

# Prolongation of QTc Duration and Increased Heart Rate in Patients with Cirrhosis of Liver

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## Abstract

**Objective:** To compare QTc duration and increased heart rate in patients with cirrhosis with non-cirrhotic controls. **Design:** Cross-sectional analytical study. **Place and Duration of Study:** Medical Unit I Allied Hospital Faisalabad between 1<sup>st</sup> March 2011 to 30<sup>th</sup> August 2011. **Patients and Methods:** 50 patients of cirrhosis were selected in Group-I. An equal number of non-cirrhotic patients were taken as control and were included in Group-II. ECG was recorded and Heart rate(HR) and QTc interval was calculated in both the groups. Comparison of increased prolongation of QTc and Heart Rate were done using independent samples t

test with significance level at 0.05. **Results:** Fifty patients of cirrhosis of liver were inducted in Group-I with same number of non-cirrhotic patients as control in Group-II. The mean  $\pm$  SD of QTc of Group-I was  $0.472 \pm 0.012$  sec and that in Group-II was  $0.434 \pm 0.014$  sec and that for HR in Group-I and II were  $79.26 \pm 10.08$  and  $74.24 \pm 7.58$  beats/min respectively. The mean QTc and HR values were significantly more in Group-I as compared to Group-II with p value = 0.0001. **Conclusion:** Means of both HR and QTc were significantly higher in cirrhotic patients as compared with non-cirrhotic controls. **Key Words:** Heart rate. Cirrhosis, QTc.

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## INTRODUCTION

Cirrhosis of liver is very common in Pakistan. Hepatitis C is the commonest underlying cause in patients presenting with cirrhosis of liver followed by Hepatitis B virus. Both the viruses account for about three-fourth of all the patients presenting with cirrhosis of liver.<sup>1</sup> According to World Health Organization (WHO), about 3% of world's population is infected with Hepatitis C virus infection (HCV), with 3-4 million new cases arising every year.<sup>2</sup> Many complications can occur as a result of cirrhosis, in which ascites, portal hypertension and varices are most common. Many new complications are being recognized which include hepatopulmonary and sleep-apnoea syndromes.<sup>3</sup> Abnormalities in cardiac electro physiology are well documented in patients with liver cirrhosis.<sup>4</sup> The use of new investigative modalities has shown several lines of evidence of impaired cardiac contractility and performance in patients with cirrhosis and has led to the introduction of the new clinical entity, cirrhotic cardiomyopathy.<sup>5</sup>

Although it was first described in 1953, but was forgotten and not much work was done on it.<sup>6</sup> Changes

in Heart Rate (HR) and QTc duration are part of this new syndrome. A prolonged QTc duration in chronic liver disease could potentially lead to ventricular arrhythmias and sudden cardiac death.<sup>6,7</sup> In one of the studies conducted in this regard, The mean QTc and HR values were significantly more in patients with cirrhosis of liver as compared to non-cirrhotic controls.<sup>8</sup> The aim of this study was to compare the HR and QTc duration in patients of cirrhosis with non-cirrhotic controls.

## PATIENTS AND METHODS

It was a cross-sectional study conducted in Medical Unit I Allied Hospital Faisalabad from 1<sup>st</sup> March 2011 to 30<sup>th</sup> August 2011. 50 patients of cirrhosis of liver were included in the study and 50 controls were also included. Using convenience sampling, confirmed patients of cirrhosis were inducted after taking informed consent. All the selected patients were allocated to Group-I. 50 normal individuals were taken as control and were allocated to Group-II. Patients

with ischemic and valvular heart disease, conduction defects, cardiac failure, hypertension, hyperkalemia and patients taking blockers, calcium channel blockers, antiarrhythmic and cardiac glycosides were excluded. Clinical details were recorded of all the selected individuals on a proforma. Three 12 lead ECG recordings were taken of each patient, 5 minutes apart, and HR and QTc were calculated for each ECG and then mean of the three were calculated and used for the analysis. QTc values were calculated for all patients by the formula:  $QTc = QT/\sqrt{R-R}$ .<sup>9</sup> Heart rate were calculated on ECG by formula  $HR=1500/R-R$ <sup>10</sup> A mean value of QTc > 0.44 seconds was taken as prolonged, while the HR > 100 was taken as increased. Blood sample were taken for complete blood counts, urea, creatinine, electrolytes, LFTs, albumin, and prothrombin time. . SPSS version 10.0 was used for statistical analysis. Means of HR and QTc were compared by independent samples 't-test' between the two groups. 95% confidence intervals and p-values were calculated with significance level set at 0.05.

## RESULTS

Fifty confirmed patients of cirrhosis of liver were inducted in Group-I with same number of non-cirrhotic individuals were included in Group-II. The mean age in Group-I was 38.2 years and that in Group-II was 37.4 years (Table I). In Group I 22 patients were male and 28 were female. In Group II 18 were male and 32 were female (Table 2). Figure 1,2 show the QTc interval and heart rate in both groups respectively. The mean  $\pm$  SD of QTc on Group-I was  $0.472 \pm 0.012$  sec and that in Group-II was  $0.434 \pm 0.014$  sec (Table 3) and that for HR in Group-I and II were  $79.26 \pm 10.08$  and  $74.24 \pm 7.58$  beats/min (Table 4) respectively. Comparing the mean QTc values of the two groups proved to be statistically significant with  $p = 0.0001$ . (Table 5) Similar comparison testing for HR also proved to be significant with  $p = 0.0001$ . (Table 5) Means of both HR and QTc were significantly higher in Group-I as compared with Group-II.

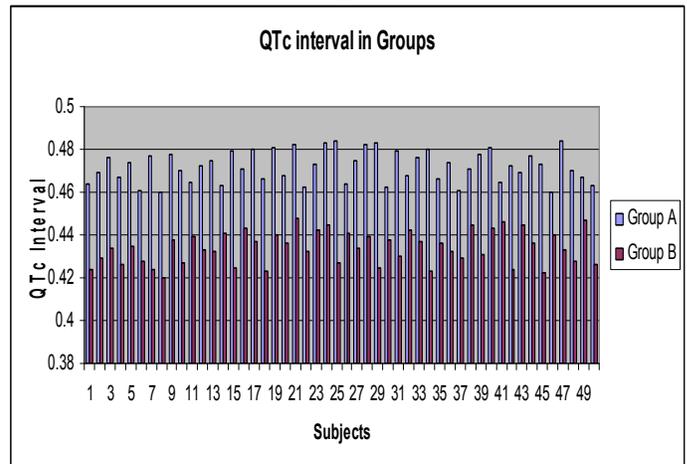
**Table-1**  
**Age Of participants**

Groups of Patients	Age
Group I	38.2
Group II	37.4

**Table-2**  
**Gender of participants**

Groups	Male	Female
Group I	22	28
Group II	18	32

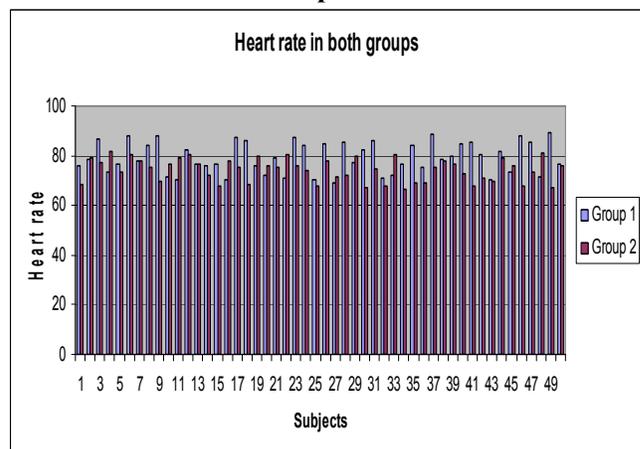
**Figure-1**  
**QTc Interval in Both Groups**



**Table-3**  
**Mean QTc Duration**

Groups I	$0.472 \pm 0.012$
Group II	$0.434 \pm 0.014$

**Figure-2**  
**Heart Rate in Both Groups**



**Table-4**  
**Mean Heart Rate**

<b>Group I</b>	79.26 ±10.08
<b>Group II</b>	74.24 ± 7.58

**Table-5**  
**Independent Samples Test**

	t	Sig. (2-tailed)
QT interval	25.418	.000
heart rate	4.515	.000

**DISCUSSION**

Cirrhotic cardiomyopathy is diagnosed infrequently because of relative unawareness regarding this entity. It has many features including prolongation of QTc, increased HR, decreased myocardial contraction force and diastolic dysfunction.<sup>6,11</sup> Several electrophysiological mechanisms like reduced beta-adrenoceptor density, postreceptor signal defects, abnormal excitation-contraction coupling have been suggested as the cause of molecular abnormalities for the conductance and impaired cardiac contractility.<sup>10</sup> Beta-receptor density and sensitivity is reduced in cirrhosis, along with altered G protein and calcium channel functions.<sup>5,11</sup> This results in both impaired chronotropic responses and electromechanical uncoupling; The coupling between the cardiac output and arterial compliance is an important factor affecting the left ventricular stress and work done by it.<sup>12</sup> The increased interval correlates with a higher incidence of sudden cardiac death. The pathogenesis of increased QT interval is unclear. The structural changes in cardiomyocyte membrane with increased cholesterol content with resultant membrane fluidity compromises the calcium and potassium pumps. In cirrhotics increased plasma levels of estrogens has also been implicated for the increased incidence of QT interval prolongation. This interval is increased in 30 to 60% of patients and level of increase relates to degree of hepatic dysfunction.<sup>5</sup> On the other hand, a too compliant arterial system will hamper prompt and timely delivery of blood to different parts

of the body and also delay flow in important vascular beds. These effects will be more prominent in the patients with excessive cardiac output, stroke volume and vascular beds of varying vascular resistance as in cirrhotic cardiomyopathy.<sup>7</sup> Prolongation of QTc duration and increased Heart rate can be used as a non-invasive and rapid diagnostic marker of cirrhotic cardiomyopathy as was proved in the study conducted in 2007.<sup>8</sup> Prolongation of QTc interval has been shown to be useful for assessment of severity of chronic liver disease.<sup>13</sup> QTc duration can be reduced by prompt usage of beta-blockers, preventing life threatening arrhythmias so early diagnosis is important. In our study, we found significant increase in mean values of QTc duration and HR between the two groups.

**CONCLUSION**

Prolongation of QTc duration and increased Heart rate can be used an important bedside marker of cirrhotic cardiomyopathy in patients with cirrhosis of liver. However larger cross-sectional studies are needed to chalk out guidelines for diagnosis of cirrhotic cardiomyopathy and cutoff values of QTc duration.

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