

## Conference Paper

# Atrial Fibrillation Triggers in Patients with Coronary Artery Disease and Subclinical Thyrotoxicosis

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## Abstract

We examined 202 patients with paroxysms of symptomatic and asymptomatic atrial fibrillation (AF), some of whom suffered from coronary artery disease (CAD) and subclinical thyrotoxicosis (ST). Healthy individuals acted as a comparison group. It was revealed that in all studied groups, extrasystoles and paroxysms of reciprocal atrioventricular orthodromic and nodal tachycardia was the role of the triggering factors of AF. In patients with ST without CAD and in healthy persons, the paroxysms of tachycardia are short and unstable. When CAD combined with ST, the number of extrasystoles and AF paroxysms is significantly higher than only in ST and in healthy individuals. It was found that in patients with asymptomatic AF the total number of extrasystoles and paroxysms of tachycardia is greater than in the case of symptomatic. Thus, the identification of concomitant subclinical thyrotoxicosis in a patient with CAD should alert the clinician to the development and progression of atrial fibrillation. It should be given great attention to screening thyroid pathology in patients with CAD.

**Keywords:** atrial fibrillation, coronary artery disease, subclinical thyrotoxicosis.

## 1. Introduction

In recent years, there has been a trend towards growth and rejuvenation of patients with atrial fibrillation (AF) [1]. In this regard, the diagnosis and treatment of this disease have acquired medical and social significance [2].

AF is often associated with thyroid pathology, which affects its course and prognosis [3, 4]. Given the importance of thyroid hormones in the regulation of the cardiovascular system, it can be assumed that functional disorders of the thyroid system can trigger the occurrence of cardiac arrhythmias [5–7]. The presence of concomitant thyroid pathology in the patient with coronary artery disease (CAD) increases the risk of the onset and progression of AF, reduces the quality of life and the prognosis [8]. According

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Received: 17 January 2018

Accepted: 25 March 2018

Published: 17 April 2018

Publishing services provided by  
Knowledge E

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Selection and Peer-review under the responsibility of the PhysBioSymp17 Conference Committee.

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to available data, the symptomatic and asymptomatic course of AF also differs [9, 10]. Despite this, the triggering factors of atrial fibrillation in this group of patients have not been described in detail so far.

The purpose was to evaluate the electrocardiographic and electrophysiological triggering factors of asymptomatic and symptomatic atrial fibrillation in the combination of CAD with subclinical thyrotoxicosis (ST).

## 2. Materials and Methods

The study, conducted at the Penza State University (Penza, Russia), involved 202 patients, who were divided into 6 groups. The first group included 38 patients with asymptomatic paroxysms of AF on the background of a combination of CAD with ST (AFas + CAD + ST), the second – 35 patients with symptomatic paroxysms of AF on the background of a combination of CAD with ST (Ass + CAD + ST), in the third – 34 patients with asymptomatic paroxysms of AF in the presence of CAD (AFas + CAD), in the fourth – 32 patients with symptomatic paroxysms of AF in the presence of CAD (AFs + CAD), in the fifth – 33 patients with ST without CAD, and in the last, sixth group – 30 healthy individuals without documented earlier AF (HI).

Symptomatic paroxysms of AF included patients with complaints of irregular heart-beat, palpitation, loss of consciousness (specific symptoms); shortness of breath, dizziness, insomnia (nonspecific symptoms). Patients, who had more than 50% of all paroxysms not accompanied by obvious symptoms, were referred to as asymptomatic.

Methods of the study included Holter monitoring of ECG (HM) and transesophageal electrophysiological study (TEEPS). HM was performed on Astrocard apparatus, and TEEPS – on the electrophysiological complex Astrocard (CJSC "Meditex", Russia). The ECG was recorded at a belt speed of 25, 50, 100 mm/s and a signal gain of 1 mV - 10, 20 mm.

Statistical processing of research results was performed on a personal computer using the Statistica for Windows software package v. 8.0 (Stat-Soft Inc., Russia) including parametric and nonparametric criteria.

## 3. Results and Discussion

As a result of the study, it was established that the starting factors of AF in all the examined groups of patients are presented by extrasystoles, a combination of extrasystoles

with paroxysms of reciprocal atrioventricular orthodromic (AVROT) and nodal (AVRNT) tachycardia (Table 1).

TABLE 1: Trigger factors of arrhythmia in patients with symptomatic and asymptomatic AF, subclinical thyrotoxicosis and healthy individuals.

Species	AF + CAD + ST		AF + CAD		ST	HI
	Asymptomatic AF	Symptomatic AF	Asymptomatic AF	Symptomatic AF		
	Group 1	Group 2	Group 3	Group 4	Group 5	Group 6
Extrasystoles, n (%)	23 (60,5)	20 (57,1)	23 (67,6)	22 (68,8)	20 (60,6)	21 (70,0)
Extrasystoles + AVROT, n (%)	7 (18,4)	8 (22,9)	6 (17,6)	5 (15,6)	6 (18,2)	4 (13,3)
Extrasystoles + AVRNT, n (%)	8 (21,1)	7 (20,0)	5 (14,7)	5 (15,6)	7 (21,2)	5 (16,7)
Total, n (%)	38 (100)	35 (100)	34 (100)	32 (100)	33 (100)	30 (100)

**Abbreviation:** AF - atrial fibrillation; CAD - coronary artery disease; ST - subclinical thyrotoxicosis; HI - healthy individuals; AVROT - atrioventricular reciprocal orthodromic tachycardia; AVRNT - atrioventricular reciprocal nodal tachycardia; n - number of patients.

As can be seen from the data obtained (Table 1), the unified mechanism of arrhythmia is preserved in all patient groups. The most frequent triggering factor of AF is extrasystole (60.5-70.0% of cases). More rarely, the appearance of AF has a more complex mechanism. Initially, the extrasystoles provokes a short AVRNT (14.7-21.2%) or AVROT (13.3-22.9%), which soon turns into AF.

The number of extrasystoles and paroxysms of tachycardia in the combination of asymptomatic AF with CAD and ST, symptomatic AF with CAD and ST, asymptomatic and symptomatic AF in patients with CAD without ST, ST, and in healthy individuals is presented in Table. 2.

As an analysis result of the total extrasystoles in patients with AF, subclinical thyrotoxicosis and healthy individuals, it was found that in patients with asymptomatic AF (group 3) their number is greater than in symptomatic AF (group 4) by 33.0% ( $p < 0.001$ ). The combination of asymptomatic AF and subclinical thyrotoxicosis (group 1) is accompanied by an increase in the number of extrasystoles by 21.2% ( $p = 0.007$ ). In patients with ST (group 5), the number of extrasystoles is greater than in HI (group 6) by 39.0% ( $p < 0.001$ ).

It was also found that the paroxysms of tachycardia are more frequent in patients with asymptomatic AF (group 3) 11.0% ( $p = 0.166$ ), and the addition of ST (group 1) to

TABLE 2: The number of extrasystoles and paroxysms of tachycardia in patients with AF, subclinical thyrotoxicosis, and healthy individuals (M ± m).

Species	AFas + CAD + ST (n = 38)	AFs + CAD + ST (n = 35)	AFas + CAD (n = 34)	AFs + CAD (n = 32)	ST (n = 33)	HI (n = 30)
	Group 1	Group 2	Group 3	Group 4	Group 5	Group 6
Total SVE	1373 ± 74,2	1098,4 ± 62,4 p <sub>1-2</sub> = 0,009	1241,5 ± 70,5 p <sub>1-3</sub> = 0,199 p <sub>2-3</sub> = 0,134	777 ± 45,8 p <sub>1-4</sub> < 0,001 p <sub>2-4</sub> < 0,001 p <sub>3-4</sub> < 0,001	715,4 ± 42,2 p <sub>1-5</sub> < 0,001 p <sub>2-5</sub> < 0,001 p <sub>3-5</sub> < 0,001 p <sub>4-5</sub> = 0,345	463,7 ± 28,2 p <sub>1-6</sub> < 0,001 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> < 0,001 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> < 0,001
Total VE	1020 ± 55,1	786,7 ± 44,6 p <sub>1-2</sub> = 0,005	926,2 ± 52,6 p <sub>1-3</sub> = 0,225 p <sub>2-3</sub> = 0,045	676,4 ± 40,1 p <sub>1-4</sub> < 0,001 p <sub>2-4</sub> = 0,069 p <sub>3-4</sub> < 0,001	434,1 ± 25,6 p <sub>1-5</sub> < 0,001 p <sub>2-5</sub> < 0,001 p <sub>3-5</sub> < 0,001 p <sub>4-5</sub> < 0,001	237,2 ± 14,5 p <sub>1-6</sub> < 0,001 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> < 0,001 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> < 0,001
Total SVE and VE	2393 ± 129,4	1885,1 ± 107 p <sub>1-2</sub> = 0,007	2167,7 ± 123 p <sub>1-3</sub> = 0,21 p <sub>2-3</sub> = 0,086	1453,4 ± 85,7 p <sub>1-4</sub> < 0,001 p <sub>2-4</sub> = 0,006 p <sub>3-4</sub> < 0,001	1149,5 ± 67,8 p <sub>1-5</sub> < 0,001 p <sub>2-5</sub> < 0,001 p <sub>3-5</sub> < 0,001 p <sub>4-5</sub> = 0,01	700,9 ± 42,6 p <sub>1-6</sub> < 0,001 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> < 0,001 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> < 0,001
AF amount (per year)	18,1 ± 1,0	14,3 ± 0,8 p <sub>1-2</sub> = 0,007	15,4 ± 0,9 p <sub>1-3</sub> = 0,042 p <sub>2-3</sub> = 0,384	13,7 ± 0,8 p <sub>1-4</sub> = 0,003 p <sub>2-4</sub> = 0,575 p <sub>3-4</sub> = 0,166	7,6 ± 0,4 p <sub>1-5</sub> < 0,001 p <sub>2-5</sub> < 0,001 p <sub>3-5</sub> < 0,001 p <sub>4-5</sub> < 0,001	3,8 ± 0,2 p <sub>1-6</sub> < 0,001 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> < 0,001 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> < 0,001
AF duration (min)	24,5 ± 1,3	18,3 ± 1 p <sub>1-2</sub> = 0,0003	22,2 ± 1,3 p <sub>1-3</sub> = 0,212 p <sub>2-3</sub> = 0,017	16,2 ± 1 p <sub>1-4</sub> < 0,001 p <sub>2-4</sub> = 0,144 p <sub>3-4</sub> < 0,001	2,6 ± 0,2 p <sub>1-5</sub> < 0,001 p <sub>2-5</sub> < 0,001 p <sub>3-5</sub> < 0,001 p <sub>4-5</sub> < 0,001	1,2 ± 0,1 p <sub>1-6</sub> < 0,001 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> < 0,001 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> < 0,001
AF HR (bpm)	146,5 ± 7,9	152,6 ± 8,6 p <sub>1-2</sub> = 0,575	145,3 ± 8,3 p <sub>1-3</sub> = 0,781 p <sub>2-3</sub> = 0,531	151,4 ± 8,9 p <sub>1-4</sub> = 0,63 p <sub>2-4</sub> = 0,785 p <sub>3-4</sub> = 0,585	154,5 ± 9,1 p <sub>1-5</sub> = 0,505 p <sub>2-5</sub> = 0,758 p <sub>3-5</sub> = 0,463 p <sub>4-5</sub> = 0,712	140,4 ± 8,5 p <sub>1-6</sub> = 0,574 p <sub>2-6</sub> = 0,336 p <sub>3-6</sub> = 0,629 p <sub>4-6</sub> = 0,393 p <sub>5-6</sub> = 0,275
Specific symptoms (in points)	0,5 ± 0,027	0,72 ± 0,041 p <sub>1-2</sub> < 0,001	0,51 ± 0,03 p <sub>1-3</sub> = 0,706 p <sub>2-3</sub> < 0,001	0,7 ± 0,04 p <sub>1-4</sub> < 0,001 p <sub>2-4</sub> = 0,661 p <sub>3-4</sub> < 0,001	0,34 ± 0,02 p <sub>1-5</sub> < 0,001 p <sub>2-5</sub> < 0,001 p <sub>3-5</sub> < 0,001 p <sub>4-5</sub> < 0,001	0,22 ± 0,02 p <sub>1-6</sub> < 0,001 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> < 0,001 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> < 0,001
Nonspecific symptoms (in points)	1,5 ± 0,081	1,28 ± 0,073 p <sub>1-2</sub> = 0,046	1,6 ± 0,09 p <sub>1-3</sub> = 0,42 p <sub>2-3</sub> = 0,054	1,3 ± 0,08 p <sub>1-4</sub> = 0,076 p <sub>2-4</sub> = 0,739 p <sub>3-4</sub> = 0,087	1,66 ± 0,098 p <sub>1-5</sub> = 0,213 p <sub>2-5</sub> = 0,006 p <sub>3-5</sub> = 0,231 p <sub>4-5</sub> = 0,008	1,8 ± 0,11 p <sub>1-6</sub> = 0,03 p <sub>2-6</sub> < 0,001 p <sub>3-6</sub> = 0,035 p <sub>4-6</sub> < 0,001 p <sub>5-6</sub> = 0,362

**Abbreviation:** AF – atrial fibrillation; AFas – asymptomatic atrial fibrillation; AFs – symptomatic atrial fibrillation; CAD – coronary artery disease; ST – subclinical thyrotoxicosis; HI – healthy individuals; SVE – supraventricular extrasystoles; VE – ventricular extrasystoles; HR – heart rate; n – number of patients; M – arithmetic mean; m – standard mean error.

them is accompanied by an increase in the number of arrhythmia by 21.1% ( $p = 0.007$ ). It should be noted that in patients with ST (group 5), the number of paroxysms of AF is greater than in HI (group 6) by 50.0% ( $p < 0.001$ ).

The duration of spontaneous paroxysms in patients with asymptomatic AF (group 3) is greater than in symptomatic (group 4) by 37.0% ( $p < 0.001$ ). It is important to note that as a result of the combination of CAD with ST, the duration of asymptomatic paroxysms of AF (group 1) increased by 33.8% compared to symptomatic (group 2) ( $p = 0.003$ ). The duration of spontaneous AF paroxysms in ST (group 5) is more by 53.9% ( $p < 0.001$ ) than in HI (group 6).

As a result of the analysis of AF clinical manifestations by the points scale, it was established that, in symptomatic AF (group 4) the number of specific symptoms is greater by 37,2% ( $p < 0,001$ ), that in asymptomatic AF (group 3); and nonspecific symptoms is less by 23,0% ( $p = 0.087$ ). A similar pattern persisted in the combination of ST with asymptomatic and symptomatic AF (groups 1 and 2). Specific symptoms are more pronounced in combination of ST with symptomatic AF (group 2) by 44.0% ( $p < 0.001$ ), and non-specific – with asymptomatic (group 1) by 17.1% ( $p = 0.046$ ).

## 4. Conclusions

1. Paroxysms of atrial fibrillation in patients with subclinical thyrotoxicosis and in healthy individuals are unstable.
2. In patients with coronary artery disease, subclinical thyrotoxicosis, healthy individuals, the triggering factors of atrial fibrillation are presented by extrasystoles, a combination of extrasystoles with paroxysms of reciprocal atrioventricular orthodontic and nodal tachycardia.
3. Triggering extrasystoles in CAD, subclinical thyrotoxicosis and in healthy individuals have different amounts, but the same structure.
4. The frequency and duration of asymptomatic paroxysms of atrial fibrillation is superior to symptomatic.
5. The combination of subclinical thyrotoxicosis with CAD is accompanied by an increase in the number and duration of asymptomatic and symptomatic paroxysms of atrial fibrillation.

## References

- [1] A. Ceornodolea, R. Bal, J. Severens, "Epidemiology and Management of Atrial Fibrillation and Stroke: Review of Data from Four European Countries", *Stroke Res Treat*, vol. 2017:8593207, 2017.
- [2] *Vserossiyskoe nauchnoe obshchestvo spetsialistov po klinicheskoy elektrofiziologii, aritmologii i kardiostimulyatsii (VNOA)* [All-Russian Scientific Society of Specialists in Clinical Electrophysiology, Arrhythmology and Cardiac Pacing], "Klinicheskie rekomendatsii po provedeniyu elektrofiziologicheskikh issledovaniy, kateternoy ablyatsii i primeneniyu implantiruemykh antiaritmicheskikh ustroystv (redaktsiya 2011)" ["Clinical guidelines on electrophysiological assessment, catheter ablation and the use of implanted antiarrhythmic devices"], Moscow, 518 p, 2011 (in Russian).
- [3] Y. Nakajima, M. Yamada, M. Akuzawa, et al, "Subclinical hypothyroidism and indices for metabolic syndrome in Japanese women: one-year followup study", *J Clin Endocrinol Metab*, vol. 98, pp. 3280-3287, 2013.
- [4] P. W. Rosario, "Natural history of subclinical hyperthyroidism in elderly patients with TSH between 0.1 and 0.4 mIU/l: a prospective study", *Clin Endocrinol (Oxf)*, vol. 72, pp. 685-688, 2010.
- [5] L. Somwaru, A. Arnold, N. Joshi, L. Fried, A. Cappola, "High frequency of and factors associated with thyroid hormone over-replacement and under-replacement in men and women aged 65 and over", *J Clin Endocrinol Metab*, vol. 94, pp. 1342-1345, 2009.
- [6] N. Tribulova, V. Knezl, A. Shainberg, et al, "Thyroid hormones and cardiac arrhythmias", *Vascul Pharmacol*, vol. 52, pp. 102-112, 2010.
- [7] O. Yonem, H. Dokmetas, S. Aslan, T. Erselcan, "Is antithyroid treatment really relevant for young patients with subclinical hyperthyroidism?", *Endocr J*, vol. 49, pp. 307-314, 2002.
- [8] P. Osuna, M. Udovcic, M. Sharma, "Hyperthyroidism and the Heart", *Methodist Debakey Cardiovasc J*, vol. 13(2), pp. 60-63, 2017.
- [9] F. K. Rakhmatullof, S. V. Klimova, A. M. Kuryaeva, N. E. Dyatlov, et al, "*Pokazateli provodjashhej sistemy serdca u zhenshhin s bessimptomnymi paroksizmami*" ["Indices of the conductive system of heart in women with asymptomatic paroxysms"], *Izvestija vysshih uchebnyh zavedenij. Povolzhskij region. Medicinskie nauki* [University proceedings. Volga region. Med Sci], vol. 1(33), pp. 78-87, 2015 (in Russian).
- [10] N. E. Dyatlov, F. K. Rakhmatullof, A. M. Kuryaeva, et al, "*Vzaimosvjaz' mezhdu srokom beremennosti i sostojaniem provodjashhej sistemy serdca pri simptomnoj mercatel'noj aritmii*" ["The relationship between gestation term and the condition

of the heart conduction system in symptomatic atrial fibrillation”], *Izvestija vysshih uchebnyh zavedenij. Povolzhskij region. Medicinskie nauki* [University proceedings. Volga region. Med Sci], vol. 1(37), pp. 54-62, 2016 (in Russian).