# Effect of Balloon Inflation-Induced Acute Ischemia on QT Dispersion during Percutaneous Transluminal Coronary Angioplasty

KUDRET AYTEMIR, M.D., VELI BAVAFA, M.D., NECLA OZER, M.D., SERDAR AKSOYEK, M.D., ALI OTO, M.D., FESC, FERHAN OZMEN, M.D., Hacettepe University, Faculty of Medicine, Department of Cardiology, Ankara, Turkey

#### Summary

Background: QT dispersion (QTd =  $QT_{max} - QT_{min}$ ) measured as interlead variability of QT interval reflects the spatial inhomogeneity of ventricular repolarization times, and increased QTd may provide a substrate for malignant ventricular arrhythmias. Ischemia is associated with regional abnormalities of conduction and repolarization.

*Hypothesis:* This study aimed to investigate the effect of acute ischemia on QTd during successful percutaneous transluminal coronary angioplasty (PTCA).

*Methods:* Forty-three patients (10 women, 33 men, mean age 56 years) were enrolled in the study. Electrocardiogram (ECG) recordings were taken before PTCA and during balloon inflation period. QT maximum (QT<sub>max</sub>), QT minimum (QT<sub>min</sub>), and QTd (QT<sub>max</sub> – QT<sub>min</sub>) values were calculated from the surface ECG.

*Results:* There was no difference among  $QT_{max}$  values (p = 0.6). Mean  $QT_{min}$  during balloon inflation was lower than before PTCA (368 ± 45 vs. 380 ± 41 ms, p = 0.002). The difference between QTd values before and during balloon inflation was statistically important (65 ± 9 vs. 76 ± 10 ms, p = 0.001). This difference is caused by a decrease in  $QT_{min}$  during balloon inflation.

Conclusion: Acute reversible myocardial ischemia induced by balloon inflation causes an increase in QTd value, and this increment is the result of a decrease in  $QT_{min}$  interval. Therefore, QTd may be a marker of reversible myocardial ischemia.

Į

Address for reprints:

Dr. Kudret Aytemir 61 Sheldrick Close SW19 2UQ London UK

Received: April 14, 1998 Accepted with revision: August 25, 1998 **Key words:** percutaneous transluminal coronary angioplasty, acute ischemia, QT dispersion

# Introduction

QT dispersion (QTd) measured as interlead variability of QT interval reflects the spatial inhomogeneity of ventricular repolarization times.<sup>1-4</sup> This inhomogeneity represents an important factor predisposing to ventricular arrhythmias.<sup>5</sup> The effects of acute ischemia and reperfusion on the homogeneity of ventricular repolarization have been studied extensively in animal models.<sup>6, 7</sup> Acute ischemia interferes with normal repolarization by changing action potential duration and conduction velocity<sup>8</sup> and can alter the QT interval and QTd.

In this study, we aimed to investigate the effect of acute ischemia on QTd during the balloon inflation period at percutaneous transluminal coronary angioplasty (PTCA).

# Methods

#### **Study Patients**

Consecutive patients with clinically stable angina pectoris referred for elective PTCA were screened for inclusion in this study. Patients had symptomatic single-vessel coronary artery disease and no history of myocardial infarction (MI). Patients were excluded if they were taking drugs that modify the QT interval, had a congenital long QT syndrome, or had an intraventricular conduction delay or left ventricular hypertrophy. Patients undergoing atherectomy or stent implantation were also excluded.

Percutaneous transluminal coronary angioplasty was performed with standard approach. Nonionic contrast medium was used for all patients. Standard paper-based 12-lead electrocardiograms (ECGs) were recorded (all 12 leads simultaneously, Cardiotest EK53R, PPG Hellige, Freiburg, Germany) at a 50 mm/s paper speed before the first inflation (baseline) and then 60 s after intracoronary balloon inflation. Only recordings of the first inflation were used to avoid ischemic preconditioning. Patients without chest pain during the preprocedure (baseline) ECG were included in the study. Electrocardiograms were also excluded if the QT interval could not be measured accurately in at least six leads. QT intervals were measured manually from the onset of QRS to the end of the T wave, defined as a return to the T-P baseline. If U waves were present, the QT interval was measured to the nadir of the curve between the T and U waves. Four consecutive cycles in each of the 12 ECG leads were used to measure the QT interval, and then mean QT intervals were calculated. QT interval dispersion was defined as the difference between the maximal and minimal QT interval measurements occurring in any of the 12 leads on ECG. All ECG measurements were made by two experienced cardiologists who were blinded to the time they were obtained. Blinded inter- and intraobserver reproducibility of QT measurement was evaluated, and comparison revealed a Spearman correlation coefficient of 0.88 and 0.90, respectively (p = 0.001).

#### ST-Segment Changes

ST-segment deviation was measured at 60 ms after the J point. ST-segment elevation was considered significant when 1 mm ST-elevation occurred in two contiguous leads. ST-segment depression was considered significant when 1 mm ST depression occurred in two contiguous leads.

#### **Statistical Analysis**

For statistical analysis, the *t*-test and the Mann Whitney U test were used. Correlational analysis was performed by using the Spearmen rank correlation test. All data were presented as mean value  $\pm$  standard deviation. A p value of <0.05 was considered significant.

# Results

Of the 62 patients screened for inclusion, 19 were excluded. The study group consisted of 43 patients (10 women and 33 men; mean age  $56 \pm 10$ , range 34–75 years). The coronary artery lesion involved the left anterior descending (LAD) coronary artery in 32 patients, the right coronary artery (RCA) in 5 patients, and the left circumflex artery (Cx) in 6 patients. During balloon inflation, ST-segment elevation in 4 patients, and ST-segment depression in 32 patients were seen, and 7 patients did not have ST-segment changes. No significant difference was found in heart rate values at baseline and during the balloon inflation period ( $68 \pm 14$  vs.  $67 \pm 15$  beats/min, p = 0.8, Table I).

TABLE I Mean heart rate, QTmax, QTmin, and QT dispersion values

	Heart rate (beats/min)	QT <sub>max</sub> (ms)	QT <sub>min</sub> (ms)	QT dispersion (ms)
Before PTCA	68±14	$440 \pm 38$	$380 \pm 41$	65±9
During balloon inflation	67±15	$446 \pm 20$	368±45	76±10
p Value	0.8	0.6	0.002	0.001

Maximum QT values  $(QT_{max})$  were  $440 \pm 38$  ms before PTCA,  $446 \pm 20$  ms during balloon inflation, and the difference was not significant (p = 0.6, Table I). Minimum QT values (QT<sub>min</sub>) were found to be  $380 \pm 41$  ms and  $368 \pm 45$  ms, and difference between QT<sub>min</sub> was significant (p = 0.002, Table I, Fig. 1). QT dispersion values were  $65 \pm 9$  ms before PTCA and  $76 \pm 10$  ms during balloon occlusion (p = 0.001, Table I, Fig. 2). There was no difference in QTd in patients developing ST-segment depression, ST-segment elevation, or in those without ST-segment changes ( $74 \pm 11$  ms;  $76 \pm 10$  ms, and  $75 \pm 12$ , respectively; p > 0.05).

# Discussion

The principal finding of our study is that QTd significantly increases during acute ischemia and that this increment is the result of a decrease in QT<sub>min</sub> interval.

Percutaneous transluminal coronary angioplasty represents a suitable model of reversible acute ischemia and reperfusion since ECG recordings can be exactly timed with these events.<sup>9, 10</sup> For determination of acute ischemia, several clinical and ECG parameters have been used.<sup>9–15</sup> The presence of QTd on the surface ECG is a marker of the dispersion of myocardial repolarization.<sup>1, 5, 16, 17</sup> As is known, QTd is defined as

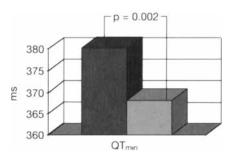


FIG. 1 Mean QT minimum values (before percutaneous transluminal coronary angioplasty and during balloon inflation).  $\blacksquare$  = Before,  $\blacksquare$  = during.

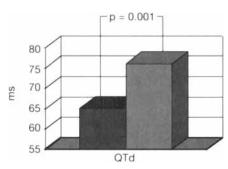


FIG. 2 Mean QT dispersion values (before percutaneous transluminal coronary angioplasty and during balloon inflation).  $\blacksquare$  = Before,  $\blacksquare$  = during.

23

the interlead difference between the longest and shortest QT interval on a standard ECG.<sup>1, 16, 17</sup> Increased QTd has also been shown to be a useful predictor of arrhythmia risk in patients with long QT syndrome, acute MI, cardiomyopathy, and heart failure.<sup>17-21</sup> Recent studies have also shown that QTd increases during stress test-induced ischemia in patients with coronary artery disease.<sup>22,23</sup> In many studies, the effects of PTCA on QTmax, QTmin, and QTd were evaluated.24-27 Pande et al.9 and Tarabey et al.27 found no significant change in QT<sub>max</sub> interval in affecting leads during balloon inflation. However, the studies in patients with acute MI suggested that there was increased QT<sub>max</sub> interval value during ischemia.4,28 During acute myocardial infarction, ischemia and many clinical factors, such as autonomic nervous system function, and hemodynamic factors affect the QT<sub>max</sub>. This may explain the difference between QT<sub>max</sub> in patients with acute MI and in those with acute ischemia during balloon inflation. In addition, we found no significant change in QT<sub>max</sub> during balloon inflation-induced ischemia, and our finding is in agreement with Tarabey et al.27 and Pande et al.9 who observed no significant change in QT<sub>max</sub> interval; hence, acute ischemia may not affect the QT<sub>max</sub> interval.

Many studies showed QT<sub>min</sub> was reduced by acute ischemia.<sup>24, 25, 27</sup> We found results very similar to these studies, that is, that QTmin during acute ischemia decreased. Our findings reported in this study also support the concept of increased QTd during acute ischemia and suggest that increased QTd is mainly caused by decreased QT<sub>min</sub> interval. Also, in a similar manner, Tarabey et al.27 and Michelucci et al.24 found that shortening of QT<sub>min</sub> in the affected leads was the main contributor to increased QTd during acute ischemia. In our study, increase in QTd was evident in all patients with or without ST-segment changes. In contrast to our study, Tarabey et al.<sup>27</sup> reported that increase in QTd was evident in patients with ST elevation but found no significant QTd change in patients with ST-segment depression. However, their results are conflicting because previous studies have shown QTd to increase during true positive stress test with ST-segment depression.<sup>22, 23, 29,30</sup> Therefore, we suggest that QTd may increase in patients with ischemic ST-segment depression. Our findings are parallel with other studies that were performed in patients with induced ischemia.22, 29, 30

Because all of our patients had coronary artery disease, we found higher mean QTd before PTCA than those found in previous studies in normal populations,<sup>31–33</sup> a finding confirmed in other studies of patients with coronary artery disease.<sup>5, 21, 32–34</sup>

Our findings support the hypothesis that in acute ischemia induced during balloon inflation, some repolarization differences occur between normal and ischemic myocardium, and this may permit the development of a reentrant circuit permitting the development of ventricular tachycardia. Therefore, ischemia might result in an increase in QTd and in ventricular tachycardia or ventricular fibrillation.

It has been shown in a previous study that repeated balloon occlusions by ischemic preconditioning may cause reduction in QTd.<sup>35</sup> To avoid the effects of ischemic preconditioning, we

did not use repeated balloon occlusion periods; measurements were taken only during the first balloon inflation period.

# Conclusion

QT dispersion is increased by balloon inflation-induced acute ischemia, and this increment is due to a decrease in QT minimum value. Increased QT dispersion may be a simple and inexpensive marker of myocardial ischemia, and additional studies are necessary to define the clinical applicability of QT dispersion in the assessment of acute myocardial ischemia.

# References

- Higham PD, Campbell RWF: QT dispersion. Br Heart J 1994;71: 508–510
- Statters BM, Malik M, Ward DE, Camm AJ: QT dispersion: Problems of methodology and clinical significance. J Cardiovasc Electrophysiol 1994;5:672–685
- Leeuwen DU, Hailen B, Wehi M: Spatial distribution of QT intervals: An alternative approach to QT dispersion. *PACE* 1996;19: 1894–1899
- Morena FL, Villanueva T, Karagounis LA, Anderson JL: Reduction in QT interval dispersion by successful thrombolytic therapy in acute myocardial infarction. TEAM-2 Study Investigators. *Circulation* 1994;90:94–100
- Zaidi M, Robert A, Fesler R, Derwael C, Brohet C: Dispersion of ventricular repolarization: A marker of ventricular arrhythmias in patients with previous myocardial infarction. *Heart* 1997;78: 371–375
- Taggart P, Sutton PM, Spear DW, Prake HF, Swanton RH, Emanuel RW: Simultaneous endocardial and epicardial monophasic action potential recordings during brief periods of coronary artery ligation in the dog: Influence of adrenaline, beta blockade and alpha blockade. *Cardiovasc Res* 1988;22:900–909
- Coronel R, Wilms-Schopmann F, Opthof T, Cinca I, Fiolet JW, Janse MI: Reperfusion arrhythmias in isolated perfused pig hearts. Inhomogeneities in extracellular potassium, ST and TQ potentials, and transmembrane action potentials. *Circ Res* 1992;71:1131–1142
- Yan GX, Yamada KA, Kleber AG, McHowat J, Corn PB: Dissociation between cellular K+ loss, reduction in repolarization time, and tissue ATP levels during myocardial hypoxia and ischemia. *Circ Res* 1993;72;3:560–570
- Pande AK, Meier B, Urban P, Moles V, Dorsaz PA, Foure J: Intracoronary electrocardiogram during coronary angioplasty. *Am Heart J* 1992;124:337–341
- Wohlgelernter D, Cleman M, Highman HA, Fetterman RC, Duncan JS, Zaret BL, Jaffe CC: Regional myocardial dysfunction during coronary angioplasty: Evaluation by two-dimensional echocardiography and 12 lead electrocardiography. J Am Coll Cardiol 1986;7:1245–1254
- Serruys WP, William W, Brower RW: Left ventricular performance, regional blood flow, wall motion, and lactate metabolism during PTCA. *Circulation* 1984;70:25–36
- Maeda T, Saikawa T, Niura H, Kahmatsu K, Shimayama N, Hara M, Maruyama T, Ito M, Takaki R: QT interval shortening and ST elevation in intracoronary ECG during PTCA. *Clin Cardiol* 1992; 15:525–528
- Vaitkus PT, Miller JM, Buxton AE, Josephson ME, Laskey WK: Ischemia induced changes in human endocardial electrograms during percutaneous transluminal coronary angioplasty. *Am Heart J* 1994;127:1481–1490

- Katsaris GA, Tsaritsoniotis EI, Tsounor IP, Panisois KD, Katsaris IA, Kaprinis IK, Roussis SX: Surface electrocardiogram in the detection of myocardial ischemia during percutaneous coronary angioplasty. *Angiology* 1993;44:797–802
- Shirota K, Ogino K, Hoshio A, Kasahara T, Katakelt H, Endo S, Mashiba H: Changes of QRS axis in transient myocardial ischemia induced by percutaneous transluminal coronary angioplasty. *Eur Heart J* 1994;15:1391–1395
- Glancy JM, Garratt CJ, Woods KL, de Bono DP: QT dispersion and mortality after myocardial infarction. *Lancet* 1995;345: 945–948
- Barr CS, Abdelwahab N, Freman M, Lang CC, Struthers AD: QT dispersion and sudden unexpected death in chronic heart failure. *Lancet* 1994;343:327–329
- Day CP, McComp JM, Cambell RWF: QT dispersion: An indication of arrhythmias in patients with long QT intervals. Br Heart J 1990;63:342–344
- Day CP, McComb J, Matthers E, Campbell RWF: Reduction in QT dispersion by sotalol following myocardial infarction. *Eur Heart J* 1991;12:423–427
- Buja G, Miorelli M, Turrini P, Melacini P, Nava A: Comparison of QT dispersion in hypertrophic cardiomyopathy between patients with and without ventricular arrhythmias and sudden death. Am J Cardiol 1993;72:973–976
- Perkiomaki JS, Kaistinen J, Yi-Mary S, Hikuri HV: Dispersion of QT interval in patients with and without susceptibility to ventricular tachyarrhythmias after previous myocardial infarction. J Am Coll Cardiol 1995;26:174–179
- Roukema G, Singh JP, Meijs M, Carvallo C, Hart G: Effect of exercise-induced ischemia on QT interval dispersion. Am Heart J 1998;135:88–92
- Sortan SC, Taggart P, Sutton P, Walker JM, Hardman SM: Acute ischemia: A dynamic influence on QT dispersion. *Lancet* 1997;349: 306–369
- 24. Michelucci A, Padeletti L, Frati M, Mininni S, Chelucci A, Stochinom L, Simonetti I, Giglioli C, Margheri M, Gensini GF: Effects

of ischemia and reperfusion on QT dispersion during coronary angioplasty. PACE 1996;19:1905–1908

- Yunus A, Gillis A, Traboulsi M, Duff HJ, Wyse DG, Knudtson ML, Mitchell LB: Effect of coronary angioplasty on precordial QT dispersion. *Am J Cardiol* 1997;79:1339–1342
- Kelly RF, Parillo JE, Hollenberg SM: Effect of coronary angioplasty on QT dispersion. Am Heart J 1997;134:399–405
- Tarabey R, Sukenik D, Molnar J, Somber CJ: Effect of intracoronary balloon inflation at percutaneous transluminal coronary angioplasty on QT dispersion. *Am Heart J* 1998;135:519–522
- Van de Loo A, Arendts W, Hohnloser S: Variability in QT dispersion measurements in the surface electrocardiogram in patients with acute myocardial infarction and in normal subjects. Am J Cardiol 1994;74:113–118
- Aytemir K, Bavafa V, Aksoyek S, Tokgozoglu L, Oto A. Ozmen F: The value of the QT dispersion in the noninvasive diagnosis of restenosis after percutaneous transluminal coronary angioplasty (abstr). PACE 1997;20:1489
- Stoletniy LN, Pai RG: Value of QT dispersion in the interpretation of exercise stress test in women. *Circulation* 1997;96:904–910
- Jordaens L, Missault L, Pelleman G, Duprez D, De Backer G, Clement DL: Comparison of athletes with life-threating ventricular arrhythmias with two groups of healthy athletes and a group of normal control subjects. *Am J Cardiol* 1994;74:1124–1128
- Molnar J, Rosenthal JE, Weiss JS, Somberg CJ: QT interval dispersion in healthy subjects and survivors of sudden cardiac death. Circadian variation in a twenty-four hour assessment. Am J Cardiol 1997;79:1190–1193
- Kautzner J, Malik M: QT interval dispersion and its clinical utility. PACE 1997;20:2625–2640
- Zareba W, Moss AJ, la Cessie S: Dispersion of ventricular repolarization and arrhythmic cardiac death in coronary artery disease. Am J Cardiol 1994;74:550–553
- Okishige K, Yamashita K, Yoshinaga H, Azegami K, Satoh T, Goseki Y, Fujii S, Ohira H, Satake S: Electrophysiologic effects of ischemic preconditioning on QT dispersion during coronary angioplasty. J Am Coll Cardiol 1996;28:70–73