

# Changes in heart rate variability caused by coronary angioplasty depend on the localisation of coronary lesions

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## Abstract

**Background:** Coronary angioplasty (PTCA) is a common treatment method in patients with coronary heart disease, but its effects on heart rate variability (HRV) have not been well established.

**Aim:** To verify whether the localisation of coronary lesion undergoing PTCA affects HRV parameters.

**Methods:** Ninety six consecutive individuals underwent elective coronary angiography with subsequent ad hoc successful PTCA. Two five-minute ECG were recorded, one before PTCA and the second 24-hour after PTCA. The HRV indices were determined by means of classical and 'new' mathematical models.

**Results:** The PTCA-induced changes in HRV variables depended on the localisation of dilated lesion. PTCA of the circumflex artery revealed the most significant HRV changes – a decrease in value of domain indices: Yeh DI ( $0.033\pm 0.031$  vs.  $0.011\pm 0.006$  un/unitless,  $p=0.005$ ), Yeh II ( $0.053\pm 0.039$  vs.  $0.032\pm 0.013$  un,  $p=0.017$ ), Organ BAND ( $9.101\pm 9.245$  vs.  $4.62\pm 2.205$  bpm/beat per minute,  $p=0.031$ ), Huey STV ( $208.821\pm 262.248$  vs.  $76.444\pm 35.281$  bpm,  $p=0.013$ ), Dalton MABB ( $15.733\pm 16.575$  vs.  $7.57\pm 4.89$  ms,  $p=0.015$ ), Dalton SD ( $48.741\pm 37.468$  vs.  $27.759\pm 10.533$  ms,  $p=0.015$ ), Zugaib STV ( $0.0129\pm 0.0132$  vs.  $0.005\pm 0.003$  un,  $p=0.005$ ), SDNN ( $27.204\pm 18.592$  vs.  $21.329\pm 32.784$  ms,  $p=0.044$ ), rMSSD ( $56.239\pm 19.751$  vs.  $51.496\pm 43.889$  ms,  $p=0.025$ ) and increased LF/HF ( $2.384\pm 2.072$  vs.  $5.632\pm 5.379$  un,  $p=0.044$ ). Angioplasty of the right coronary artery resulted in decreased AR TP ( $18.273\pm 2.296$  vs.  $17.085\pm 2.256$  ms<sup>2</sup>,  $p=0.017$ ) and alteration of the sympathovagal balance of the autonomic nervous system towards predominance of sympathetic activity: AR LF ( $0.264\pm 0.029$  vs.  $0.284\pm 0.040$  un,  $p=0.007$ ), LF/HF ( $4.310\pm 4.457$  vs.  $6.958\pm 7.013$  un,  $p=0.018$ ), HF ( $0.199\pm 0.165$  vs.  $0.141\pm 0.157$  un,  $p=0.031$ ), AR HF ( $0.647\pm 0.043$  vs.  $0.621\pm 0.054$  un,  $p=0.014$ ). PTCA of the left anterior descending artery caused no change.

**Conclusion:** Changes in heart rate variability caused by coronary angioplasty depend on the localisation of coronary lesions.

**Key words:** heart rate variability, percutaneous transluminal coronary angioplasty, stable angina

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## Introduction

Heart rate variability (HRV) refers to variability of the intervals between successive sinus beats, i.e. R-R intervals. The HRV changes result from the impact of autonomic nervous system tone on sinus node function. Moreover, they reflect the effects of the humoral substances released locally and peripherally. Up to now, HRV parameters have been shown to adequately predict poor prognosis in ischaemic heart disease patients, particularly after myocardial infarction (MI). These parameters are also used to detect autonomic neuropathy. The HRV analysis is not used routinely in clinical practice mainly due to either the fact that it is

a time-consuming technique associated with 24-hour ECG (electrocardiographic) signal recordings or a lack of standardisation of diagnostic methods as well as some technical difficulties. An attempt to make analysis easier led to successful replacement of 24-hour signal registration by short-term measures lasting 2 to 15 minutes [1]. New HRV parameters as well as possibilities for their more common application in clinical practice have been sought [2-4].

It has been shown that HRV indices decrease after percutaneous revascularisation [5], in particular parasympathetic-related ones [6,7]. The results of some studies indicate that this drop is accompanied by increased sympathetic tone [6]. Moreover, variable reaction was also

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observed, i.e. a decrease in some patients and an increase in others [8]. It was also shown previously that percutaneous transluminal coronary angioplasty (PTCA) of the infarct-related artery over a period of approximately 3 months following MI resulted in an elevation of overall autonomic tone, especially regarding the parasympathetic-related parameters [9]. The reasons for these discrepancies may be due to not completely disclosed underlying phenomena of HRV changes associated with myocardial ischaemia and improved perfusion. This may explain why in spite of many promising reports regarding the possibility of HRV use in diagnosis of either early atherosclerotic lesions [2] or restenosis following PTCA [3, 8], other studies [10] did not confirm such a possibility. However, the degree of coronary artery disease (CAD) [11] or localisation of the targeted artery [6] have not been shown to have any influence on the HRV results after the procedure.

The aim of this study was to assess the impact of successful PTCA on HRV variables with respect to localisation of the treated coronary artery in patients with a stable form of CAD.

## Methods

### Patients

Out of 500 consecutive individuals who underwent elective coronary angiography, 96 subjects (75 men and 21 women at the mean age of  $56.8 \pm 8.7$  years) were enrolled in the study. They presented stable CAD and had successful primary PTCA following angiography. Patients selected for this examination for reasons other than angina were not included. Moreover, patients who did not undergo PTCA due to either a lack of significant lesions in the coronary arteries or culprit lesions not qualified for PTCA were not entered into the further analysis. Patients with suboptimal effect of PTCA and also with an inadequate quality of the ECG recording were not included in the further analysis.

Patients received typical medical treatment for CAD. In the majority of them concomitant cardiovascular risk factors were also present. In order to evaluate the impact of location of the treated artery on HRV, patients were split into three subgroups. The LAD group involved 43 patients who underwent angioplasty of the left anterior descending (LAD) artery. The RCA group comprised 36 individuals with PTCA of the right coronary artery (RCA). Finally, the CX group consisted of 17 patients who underwent PTCA of the left circumflex (CX) artery.

### Coronary artery angiography

Coronary angiography and angioplasty were performed according to the well-established standards. The atherosclerotic lesions in the coronary arteries were treated with PTCA when stenosis exceed 70% of the arterial lumen. A procedure was considered successful if the final flow was TIMI 3 (complete, rapid filling of the coronary

artery with contrast medium, the presence of residual stenosis not exceeding 20%) and no complications during either the procedure or in-hospital stay.

### HRV analysis

Confirmation of the clinical relevance of the HRV analysis derived from 2 to 15-minute ECG recordings [1] allowed us to use 5-minute ECG records. The first ECG was acquired on the admission and the second one within 24 hours after angioplasty. ECG registration was always performed in the same conditions, in the morning, in the horizontal position with maximal limitation of the affecting external stimuli. The electrocardiogram was recorded using the computer software and cardiograph Kardio PC (® MEDEA Gliwice). Acquired ECG analogue signals were transformed by an analogue-digital converter with 500 Hz sampling frequency. The digital signals were then analysed on the basis of the graphic environment of LabVIEW 7.1 Express (National Instruments). The series of the heart rate (HR) derived from the ECG record was analysed using the Berger splin filtration algorithm, where sampling of the flattened HR function occurred with frequency  $f_s=1.07$  Hz [12].

A standard time domain parameters, SDNN (standard deviation of subsequent R-R intervals), rMSSD (root-mean-square of successive interval differences) and frequency domain indices such as TP (total power, including a frequency range  $\leq 0.4$  Hz), HF (high frequency, frequencies within a range of 0.15-0.4 Hz), LF (low frequency, frequencies within a range of 0.04-0.15 Hz) and LF/HF index were analysed. Spectral analysis was performed using either the Fourier rapid transformation technique or autoregressive method (AR). It allowed us to reach high resolution of frequency, stabilisation of the spectral variation estimators and spectral parameterisation. Except for total spectrum power expressed as  $ms^2$ , the frequency domain parameters were expressed as relative units. They were calculated from the rate of the given index to TP after subtraction of the value of ULF (ultra low frequency, a range of frequencies  $< 0.003$  Hz).

Moreover, the analysis was extended by evaluation of the time domain parameters calculated using short- and long-term variability models described by Yeh, Zugaib, Organ, Dalton and Huey [4]. The methods of signal transformation with respect to just either time or frequency are sometimes insufficient for practical use in biomedicine where the signal changes its spectra values as time elapses. Thus, there is a need to employ an analysis combining time and frequency domains with the use of wavelet transform  $w1-w5$  parameters [13, 14]. A description of the time domain parameters and combined time and frequency domains employed in this study is outlined in Table I.

The PTCA impact on HRV with respect to both the whole studied patient population and subgroups was

**Table I.** A description of HRV indices of time domain and combined time with frequency domains used in the study

Index	Definition	Unit
Yeh DI	Short-term variability index: standard deviation derived from a sequence of a rate difference/sum of two successive $T_{R-R}$ , $T_i - T_{i+1}$ intervals. It refers to 30-second time period.	Without unit of measure [un]
Yeh II	Long-term variability index: a rate of standard deviation to mean value of successive $T_i$ intervals. It refers to a 30-second time period.	Without unit of measure [un]
Organ BAND	In order to determine long-term variability every temporary HR value corresponding with RR interval is compared to averaged HR value calculated over a 30-second time period.	Beat per minute [bpm]
Huey's model	It is based on temporary HR value and an analysis of change trend of its successive values within time period corresponding over hundred RR periods.	
Huey STV	Index of short term variability: is a sum of modules of the successive HR value differences in a case of monotony analyzed signal change.	Beat per minute [bpm]
Huey LTV	Long-term variability is defined if no change in a sign of difference between adjacent HR values is observed over at least three subsequent intervals.	Beat per minute [bpm]
Dalton MABB	Short-term variability index: a mean value out of absolute difference between two successive $T_i$ intervals divided by 2 and measured over a 2-minute time period.	milliseconds [ms]
Dalton SD	Long term variability index: standard deviation of the consecutive $T_i$ intervals over 2-minute HR signal assessment.	milliseconds [ms]
Zugaib STV	Variability defined on the basis of $T_i$ time interval values. Supplementary $D_i$ index is calculated as time intervals difference/sum ratio. Zugaib STV index is the mean value of absolute differences between successive $D_i$ values and their median over 128 subsequent RR intervals.	Without unit of measure [un]
SDNN	Standard deviation of all sinus rhythm RR intervals duration on the basis of all 5-minute ECG blocks over the examined time period.	milliseconds [ms]
rMSSD	The square root of the mean of the squared sum of the differences between successive sinus rhythm RR intervals in the examined time period.	milliseconds [ms]
w1-w5 (wavelet)	Standard deviation of the wavelet transform is a index of combined time and frequency domains, wavelet $j$ indices describe scale $m$ factor for wavelet function and are connected by the following correlation $m=2^j$ , according to the definition, low values of the scale factor refer to spectrum of the higher frequencies, a result $m=1$ corresponds with rMSSD index, an increase of calibration parameter reduces a width of wavelet thus improving time-related resolution of the analysis accompanied by simultaneous deterioration of frequency resolution.	milliseconds [ms]

Abbreviations: DI – differential index, II – interval index, STV – short-term variability, LTV – long-term variability, rMSSD – root mean square of successive differences, SD – standard deviation, SDNN – standard deviation of NN, un – unitless

evaluated. Baseline and postprocedural HRV profiles were compared. A comparative analysis of the delta parameters following PTCA of the specific coronary arteries was carried out (delta:  $\Delta$  = the value of a given parameter before PTCA – its value following PTCA). Additionally, the impact of the group differentiating factors was analysed.

The study protocol was approved by the Bioethical Committee of the Poznań University of Medical Sciences (No. 887/04 dated 6.05.2004). All patients expressed informed consent to participate in this study.

### Statistical analysis

Data were analysed using the statistical computer software STATISTICA 7.0 (StatSoft Inc.). In the case of a lack of normal distribution, checked with Shapiro-Wilk's W test, the Mann-Whitney U test for unrelated variables was used. To compare the HRV parameters in the specific groups

before and after PTCA procedures Wilcoxon pair rank test was employed. Due to the fact that the parameters are expressed on the interval scale, the results are presented as the arithmetic mean and standard deviation. To assess the correlation on a nominal scale the exact Fisher's test and  $\chi^2$  test with Yates' correction were used. A p value of 0.05 was considered as statistically significant.

### Results

The examined group of patients was not equal regarding coexistence of diabetes mellitus and severity of the atherosclerosis assessed according to the Gensini scale. They also differed with respect to location of MI and previous angioplasty. Complete characteristics of the groups are presented in Table II.

Comparison of HRV indices before and after PTCA in all enrolled patients revealed that myocardial revascularisation

caused a drop in spectrum power within the range of high frequencies (HF) ( $0.199 \pm 0.170$  vs.  $0.141 \pm 0.153$  un,  $p=0.002$ ) and elevation of LF/HF ( $4.017 \pm 4.125$  vs.  $6.042 \pm 5.615$  un,  $p=0.0002$ ) as well as wavelet w3 index ( $3.176 \pm 3.533$  vs.  $3.503 \pm 2.988$  ms,  $p=0.048$ ).

The HRV indices that differentiated the three examined subgroups of patients at the baseline were as follows: rMSSD was found to be significantly higher in the RCA group vs. the LAD group ( $70.742 \pm 53.067$  vs.  $53.59 \pm 31.46$  ms,  $p=0.019$ ), AR TP higher in the RCA group vs. the LAD group ( $18.273 \pm 2.296$  vs.  $17.096 \pm 2.0$  ms<sup>2</sup>,  $p=0.005$ ) or CX group ( $18.273 \pm 2.296$  vs.  $16.839 \pm 2.568$  ms<sup>2</sup>,  $p=0.030$ ) (see Figures 1 and 2).

The PTCA of LAD caused no significant changes in the HRV indices.

Angioplasty of the RCA resulted in a statistically significant decrease of AR TP as well as HF or AR HF accompanied by an increase in AR LF and LF/HF (Table III).

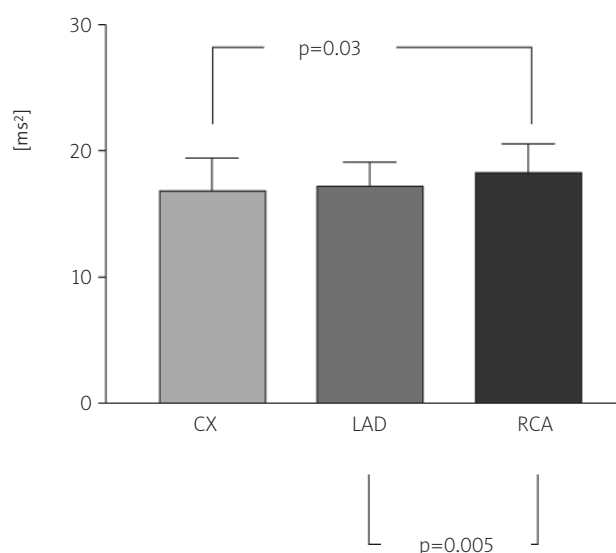


Figure 1. Comparison of baseline index AR TP

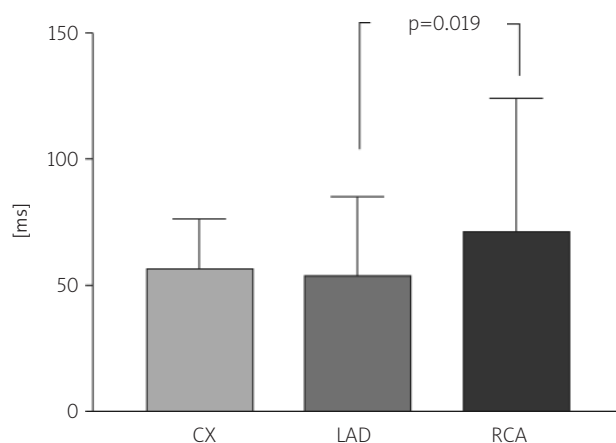


Figure 2. Comparison of baseline rMSSD values

Table II. Characteristics of the examined groups of patients

Variable	LAD n=43 n (%)	CX n=17 n (%)	RCA n=36 n (%)	p	
Gender	Women	12 (28)	2 (2)	7 (19)	NS
	Men	31 (72)	15 (88)	29 (81)	NS
Age [years]	56.98±9.2	57.4±7.1	56.1±9.1	NS	
BMI [kg/m <sup>2</sup> ]	28.4±4.2	27±3.7	26.7±3.4	NS	
Arterial hypertension	33 (77)	11 (65)	19 (53)	NS	
Diabetes mellitus type 2	13 (30)	0	5 (14)	*	
Smokers	21 (49)	13 (76)	23 (64)	NS	
Hipercholesterolemia	29 (67)	12 (71)	26 (72)	NS	
Previous myocardial infarction	Q-wave	16 (37)	5 (29)	16 (44)	NS
	Non-Q-wave	6 (14)	6 (35)	6 (17)	NS
Inferior wall infarction	8 (18)	3 (18)	19 (53)	**	
Anterior wall infarction	12 (28)	0	1 (3)	***	
Lateral wall infarction	1 (2)	4 (24)	1 (3)	****	
Indefinite location of myocardial infarction	3 (7)	4 (24)	3 (8)	NS	
Time from myocardial infarction [months]	7.3±2.7	54.7±108.9	10.3±7.6	NS	
Previous CABG	1 (2)	0	0	NS	
Previous PTCA	7 (16)	4 (24)	6 (17)	NS	
Previous PTCA LAD	3 (7)	0	2 (6)	NS	
Previous PTCA CX	0	3 (18)	3 (8)	*****	
Previous PTCA RCA	5 (12)	1 (6)	4 (11)	NS	
Time from procedure [months]	13.7±16.8	41.3±33.7	17.3±16.1	NS	
ACE inhibitors	26 (60)	9 (53)	18 (50)	NS	
Diuretics	10 (23)	3 (18)	5 (14)	NS	
Ca blockers	6 (14)	1 (6)	4 (11)	NS	
Anti-arrhythmic agents	2 (5)	0	3 (8)	NS	
Statins	34 (79)	12 (71)	32 (89)	NS	
Fibrats	0	0	3 (8)	NS	
Beta-blockers	35 (81)	11 (65)	26 (72)	NS	
Digoxin	0	0	1 (3)	NS	
Vasodilators	21 (49)	7 (41)	14 (39)	NS	
Gensini score	51.88±38.59	59.38±40.28	35.67±24.93	*****	
One-vessel disease	23 (53)	6 (35)	16 (44)	NS	
Two-vessel disease	14 (32)	8 (47)	13 (36)	NS	
Multi-vessel disease	6 (14)	3 (18)	7 (19)	NS	
Balloon angioplasty	4 (9)	4 (24)	2 (6)	NS	
1 stent implanted	31 (72)	11 (65)	27 (75)	NS	
≥2 stents implanted	8 (19)	2 (12)	7 (19)	NS	

\* LAD vs. CX,  $p < 0.001$

\*\* LAD vs. RCA,  $p=0.005$

\*\*\* LAD vs. CX,  $p=0.045$

\*\*\*\* LAD vs. CX,  $p=0.022$ ; CX vs. RCA,  $p=0.034$

\*\*\*\*\* LAD vs. CX,  $p=0.018$

\*\*\*\*\* RCA vs. CX,  $p=0.015$

In the CX group, PTCA caused a significant decrease of nine time domain indices and a marked increase of the LF/HF ratio (Table IV). Analysis of PTCA-induced changes in HRV parameters ( $\Delta$ ) revealed several significant differences (Table V).

**Table III.** Significant changes of the HRV indices following PTCA of the right coronary artery (RCA)

Index	Before PTCA		After PTCA		p
	Mean	SD	Mean	SD	
HF [un]	0.199	0.165	0.141	0.157	0.031
LF/HF [un]	4.310	4.457	6.958	7.013	0.018
AR TP [ms <sup>2</sup> ]	18.273	2.296	17.085	2.256	0.017
AR LF [un]	0.264	0.029	0.284	0.040	0.007
AR HF [un]	0.647	0.043	0.621	0.054	0.014

**Table IV.** Significant changes of the HRV indices following PTCA of the circumflex branch of the left coronary artery (CX)

Index	Before PTCA		After PTCA		p
	Mean	SD	Mean	SD	
Yeh DI [un]	0.033	0.031	0.011	0.006	0.005
Yeh II [un]	0.053	0.039	0.032	0.013	0.017
Organ BAND [bpm]	9.101	9.245	4.62	2.205	0.031
Huey STV [bpm]	208.821	262.248	76.444	35.281	0.013
Dalton MABB [ms]	15.733	16.575	7.57	4.89	0.015
Dalton SD [ms]	48.741	37.468	27.759	10.533	0.015
Zugaib STV [un]	0.0129	0.0132	0.005	0.003	0.005
SDNN [ms]	27.204	18.592	21.329	32.784	0.044
rMSSD [ms]	56.239	19.751	51.496	43.889	0.025
LF/HF [un]	2.384	2.072	5.632	5.379	0.044

**Table V.** Changes of the means of the HRV parameters ( $\Delta$ ) after PTCA of the individual coronary arteries

Index	$\Delta$ LAD	$\Delta$ CX	$\Delta$ RCA	p		
				$\Delta$ LAD vs. $\Delta$ CX	$\Delta$ LAD vs. $\Delta$ RCA	$\Delta$ CX vs. $\Delta$ RCA
Yeh DI [un]	0.0002±0.015	0.022±0.03	-0.005±0.038	0.002	NS	0.006
Yeh II [un]	-0.003±0.033	0.021±0.035	-0.009±0.043	0.007	NS	0.021
Organ BAND [bpm]	-0.725±5.39	4.481±8.377	-1.192±6.995	0.004	NS	0.033
Huey STV [bpm]	40.117±253.735	132.38±258.89	-2.888±361.89	0.014	NS	0.01
Huey LTV [bpm]	-5.642±81.876	53.055±124.15	-2.883 ±135.02	0.007	NS	NS
Dalton MABB [ms]	0.955±6.906	8.163±13.776	-2.231±25.639	0.022	NS	0.028
Dalton SD [ms]	-2.469±23.792	20.982±33.774	-8.375±41.103	0.008	NS	0.02
Zugaib STV [un]	0.0007±0.007	0.008±0.012	-0.004±0.021	0.003	NS	0.003
SDNN [ms]	8.671±39.324	5.875±37.481	-1.087±47.971	0.014	NS	NS
rMSSD [ms]	-11.258±59.757	4.743±50.879	2.482± 62.337	0.003	NS	NS
TP [ms <sup>2</sup> ]	-0.899±25.296	14.067±37.223	-4.692±23.592	0.036	NS	NS
w 2 [ms]	-0.202±2.964	2.043±4.204	-0.641±2.293	0.013	NS	0.018
AR TP [ms <sup>2</sup> ]	0.086±2.776	-0.385±2.477	1.188±3.101	NS	0.048	0.038

A trend of the most changes of HRV parameters following PTCA of CX was found to be opposite to that in the case of other coronary artery revascularisation (Figures 3 and 4).

Following angioplasty, the examined subgroups did not differ significantly between each other, although in the CX group the results of most parameters of sinus rhythm variability were the lowest.

No significant differences in the delta analysis including the impact of coexistence of diabetes mellitus, previous MI and infarct location were noted. No differences between patients with single- and multi-vessel disease were seen either. However, a comparative analysis of the HRV parameters calculated after angioplasty in the group of patients with mild ( $GS \leq 32$ ) and significant ( $GS > 32$ ) severity of atherosclerotic changes expressed on the Gensini scale revealed significant changes regarding  $\Delta$  HF and  $\Delta$  LF/HF values (Table VI).

## Discussion

Our results indicating a decrease of parasympathetic tone and autonomic balance shift towards sympathetic domination follow PTCA are consistent with the findings of Kanadasi et al. [6] and Ohler et al. [15]. However, they are in opposition to the results reported by Tseng et al. [8], who showed that the profile of the HRV parameters recorded before the procedure did not differ significantly from the profile assessed directly after the procedure.

The observed HRV changes as a result of RCA angioplasty are probably associated with sinus node blood supply most commonly from this artery. However, the results of CX and LAD angioplasty indicate that not only sinus node blood supply plays an important role in HRV profile. So far there is no known mechanism that would cause the observed phenomena. It may be

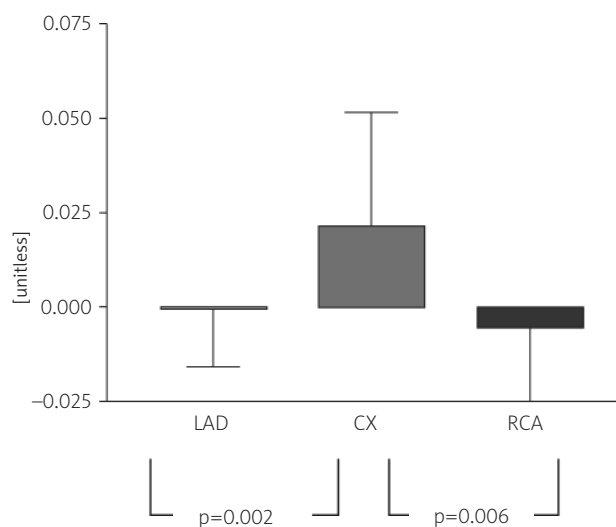


Figure 3. Differences in ( $\Delta$ ) Yeh DI index after PTCA

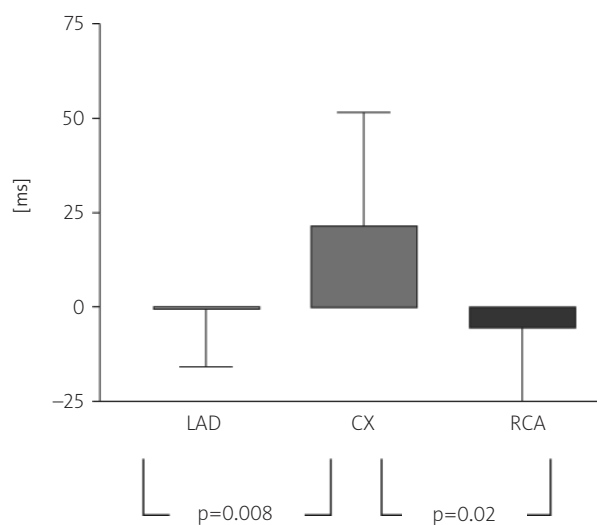


Figure 4. Differences in ( $\Delta$ ) Dalton SD index after PTCA

Table VI. Significant changes of the HRV indices after PTCA in a group of patients with less ( $GS \leq 32$ ) and more advanced atherosclerotic lesions ( $GS > 32$ ), expressed in Gensini scale (GS)

Index	$GS \leq 32$			$GS > 32$			$\Delta GS \leq 32$	$\Delta GS > 32$	p $\Delta GS \leq 32$ vs. $\Delta GS > 32$
	Before PTCA	After PTCA	p	Before PTCA	After PTCA	p			
HF [un]	0.222±0.174	0.115±0.132	0.0002	0.181±0.168	0.148±0.145	NS	0.107±0.183	0.033±0.202	0.044
LF/HF [un]	3.110±3.117	6.877±5.846	0.00001	4.701±4.654	5.547±5.427	NS	-3.767±4.519	-0.846±5.721	0.002

speculated that CX usually supplies the myocardium close to the sinus node compared to LAD. Our observations are not consistent with the findings of Kanadası et al. [6], who did not find any impact of the location of the artery treated with angioplasty on HRV changes (a relatively small population is a limitation of their study).

One may assume that unequal distribution of individuals with diabetes might affect the findings of the study. It is commonly accepted that diabetic patients and those with diabetes-induced autonomic neuropathy present impaired capacity for vasodilatation and reduced coronary flow. Rich et al. reported an inverse correlation between HRV and diabetes mellitus. However, a study published by Szot et al. [17], like our own analyses, did not reveal a significant impact of diabetes mellitus on the values of HRV parameters in response to PTCA.

Hypothetically, differences in the findings in the particular groups might have resulted from collateral circulation. Its presence may produce only slight improvement in myocardial perfusion following PTCA, contrary to the subjects without collateral circulation presenting a sudden change in myocardial perfusion [18].

Similarly to our study, Rich et al. [16] as well as Tseng et al. did not observe any relationship between Gensini scale and basic HRV profile [8]. In contrast, Hayano et al. noted a significant negative correlation between vagal

nerve tone and severity of the lesions in the coronary arteries [19]. Wachowiak-Baszyńska et al. also revealed decreased SDNN and rMSSD in patients with more extensive atherosclerotic coronary lesions [20].

An assessment of the impact of atherosclerotic lesions' severity on HRV following PTCA in our study was consistent with the findings reported by Osterhues et al. only if the classification proposed by Bruske, Proudfit and Sonnes was employed [11].

The findings of Szot et al. regarding no correlation between HRV parameters after PTCA and previous MI [17] are consistent with our results. Osterhues et al. [11] noted marked differences at baseline between patients with previous MI and subjects without it, in opposition to our results. However, the changes in HRV following PTCA in the group of patients after MI, contrary to individuals without it, were comparable with our and Osterhues studies (an increase in sympathetic parameters accompanied by a drop in parasympathetic indices) [11]. Like in our studies, Osterhues et al. did not find any relationship with the location of the previous MI [11].

The time period adopted in our study enables us to assume that our findings resulted mainly from the changes in myocardial perfusion. The effects of reperfusion-induced myocardial injury may be also taken into account. It presumes

that an increase in coronary blood flow may implicate an adverse influence on the myocardium through pressure overload and muscular filaments strain. In consequence, it leads to oedema, excessive constriction or even myocytes' death [22]. It was shown in experimental settings that many metabolic changes took place after transient coronary artery occlusion such as potassium and adenosine concentration elevation, acidosis, rapid release of free agents and mitochondria overload with calcium and also more pronounced endothelium dysfunction. These phenomena may stimulate functional denervation of the myocardial nervous endings [23]. First, consequences of ischaemia and reperfusion involve the subendocardial layer (the place where the fibre endings of the vagal nerve are located). Then they propagate to subepicardial regions. Thus, the assumption that lowered parasympathetic modulation of the sinus node, in the early period following PTCA, is a consequence of the rapid changes in blood flow. The impact of ischaemia on myocardial performance during angioplasty is seen within a short time period [24]. Meanwhile, a longer period is necessary to restore normal function to the damaged cellular structure or biochemical processes [25]. The findings of the studies published by Kanadasi et al. [6], Osterhues et al. [11] and Wennerblom et al. [7] showing a gradual increase in parasympathetic tone accompanied by a drop in sympathetic activity in the long-term follow-up suggest a delayed restoration of normal autonomic activity in consequence of successful coronary revascularisation. The results of these reports indicate improvement in vagal nerve function although complete functional restoration was not achieved. This may support the thesis that myocardial ischaemia is not the only mechanism of reduced parasympathetic tone in uncomplicated coronary artery disease.

We are aware of the limitations of our study such as a lack of long-term follow-up and heterogeneous patient population (with concomitant diseases and influenced by medical therapy). However, a lack of many often accepted exclusion criteria was an intentional step to create the most representative group for daily clinical practice.

## Conclusions

1. Percutaneous transluminal coronary angioplasty has an impact on the parameters of HRV in the early assessment, i.e. within 24 hours following the percutaneous procedure.
2. Location of the treated coronary artery significantly affects HRV; angioplasty of the left circumflex artery causes a decrease of many time domain parameters but flow restoration in the left anterior descending artery does not manifest in HRV changes. The PTCA of RCA causes changes in the frequency domain that indicate predominance of the sympathetic arm of the autonomic nervous system.
3. The degree of atherosclerotic lesion severity calculated according to the Gensini scale has an impact on the HRV changes following PTCA.

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# Wpływ angioplastyki wieńcowej na kształtowanie się zmienności rytmu serca u osób ze stabilną chorobą niedokrwienną serca z uwzględnieniem lokalizacji tętnicy poddanej zabiegowi

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## Streszczenie

**Wstęp:** Kardiologia inwazyjna stworzyła ogromne możliwości terapeutyczne. Nadal jednak brakuje skutecznych, nieinwazyjnych i łatwo dostępnych metod diagnostycznych służących wyselekcjonowaniu osób z podwyższonym ryzykiem rozwoju chorób sercowo-naczyniowych, a co za tym idzie – ukierunkowaniu dalszej diagnostyki i zminimalizowaniu czasu potrzebnego do wdrożenia optymalnego modelu leczenia. Na uwagę zasługuje analiza zmienności rytmu serca (ang. *heart rate variability*, HRV), jednakże zjawiska leżące u podłoża zmian HRV związanych z niedokrwieniem i poprawą ukrwienia nie zostały dotychczas wyjaśnione.

**Cel:** Ocena wpływu skutecznej angioplastyki wieńcowej na HRV u osób ze stabilną chorobą niedokrwienną serca (ang. *ischaemic heart disease*, IHD) z uwzględnieniem lokalizacji naczynia poddawanego zabiegowi.

**Metody:** Spośród 500 osób kolejno poddanych planowej koronarografii do badania zakwalifikowano 96 chorych w stabilnym okresie IHD, u których przeprowadzono następczą bezpośrednią angioplastykę. Chorych podzielono na trzy grupy: grupę GPZ (43 osoby), u których wykonano plastykę gałęzi przedniej zstępującej, grupę PTW (36 osób) z PTCA (ang. *percutaneous transluminal coronary angioplasty*) prawej tętnicy wieńcowej i grupę GO (17 osób) poddanych angioplastyce gałęzi okalającej lewej tętnicy wieńcowej. Dokonano dwóch 5-minutowych rejestracji EKG przed interwencją i w okresie 24 godz. po niej. Cyfrowe sygnały EKG analizowano na podstawie graficznego środowiska tworzenia aplikacji LabVIEW 7.1 Express firmy National Instruments. Zastosowano standardowe parametry: SDNN (ang. *standard deviation of NN*), RMSSD (ang. *root mean square of successive differences*), TP (ang. *total power*), HF (ang. *high frequency*), LF (ang. *low frequency*), współczynnik LF/HF. Analizę spektralną przeprowadzono metodą szybkiej transformacji Fouriera i autoregresyjną (AR). Badanie poszerzono o ocenę wskaźników w dziedzinie czasu opisanych przez Yeha, Zugaiba, Organa, Daltona, Hueya oraz analizę w połączonych dziedzinach czasu i częstotliwości, posługując się wskaźnikami falkowymi w1-w5. Do analizy statystycznej zastosowano testy U Manna-Whitneya oraz kolejności par Wilcoxa. Jako poziom istotności statystycznej przyjęto wartość  $\alpha=0,05$ . Wartości wskaźników HRV opisano średnią arytmetyczną i odchyleniem standardowym.

**Wyniki:** W wyniku PTCA w całej ocenianej populacji odnotowano spadek mocy widma HF ( $p=0,002$ ) z jednoczesnym wzrostem wskaźników: LF/HF ( $p=0,0002$ ) i w3 ( $p=0,048$ ). Angioplastyka PTW spowodowała spadek mocy całkowitej widma: AR TP ( $p=0,017$ ), wzrost aktywności współczulnej: AR LF ( $p=0,007$ ), LF/HF ( $p=0,018$ ), i spadek przywspółczulnej: HF ( $p=0,031$ ), AR HF ( $p=0,014$ ). W wyniku PTCA GPZ nie zaobserwowano istotnych statystycznie zmian wskaźników HRV. Z kolei angioplastyka GO spowodowała obniżenie wielu wskaźników czasowych: Yeh DI ( $p=0,005$ ), Yeh II ( $p=0,017$ ), Organ BAND ( $p=0,031$ ), Huey STV ( $p=0,013$ ), Dalton MABB ( $p=0,015$ ), Dalton SD ( $p=0,015$ ), Zugaib STV ( $p=0,005$ ), SDNN ( $p=0,044$ ), RMSSD ( $p=0,025$ ) i wzrost współczynnika LF/HF ( $p=0,044$ ). Analiza porównawcza delt parametrów HRV uzyskanych w wyniku PTCA poszczególnych tętnic wieńcowych ( $\Delta$  = wartość wskaźnika przed PTCA – wartość po PTCA) potwierdziła zjawiska zaobserwowane w analizach przeprowadzonych dla pojedynczych naczyń.

**Wnioski:** Zabieg angioplastyki naczyń wieńcowych wpływa na kształtowanie się parametrów zmienności rytmu zatokowego we wczesnej ocenie, to jest w okresie 24 godz. od zabiegu. Lokalizacja tętnicy poddanej rewaskularyzacji ma istotny wpływ na HRV; angioplastyka GO lewej tętnicy wieńcowej wywołuje obniżenie wielu wskaźników czasowych, udrożnienie GPZ nie przekłada się na zmiany HRV, PTCA PTW powoduje zmiany w domenie częstotliwościowej świadczące o przewodze składowej współczulnej układu autonomicznego.

**Słowa kluczowe:** zmienność rytmu serca, angioplastyka wieńcowa, stabilna choroba wieńcowa

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