782x2008000400014]
5. Serhiyenko VA, Serhiyenko AA(2018). Cardiac autonomic neuropathy: Risk factors, diagnosis and treatment. *World J Diabetes*; 9: 1-24 [PMID: 29359025 DOI: 10.4239/wjd.v9.i1.1]
6. Agashe S, Petak S(2018). Cardiac Autonomic Neuropathy in Diabetes Mellitus. *Methodist Debaque Cardiovasc J*; 14: 251-256 [PMID: 30788010]

7. Diabetes Foundation (India). Available at www.diabetesfoundationindia.org.
8. ADA (American Diabetes Association). Clinical Practice Recommendations. American Diabetes Association. Diabetes Care.
9. Breder ISS, Sposito AC(2019). Cardiovascular autonomic neuropathy in type 2 diabetic patients. *Rev Assoc Med Bras* (1992); 65: 56-60 [PMID: 30758421 DOI: 10.1590/1806-9282.65.1.56]
10. Colombo J, Arora R, Depace N, Vinik A(2015). Clinical autonomic dysfunction: Measurement, indications, therapies, and outcomes. 1st ed. Springer International Publishing : 1- 452
11. Thapa R, Bajimaya S, Sharma S, et al(2015). Systemic association of newly diagnosed proliferative diabetic retinopathy among type 2 diabetes patients presented at a tertiary eye hospital of Nepal[J]. *Nepal J Ophthalmol* ; 7:26-32.
12. Aidenloo NS, Mehdizadeh A, Valizadeh N, et al(2016). Optimal glycemic and hemoglobin a1c thresholds for diagnosing diabetes based on prevalence of retinopathy in an Iranian population[J]. *Iran Red Crescent Med J*; 18:e31254.
13. Wang Z, Chen Z, Zhang L(2018). Status of hypertension in China: results from the China hypertension survey, 2012-2015[J]. *Circulation*; 137:2344-56.
14. Frank B(2011). Globalization of diabetes, the role of diet, lifestyle and genes. *Diabetes Care*; 34(6):1249-57.
15. Veglio M, Borra M, Stevens LK, et al(2009). The relation between QTc interval prolongation and diabetic complications: The EURODIAB IDDM Complications Study Group; 42(1):68-75.
16. Banthia S, Bergner DW, Chicos AB(2013). Detection of cardiovascular autonomic neuropathy using exercise testing in patients with type 2 diabetes mellitus; 27(1):64-9.
17. Whitsel EA, Boyko EJ, Siscovick DS(2000). Reassessing the role of QTc in the diagnosis of autonomic failure among patients with diabetes: a meta- analysis. *Diabetes Care*; 23(2):241-7.
18. DiabetesControl and Complications Trial Research Group. (1998). The effect of intensive diabetes therapy on measures of autonomic nervous system function in the Diabetes Control and Complications Trial (DCCT). *Diabetologia*, 41(4), 416-423.
19. Gaede P, Lund-Andersen H, Parving HH, Pedersen O(2008). Effect of a multifactorial intervention on mortality in type 2 diabetes. *N Engl J Med*; 358: 580-591 [PMID: 18256393 DOI: 10.1056/ NEJMoa0706245]
20. Dimitropoulos G, Tahrani AA, Stevens MJ(2014). Cardiac autonomic neuropathy in patients with diabetes mellitus. *World J Diabetes*; 5(1):17-39.
21. Vinik AI, Erbas T, Casellini CM(2013). Diabetic cardiac autonomic neuropathy, inflammation and cardiovascular disease. *J Diabetes Investig*; 4(1):4- 18.
22. Spallone V, Ziegler D, Freeman R, Bernardi L, Frontoni S, PopBusui R, Stevens M, Kempler P, Hilsted J, Tesfaye S, Low P, Valensi P(2011). Toronto Consensus Panel on Diabetic Neuropathy. Cardiovascular autonomic neuropathy in diabetes: clinical impact, assessment, diagnosis, and management. *Diabetes Metab Res Rev*; 27: 639-653 [PMID: 21695768 DOI: 10.1002/dmrr.1239]
23. Vinik AI, Ziegler D(2007). Diabetic cardiovascular autonomic neuropathy. *Circulation*; 115(3):387-97.
24. Pop-Busui R(2010). Cardiac autonomic neuropathy in diabetes: a clinical perspective. *Diabetes Care*; 33(2):434-41.
25. Boulton AJ, Vinik AI, Arezzo JC, et al.(2005). American Diabetes Association. Diabetic neuropathies: a statement by the American Diabetes Association. *Diabetes Care*; 28(4):956-62.
26. Vinik AI, Erbas T, Casellini CM(2013). Diabetic cardiac autonomic neuropathy, inflammation and cardiovascular disease. *J Diabetes Investig*; 4: 4-18 [PMID: 23550085 DOI: 10.1111/jdi.12042]
27. Verma S, Alam R, Ahmad I(2018). Effect of glycemic control and disease duration on cardiac autonomic function and oxidative stress in type 2 diabetes mellitus; 17(2):149-158.
28. Chih-Cheng Huang(2016). "Diabetic Retinopathy Is Strongly Predictive of Cardiovascular Autonomic Neuropathy in Type 2 Diabetes", *Journal of Diabetes Research*, Article ID 6090749, 7 pages, 2016.
29. Boulton AJ, Vinik AI, Arezzo JC, et al.(2005). American Diabetes Association. Diabetic neuropathies: a statement by the American Diabetes Association. *Diabetes Care*; 28(4):956-62.