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Long-term outcomes of catheter ablation pulmonary veins on example of a clinical case patient with paroxysmal atrial fibrillation

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INTRODUCTION

- Despite good progress in the management of patients with atrial fibrillation (AF), this arrhythmia remains one of the major causes of stroke, heart failure, sudden death, and cardiovascular morbidity in the world
- Since the initial description of triggers in the pulmonary veins that initiate paroxysmal AF, catheter ablation (CA) of AF has developed from a specialized, experimental procedure into a common treatment to prevent recurrent AF
- As first-line treatment for paroxysmal AF, randomized trials showed only modestly improved rhythm outcome with CA compared to antiarrhythmic drug therapy

RADIOFREQUENCY CATHETER ABLATION (RFA) IN AF 1.2.

Complete pulmonary vein isolation (PVI) on an atrial level is the best documented target for catheter ablation, achievable by point-by-point radiofrequency ablation, linear lesions encircling the pulmonary veins

PVI was initially tested in Catheters patients with paroxysmal AF, but appears to be noninferior to more extensive ablation in persistent AF as well



RADIOFREQUENCY CATHETER ABLATION (RFA) IN AF 2.2.

Left superior pulmonary vein

Right superior pulmonary vein

Left inferior pulmonary vein Right inferior pulmonary vein Electroanatomical image of the left atrium and pulmonary veins (posterior view) showing lesions (red circles) for antral pulmonary vein isolation during catheter ablation of atrial fibrillation

CATHETER ABLATION OF PULMONARY VEINS VIDEO



https://www.youtube.com/watch?v= M3BBzmhL4Fc

COMPLICATIONS RELATED TO CA OF AF

| Complication severity | Complication type | Rate ^{727, 748} 750, 754-759 |
|-----------------------|--|--|
| Life-threatening | Periprocedural death | <0.2% |
| complications | Oesophageal injury (perforation/fistula)ª | <0.5% |
| | Periprocedural stroke (including TIA/air embolism) | <1% |
| | Cardiac tamponade | I-2% |
| Severe complications | Pulmonary vein stenosis | <1% |
| | Persistent phrenic nerve palsy | I-2% |
| | Vascular complications | 2-4% |
| | Other severe complications | ≈1% |
| Other moderate or mi | nor complications | I-2% |
| Unknown significance | Asymptomatic cerebral embolism (silent stroke) ^b | 5-20% |
| | Radiation exposure | |

COMBINING ANTIARRHYTHMIC DRUGS AND CATHETER ABLATION

- Antiarrhythmic drug therapy is commonly given for 8–12 weeks after ablation to reduce early recurrences of AF after catheter ablation, supported by a recent controlled trial where amiodarone halved early AF recurrences compared with placebo
- Prospective studies have not been done, but a meta-analysis of the available (weak) evidence suggests slightly better prevention of recurrent AF in patients treated with antiarrhythmic drugs after catheter ablation
- Many patients are treated with antiarrhythmic drug therapy after catheter ablation (most often amiodarone or flecainide), and this seems a reasonable option in patients with recurrent AF after ablation
- The clinical case described below shows long-term outcomes of CA pulmonary veins of the patient with paroxysmal atrial fibrillation

OUR PATIENT

- A 66 years old female (25.08.1950)
- Retired
- Lives in Kharkiv
- Admitted to our polyclinic 14/10/2016

COMPLAINTS

Dyspnea during ordinary physical activity, absent at rest

MEDICAL HISTORY 1.2.

- 2000 autoimmune thyroiditis III degree with nodular goiter, euthyroid state; right – sided thyroidectomy and isthmus resection; according patient's report euthyroidism all the time
- Since 2001- hypertension (max 220/110 mmHg, usual BP 150/90 mmHg (with drugs))
- 2010- paroxysmal tachycardia and palpitations, AF was first diagnosed
- Since 2012 dignosis: <u>PAROXYSMAL ATRIAL FIBRILLATION, EHRA III.</u> <u>CHA2DS2-VASc - 2. HAS-BLED score - 2. ESSENTIAL ARTERIAL</u> <u>HYPERTENSION STAGE II, 3 GRADE. HYPERTENSIVE HEART (LVH).</u> <u>HEART FAILURE WITH PRESERVED EJECTION FRACTION</u>
- 2014- catheter ablation surgery of pulmonary veins
- After 3 days of ablation the patient had a paroxysm of AF- an electrical cardioversion was performed, continued to intake prescribed anthyarritmic treatment for 3 months (betaxolol 10 mg/day, propafenon 300 mg/day)

MEDICAL HISTORY 2.2.

- Despite of drug intaking, ones in 3 weeks she had episodes of AF which were being stopped by intaking additional 300 mg of propafenon
- After 3 months paroxysms of AF became more infrequent (once in 3 months), shorter duration (1-2 hours), stopped after intaking propafenon 300 mg with mild/moderate symptoms of paroxysms of AF
- 2015 gross hematuria on warfarin (the drug intaking was stopped); since 2015 takes aspirin for prevention thromboembolic complications
- Following months (over 3) notes poor control of BP (increasing of it despite taking hypotension drugs)
- After 8 months to the present day of CA she started suffer from paroxysmal tachycardias and heart palpitations with HR 120-130 bpm with mild symptoms, which are not related to physical exercise (mostly at night) 1 time per month, sometimes related to increasement of blood pressure (BP) with duration from 1-2 min to 6 hours and converted to sinus rhythm by taking additional propafenon 300 mg and sometimes procainamide 500 mg

HISTORY OF DISEASES

- 1981- appendectomy
- 1993 acute pyelonephritis
- 2004 radical hysterectomy, iatrogenic menopause
- 2007 cyst in the right breast was removed

OBJECTIVE SUBJECT 1.2.

- The general condition is satisfactory, consciousness is clear, emotionally stable, optimistic mood
- Hypersthenic, height 174 cm, weight 105kg, BMI = 34.68 kg / m 2, waist-to-hip ratio 1,07
- Skin, visible mucous membranes are pale pink and clean
- Peripheral lymph nodes are not palpable
- The thyroid is not palpable in the right side, slightly in the left
- Signs of eyelid retraction, periorbital edema, proptosis are absent

OBJECTIVE SUBJECT 2.2.

 Respiratory System: pulmonary percussion –normal

auscultation - weakened vesicular breathing, no adventitious sounds

Cardiovascular system:

heart borders extended to the left on 1,5 cm of midclavicular line, HR =78 bpm, regular. no pulse deficiency; heart sounds are muted

- BP left hand = BPsin= 175/100 mmHg (on the background of antihypertensive therapy), right hand = BPdex= 150/90 mmHg
- Gastrointestinal system:

abdomen is soft, painless, symmetrical, no discrepancies of the abdominal muscles, no visible peristalsis

liver edge is smooth, painless, palpated 1.5 cm below the costal arch

spleen and pancreas are not palpable

Symmetrical mild shin pitting edema

Recommendations for diagnostic workup of atrial fibrillation patients

| Recommendations | Class ^a | Level ^b | Refc |
|---|--------------------|--------------------|------|
| ECG documentation is required to establish the diagnosis of AF. | | в | 349 |
| A full cardiovascular evaluation, including an accurate history, careful clinical examination, and assessment of concomitant conditions, is recommended in all AF patients. | | C | |
| Transthoracic echocardiography is recommended in all AF patients to guide management. | | C | 339 |
| Long-term ECG monitoring should be considered in selected patients to assess the adequacy of rate control in symptomatic patients and to relate symptoms with AF episodes. | lla | C | |

AF = atrial fibrillation; ECG = electrocardiogram.

^aClass of recommendation.

^bLevel of evidence.

^cReference(s) supporting recommendations.

2016 ESC Guidelines for the management of atrial fibrillation developed in collaboration with EACTS

PRESCRIBED EXAMINATIONS

- Complete blood test
- General urine test
- Fasting glucose level
- ECG
- EchoCG
- 24-h ambulatory ECG monitoring
- Biochemical blood test (liver (ALT, AST, AP)* and renal function tests (BUN*, creatinine), coagulogram* *not peformed examinations due to socio-economic problems

- Blood lipid spectrum
- Blood electrolytes (K, Na)*
- ► TSH, T4, TPO AB*
- Chest X-Ray*
- Ultrasound of thyroid gland*
- Ultrasound of abdomen*

COMMON BLOOD TEST (16/10/2016)

| Measure | Result | Rate | Measure | Result | Rate |
|------------------|--------|-----------------|--------------------------|--------|-----------|
| Hemoglobin | 137 | F 120-158 g / l | Segmented Neutrophils | 59 | 47-72 % |
| Erythrocytes | 4.52 | F 4.0 – 5.2 T/L | Eosinophils | 2 | 0.5-5.0% |
| Color index | 1.06 | 0.85-1.15 | Basophils | 0 | 1-1.0 % |
| Leukocytes | 8.3 | 4.0 – 9.0 g/L | Monocytes | 4 | 3-11% |
| ESR | 10 | F 2-20 mm/h | Lymphocytes | 33 | 19-37% |
| Platelets | 285 | 160-320 g/L | Hematocrit | 39 | F 35-44 % |
| Band Neutrophils | 1 | 1-6% | <u>Conclusion : norm</u> | nal | |

GENERAL URINE TEST (16/10/16)

| MEASURE | RESULT | NORMAL RANGE | |
|--------------------------|--------------|----------------|--|
| SPECIFIC GRAVITY | 1.014 | 1.001-1.040 | |
| REACTION | 6.4 | 5.0-7.0 | |
| PROTEIN | 0.021 | to 0.033 g / I | |
| GLUCOSE | Absent | Absent | |
| LEUCOCYTES | 2-3 | 6-8 | |
| EPITHELIUM TRANSITION | Not detected | Not detected | |
| BACTERIA | Not detected | Not detected | |
| | | | |

Conclusion: normal

BIOCHEMICAL BLOOD TEST (16/10/16)

| MEASURE | RESULT | NORMAL RANGE | |
|--|-------------------|---------------------------------|--|
| Creatinine | 92 | 53-123 mcmol/L | |
| Conclusion: normal | | | |
| ESTIMATED GFR | | NORMAL RANGE | |
| | | | |
| The Modification of Diet in Renal Disease (MDRD) Study equation | 54 ml/min/1.73 m2 | Age (60–69) - 85 ml/min/1.73 m2 | |
| GFR (mL/min/1.73 m²) = 175 × (S _{cr}) ^{-1.154} × (Age) ^{-0.203} × (0.742 if female) × (1.212 if African American) | | | |

<u>Conclusion:</u> decreased kidney function



THYROID-STIMULATING HORMONE (TSH) (16/10/16)



Conclusion: normal

BLOOD LIPID SPECTRUM (16/10/16)

| MEASURE | RESULT | RATE |
|-------------------------------|--------|----------------|
| TOTAL CHOLESTEROL | 7.34 | ≤ 5.2 mmol / I |
| VLDL | 0.97 | <1.0 mmol / I |
| LDL | 5.29 | <3.5 mmol / I |
| HDL- cholesterol levels | 1.08 | >0.9 mmol / I |
| Triglycerides | 2.12 | ≤2.3 mmol / I |
| COEFFICIENT of atherogenicity | 5.8 | To 3.0 mmol/l |

Conclusion: dyslipidemia

ECG with paroxysm of AF before CA



Conclusion: paroxysm of AF with HR 115-125 bpm

ECG with paroxysm of AF 2 years after CA



<u>Conclusion:</u> paroxysm of AF with HR 90-110 bpm. Violation of the repolarization processes on the left ventricular posterior wall (also artifacts on the ECG)

ECG 2 years after CA



<u>Conclusion:</u> sinus rhytm, regular, heart rate 78bpm, signs of left ventricular hypertrophy

24-h AMBULATORY ECG MONITORING 2 YEARS AFTER CA 1.2.

Conclusion :

- During the monitoring 22 h 38 min was registered sinus rhythm with a mean heart rate 74 bpm (maximum HR 120 pm, at 20:05:15, minimum HR 66 bpm - 16:50:55)
- Was recorded: single supraventricular premature contractions (total 266); single monomorphic ventricular premature contractions (total 49); short episodes of supraventricular tachyarrhythmias (total 4) with an average heart rate of 160 bpm with max duration for up to 5 seconds
- Ischemic changes have not been identified
- Circadian index 1.07 (N 1.24-1.44)

24-h AMBULATORY ECG MONITORING 2 YEARS AFTER CA 2.2. :

Episodes of short supraventricular tachyarrhythmia

Парные и групповые НЖЭ. Всего парных и групповых наджелудочковых экстрасистол 3 (днем: 2, ночью: 1).



HEART RATE VARIABILITY 2 YEARS AFTER CA



Conclusion:

- This character of the rhythmogram and HRV indicates the structure to stabilize the heart rhythm with the transition of its regulation from the reflex autonomic level to a lower humoral-metabolic, are not able to quickly provide homeostasis
- Functional heart capabilities are reduced
- Condition of a poor adaptation with a sharp decline in the functional capacity of the body



ECHOCARDIOGRAPHY 1.2.

| Name | Result before ablation | Result 2 years after ablation | Normal |
|----------------------------|---------------------------|-------------------------------------|---------------|
| 1) Acoustic window | normal | normal | normal |
| 2) Aorta | 34 | 34 | 20-37 mm |
| 3) Aortic Valve | 17.5 | 16 | 17-26 mm |
| 4) Left Atrium | 44 | 42 | To 38 mm |
| 5) Mitral Valve | Regurgitation I degree | Regurgitation I degree | |
| 6)Posterior wall of the LV | 13 | 12 | 6-11 mm |
| 7) Interventricular septum | 13 | 12 | 6-11 mm |
| 8) Right Ventricle | 26 | 26 | D.: (9-26 mm) |

ECHOCARDIOGRAPHY 2.2.

| Name | Result before ablation | Result 2 years after ablation | Normal |
|-----------------------|---------------------------|-------------------------------------|--------|
| 9) Right Atrium | 35 | 35 | <38 mm |
| 10) Tricuspid Valve | | | |
| 11) Ejection Fraction | 52 | 57 | 55-78% |

<u>Conclusion 2 years after ablation:</u> Atherosclerosis of aorta and aortic valves mild degree. Moderate dilatation of left atrium. Concentric left ventricle hypertrophy (LV Mass Index 100 g/m². RWT 0.49). Dyssynergic areas were not identified. Diastolic function - relaxation violation (E/A-0.8)

BASIC CLINICAL SYNDROMES

- Atherosclerosis (sclerotic changes of aorta and aortic valve)
- Arterial hypertension
- Arrhythmias (paroxysmal AF)
- Reduction of circadian index and heart spectrum, as a manifestation of reducing humoral and autonomic regulation with non-dipper HR
- Heart failure
- Dyslipidemia
- Hypertensive heart (LVH, atrial enlargement, diastolic dysfunction)
- Obesity

The clinical diagnosis according to current classifications

TYPES OF AF ACCORDING GUIDELINES

| AF pattern | Definition |
|--------------------------------|---|
| First diagnosed AF | AF that has not been diagnosed before, irrespective of the duration of the arrhythmia or the presence and severity of AF-related symptoms. |
| Paroxysmal AF | Self-terminating, in most cases within 48 hours. Some AF paroxysms may continue for up to 7 days. AF episodes that are cardioverted within 7 days should be considered paroxysmal. |
| Persistent AF | AF that lasts longer than 7 days, including episodes that are terminated by cardioversion, either with drugs or by direct current cardioversion, after 7 days or more. |
| Long-standing persistent AF | Continuous AF lasting for ≥1 year when it is decided to adopt a rhythm control strategy. |
| Permanent AF | AF that is accepted by the patient (and physician). Hence, rhythm control interventions are, by AF. Should a rhythm control strategy be adopted, the arrhythmia would be re-classified as 'long-standing definition, not pursued in patients with permanent persistent AF'. |

MODIFIED EUROPEAN HEART RHYTHM ASSOCIATION SYMPTOM SCALE (MODIFIED FROM WYNN ET AL)

| Modified EHRA score | Symptoms | Description |
|------------------------|-----------|---|
| | None | AF does not cause any symptoms |
| 2a | Mild | Normal daily activity not affected by symptoms related to AF ^a |
| 2Ь | Moderate | Normal daily activity not affected by symptoms related to AF, but patient troubled by symptoms ^a |
| 3 | Severe | Normal daily activity affected by symptoms related to AF |
| 4 | Disabling | Normal daily activity discontinued |

AF = atrial fibrillation; EHRA = European Heart Rhythm Association. ^aEHRA class 2a and 2b can be differentiated by evaluating whether patients are functionally affected by their AF symptoms. AF-related symptoms are most commonly fatigue/tiredness and exertional shortness of breath, or less frequently palpitations and chest pain.^{42,194,200–202}

CHA2-DS2-VASC RATING SCALE RISK OF THROMBOEMBOLIC COMPLICATIONS IN PATIENTS WITH ATRIALFIBRILLATION / FLUTTER

| | RISK FACTOR | POINTS | |
|---|--|--------|--|
| С | Congestive heart failure/left ventricular dysfunction | 1 | |
| Н | Hypertension | 1 | |
| A2 | Age ≥75 years | 2 | |
| D | Diabetes mellitus | 1 | |
| S2 | Stroke/transient ischaemic attack/thromboembolism | 2 | |
| V | Vascular disease (prior myocardial infarction, peripheral artery disease, aortic plaque) | 1 | |
| Α | Age 65–74 years | 1 | |
| SC | Sex category (i.e. female gender) | 1 | |
| Conclusion: total points 5. The expected frequency of strokes per year 6.7% | | | |

SCALE HAS-BLED: RISK FACTORS FOR BLEEDING (ESC GUIDELINES FOR THE MANAGEMENT OF ATRIAL FIBRILLATION, 2011)

| | Condition | Points |
|---|--|--------|
| Н | Hypertension: (uncontrolled, >160 mmHg systolic) | 1 |
| Α | Abnormal renal function: dialysis, transplant, Cr >2.6 mg/dL or >200 µmol/L Abnormal liver function: Cirrhosis or Bilirubin >2x Normal or AST/ALT/AP >3x Normal | 1 |
| S | Stroke: Prior history of stroke | 1 |
| В | Bleeding: Prior Major Bleeding or Predisposition to Bleeding | 1 |
| L | Labile INR: (Unstable/high INRs), Time in Therapeutic Range < 60% | 1 |
| Е | Elderly: Age > 65 years Medication | 1 |
| D | Prior Alcohol or Drug Usage History (Antiplatelet agents, NSAIDs) | 1 |
| Conclusion: HAS-BLED score- 4. The patient has a HIGH risk of bleeding. The risk of major bleeding within 1 year in patients with atrial expressed as bleeds per 100 patient years: 3.76 – 6.4% | | |

CLASSIFICATION OF OFFICE BLOOD PRESSURE LEVELS (MMHG)

| Category | Systolic | | Diastolic |
|-------------------|----------|--------|-----------|
| Optimal | <120 | and | <80 |
| Normal | 120–129 | and/or | 80–84 |
| High normal | 130–139 | and/or | 85–89 |
| Grade 1 | 140–159 | and/or | 90–99 |
| hypertension | | | |
| Grade 2 | 160–179 | and/or | 100–109 |
| hypertension | | | |
| Grade 3 | >/=180 | and/or | >/=110 |
| hypertension | | | |
| Isolated systolic | >/=140 | and | <90 |
| hypertension | | | |

CLASSIFICATION OF HYPERTENSION STAGES (RECOMMENDATIONS OF THE ASSOCIATION OF CARDIOLOGISTS OF UKRAINE 2008)

| Stage | The degree of target organ damage |
|-------|---|
| I | Objective changes in the target organs are absent |
| I | There is objective evidence of target organ damage without symptoms with their hand or dysfunction:Left ventricular hypertrophy (on ECG, ultrasound, Ro) Generalized narrowing of retinal arteries Microalbuminuria and / or a small increase in serum creatinine (y m 115 - 133 mmol / L at x107 - 124 mmol / I) Carotid artery disease - a thickening of the intima-media> 0.9 mm or the presence of atherosclerotic plaques. |
| III | There is objective evidence of target organ damage with symptoms from their side and impaired heart - myocardial infarction, heart failure II A - III stage; brain - stroke, transient ischemic attack, acute hypertensive encephalopathy, vascular dementia; fundus - hemorrhage and retinal exudates with papilledema the optic nerve or without; kidney - concentration of plasma creatinine in males> 133 umol / L, y Women> 124; vessels - dissecting aortic aneurysm; peripheral arterial occlusion |

THE NEW YORK HEART ASSOCIATION (NYHA) FUNCTIONAL CLASSIFICATION (FUNCTIONAL CAPACITY) OF CHRONIC HEART FAILURE

NYHA Class Level of Clinical Impairment



No limitation of physical activity. Ordinary physical activity does not cause undue breathlessness, fatigue, or palpitations.



Slight limitation of physical activity. Comfortable at rest, but ordinary physical activity results in undue breathlessness, fatigue, or palpitations.



Marked limitation of physical activity. Comfortable at rest, but less than ordinary physical activity results in undue breathlessness, fatigue, or palpitations.



Unable to carry on any physical activity without discomfort. Symptoms at rest can be present. If any physical activity is undertaken, discomfort is increased.

AMERICAN HEART ASSOCIATION HEART FAILURE STAGES

| Class | Objective Assessment |
|-------|---|
| Α | No objective evidence of cardiovascular disease. No symptoms and no limitation in ordinary physical activity. |
| В | Slight limitation of physical activity. Comfortable at rest. Ordinary physical activity results in fatigue, palpitation, dyspnea (shortness of breath). |
| С | Objective evidence of moderately severe cardiovascular disease. Marked limitation in activity due to symptoms, even during less-than-ordinary activity. Comfortable only at rest. |
| D | Objective evidence of severe cardiovascular disease. Severe limitations. Experiences symptoms even while at rest. |

DEFINITION OF HEART FAILURE WITH PRESERVED (HFpEF), MID-RANGE (HFmrEF) AND REDUCED EJECTION FRACTION (HFrEF)

| Type of HF | | HFrEF | HFmrEF | HFpEF |
|------------|---|-------------------------------|---|---|
| | I | Symptoms ± Signs ^a | Symptoms ± Signs ^a | Symptoms ± Signs ^a |
| NIA | 2 | LVEF <40% | LVEF 40-49% | LVEF ≥50% |
| CRITER | 3 | | Elevated levels of natriuretic peptides^b; At least one additional criterion: a. relevant structural heart disease (LVH and/or LAE), b. diastolic dysfunction (for details see Section 4.3.2). | Elevated levels of natriuretic peptides^b; At least one additional criterion: a. relevant structural heart disease (LVH and/or LAE), b. diastolic dysfunction (for details see Section 4.3.2). |

BNP = B-type natriuretic peptide; HF = heart failure; HFmrEF = heart failure with mid-range ejection fraction; HFpEF = heart failure with preserved ejection fraction; HFrE heart failure with reduced ejection fraction; LAE = left atrial enlargement; LVEF = left ventricular ejection fraction; LVH = left ventricular hypertrophy; NT-proBNP = N-tern pro-B type natriuretic peptide.

^aSigns may not be present in the early stages of HF (especially in HFpEF) and in patients treated with diuretics. ^bBNP>35 pg/ml and/or NT-proBNP>125 pg/mL.

CARDIOVASCULAR RISK STRATIFICATION CHART WITH RECOMMENDED FOLLOW-UP FREQUENCY FOR EACH CATEGORY

| | | BI | ood pressure (mr | nHg) | | |
|--------------------------------|---------------------------|---------------------------------------|--|---|---|---------------------------------------|
| Other risk fac OD or diseas | ctors, e | Normal SBP 120-129 or DBP 80-84 | High normal SBP 130-139 or DBP 85-89 | Grade 1 HT SBP 140-159 or DBP 90-99 | Grade 2 HT SBP 160-179 or DBP 100-109 | Grade 3 HT SBP ≥180 or DBP ≥110 |
| No other Risk level | | Average risk | Averago risk | Low added risk | Moderate added risk | High added risk |
| risk factor | Follow up visits /year | | | 2 | 2 | 3.5 |
| 1-2 risk | Risk level | Low added risk | Low added risk | Moderate added risk | Moderate added risk | Very high addet risk |
| factors | Follow up visits /year | 3.5 | 3.5 | 2 | 2 | - |
| 3 or more risk factors, | Risk level | Moderate added risk | High added risk | High added risk | High added risk | Very high added risk |
| MS, OD or Diabetes | Follow up visits /year | 3.5 | 3.5 | 3.5 | 3.5 | 3.5 |
| Established | Risk level | Very high added risk | Very high added risk | Very high added risk | Very high added risk | Very high added risk |
| disease | Follow up visits /year | 3.5 | 3.5 | 3.5 | 3.5 | 3.5 |

LIPOPROTEIN PATTERNS (FREDRICKSON PHENOTYPES)

| Phenotype | Elevated Lipoprotein(s) | Elevated Lipids |
|-----------|-------------------------------|---------------------|
| I | Chylomicrons | TGs |
| lla | LDL | Cholesterol |
| IIb | LDL and VLDL | TGs and cholesterol |
| | VLDL and chylomicron remnants | TGs and cholesterol |
| IV | VLDL | TGs |
| V | Chylomicrons and VLDL | TGs and cholesterol |

LDL = low-density lipoprotein; TGs = triglycerides; VLDL = very-low-density lipoprotein

CLASSIFICATION OF CHRONIC KIDNEY DISEASE (CKD STAGES)

| NKF CKD Stage (USA) | KDIGO GFR Category (International) | Glomerular Filtration Rate (mL/min/1.73 m ²) | Terms |
|------------------------|---------------------------------------|--|--|
| Stage 1 | G1 | ≥90 | Normal or high In the absence of evidence of kidney damage and abnormal urinalysis, neither GFR category G1 nor G2 fulfill the criteria for CKD |
| Stage 2 | G2 | 60–89 | Mildly decreased relative to a young adult level In the absence of kidney damage and abnormal urinalysis, neither GFR category G1 nor G2 fulfill the criteria for CKD |
| Stage 3A | G3a | 45–59 | Mildly to moderately decreased |
| Stage 3B | G3b | 30–44 | Moderately to severely decreased |
| Stage 4 | G4 | 15–29 | Severely decreased |
| Stage 5 | G5 | <15 | Kidney failure |
| Stage 5D | G5 | <15 | Dialysis |
| Stage 5T | G5 | <15 | Kidney transplant |

https://www.google.com.ua/url?sa=t&rct=j&q=&esrc=s&source=web&cd=3&ved=0ahUKEwjF56uN4r_QAhXGVSwKHTfmDl4QF FHS_CKD_GUIDELINES_V7.0.pdf&usg=AFQjCNE5Ww605G9DQnR28wwTya

WAIST TO HIP RATIO

Waist to Hip Circumference Ratio Standards for Men and Women

| | | ÷ | Disease Risk R | elated to Obesity | y |
|-------|----------------|--------|----------------|-------------------|-----------|
| | Age (years) | Low | Moderate | High | Very High |
| MEN | 20-29 | <0.83 | 0.83-0.88 | 0.89-0.94 | >0.94 |
| | 30-39 | <0.84 | 0.84-0.91 | 0.92-0.96 | >0.96 |
| | 40-49 | <0.88 | 0.88-0.95 | 0.96-1.00 | >1.00 |
| | 50-59 | <0.90 | 0.90-0.96 | 0.97-1.02 | >1.02 |
| | 60-69 | <0.91 | 0.91-0.98 | 0.99-1.03 | >1.03 |
| WOMEN | 20-29 | <0.71 | 0.71-0.77 | 0.78-0.82 | >0.82 |
| | 30-39 | <0.72 | 0.72-0.78 | 0.79-0.84 | >0.84 |
| | 40-49 | < 0.73 | 0.73-0.79 | 0.80-0.87 | >0.87 |
| | 50-59 | <0.74 | 0.74-0.81 | 0.82-0.88 | >0.88 |
| | 60-69 | <0.76 | 0.76-0.83 | 0.84-0.90 | >0.90 |

(Adapted from Heyward VH, Stolarcyzk LM: Applied Body Composition Assessment. Champaign IL, Human Kinetics, 1996, p82.)

CLASSIFICATION OF OVERWEIGHT AND OBESITY BMI CLASSIFICATION

| Classification | BMI Category (kg/m ²) | Risk of developing health problems |
|-----------------|--------------------------------------|---------------------------------------|
| Underweight | < 18.5 | Increased |
| Normal Weight | 18.5 - 24.9 | Least |
| Overweight | 25.0 - 29.9 | Increased |
| Obese class I | 30.0 - 34.9 | High |
| Obese class II | 35.0 - 39.9 | Very high |
| Obese class III | >= 40.0 | Extremely high |

COMPLETE MAIN DIAGNOSIS OF OUR PATIENT

Diagnosis before ablation

Diagnosis 2 years after ablation

- PAROXISMAL ATRIAL **FIBRILLATION, EHRA III**
- CHA2DS2-VAS score 2
- HAS-BLED score 2
- ESSENTIAL ARTERIAL **HYPERTENSION STAGE II,3** GRADE
- HYPERTENSIVE HEART (LVH)
- HEART FAILURE WITH PRESERVED EJECTION FRACTION

- CONDITION AFTER CA OF PULMONARY VEINS DUE TO PAROXYSMAL AF (25/04/14), WITH DECREASEMENT IN **FREQUENCY OF PAROXYSMS FROM ONES IN 3 WEEKS TO ONES PER 2 MONTHS**
- CHA2DS2-VAS score 5
- HAS-BLED score 4
- ESSENTIAL ARTERIAL HYPERTENSION STAGE II,3 GRADE
- HYPERTENSIVE HEART (LVH)
- ► HEART FAILURE WITH PRESERVED EJECTION FRACTION II FC, **STAGE B**
- SYSTEMIC ATHEROSCLEROSIS (ATHEROSCLEROSIS OF THE **AORTA AND AORTIC VALVES, DYSLIPIDEMIA II A TYPE AFTER FREDRICKSON**)
- VERY HIGH ADDED TOTAL CV RISK
- AUTOIMMUNE THYROIDITIS (FOCAL? Riedel's? Hashimoto's?)
 CONDITION AFTER RIGHT SIDED THYROIDECTOMY
- CONDITION AFTER RIGHT SIDED THYROIDECTOMY AND ISTHMUS RESECTION (2000), EUTHYROID DEEP DECLINE THE POWER OF ALL BRANCHES AUTONOMIC
- **REGULATION: NON-DIPPER HR WITH LOW DEGREE OF TP**

CO-MORBIDITY OF OUR PATIENT

2 years after ablation

- CKD 3A: HYPERTENSIVE NEPHROPATHY (eGFR 54 ML/MIN/1.73 m2)
- OBESITY I CLASS
- ► NON-ALCOHOLIC FATTY LIVER DISEASE?

TREATMENT

GOAL-BASED FOLLOW-UP

| Category | Intervention | Follow-up aspects | Performance indicator (examples) |
|--|---|--|--|
| Prognostic | Comorbidity control (relevant examples given) | Obesity Arterial hypertension Heart failure Coronary artery disease Diabetes Valvular heart disease | Weight loss Blood pressure control Heart failure therapy and hospitalizations Statin and antiplatelet therapy; revascularization Glycaemic control Valve repair or replacement |
| Prognostic | Anticoagulation | Indication (risk profile; timing, e.g. post-cardioversion). Adherence (NOAC or VKA) and INR (if VKA). NOAC dosing (co-medications; age; weight; renal function). | Stroke Bleeding Mortality |
| Mainly symptomatic Partly prognostic | Rate control | Symptoms Average resting heart rate <110 bpm | Modified EHRA score Heart failure status |
| Symptomatic at present | Rhythm control | Symptoms vs. side effects Exclusion of pro-arrhythmia (PR; QRS; QTc interval) | LV function Exercise capacity Hospitalization Therapy complications |
| Relevant for implementation of therapy and adherence | Patient education and self-care capabilities | Knowledge (about disease; about treatment; about management goals) Capabilities (what to do if) | Adherence to therapy Directed evaluation, preferably based on systematic checklists |
| Relevant for chronic care management | Caregiver involvement | Who? (spouse; GP; home nurse; pharmacist) Clearly spelling out participation roles Knowledge and capabilities | Directed evaluation of task performance (e.g. via patient card) Dispensed medication Log of follow-up visits |

p.m. = beats per minute; mEHRA symptoms scale = modified European Heart Rhythm Association symptoms scale; GP = general practitioner; INR = international normalized tio; LV = left ventricular; NOAC = non-vitamin K antagonist oral anticoagulant; VKA = vitamin K antagonist.

LIFESTYLE MODIFICATION

Intensive weight reduction in addition to the management of other cardiovascular risk factors (in the range of 10–15 kg weight loss achieved), led to fewer AF recurrences and symptoms compared with an approach based on general advice in obese patients with AF

DASH diet

 Control of compliance to medical recommendations



AAD = antiarrhythmic drug;AF = atrial fibrillation; PVI = pulmonary vein isolation.

catheter ablation should target PVI. IA for paroxysmal AF, IIaB for persistent and long-standing persistent AF.

AF surgery may be PVI (e.g. in paroxysmal AF) or maze surgery (e.g. in therapy-refractory or persistent and long-standing persistent AF).

Hybrid therapy involves combination of antiarrhythmic drugs, catheter ablation, and/or AF surgery.

2016 ESC Guidelines for the management of atrial fibrillation developed in collaboration with EACTS

STROKE PREVENTION IN AF



AF = atrial fibrillation; LAA = left atrial appendage; NOAC = non-vitamin K antagonist oral anticoagulant; OAC = oral anticoagulation; VKA = vitamin K antagonist.

ANTICOAGULATION AFTER SUCCESSFUL' CATHETER ABLATION

- In view of the long-term recurrence rates of AF according 2016 guidelines OAC is recommended to continue in AF patients after 'successful' catheter ablation
- Nonetheless, observational data suggest that the stroke risk may be lower after catheter ablation of AF compared with other AF patients
- The ongoing EAST AFNET 4 trial will inform, in a more general way, whether rhythm control therapy can reduce stroke rates in anticoagulated AF patients
- In addition, there seems to be a place for a controlled trial evaluating the termination of OAC therapy at an interval after 'successful' catheter ablation

AF AND HYPERTENSION

Secondary analyses of trials in patients with LVH and hypertension have found that angiotensin receptor blockers (ARBs) (losartan, valsartan) are better in preventing first occurrence of atrial fibrillation than beta-blocker (atenolol) or calcium antagonist (amlodipine) therapy, consistent with similar analyses in patients with heart failure

SUMMARY OF RECOMMENDATIONS ON THERAPEUTIC STRATEGIES IN HYPERTENSIVE PATIENTS WITH HEART

DISEASE 1.2.

| Recommendations | Class * | Level " | Ret | |
|--|---------|----------------------|-----------------------------------|---------------------|
| In hypertensive patients with CHD, a SBP goal <140 mmHg should be considered. | lla | 8 | 141, <mark>2</mark> 65 | - |
| In hypertensive patients with a recent myocardial infarction beta-blockers are recommended. In case of other CHD all antihypertensive agents can be used, but beta-blockers and calcium antagonists are to be preferred, for symptomatic reasons (angina). | | . | 284 | |
| Diuretics, beta-blockers, ACE inhibitors, angiotensin receptor blockers, and/or mineralocorticoid receptor antagonists are recommended in patients with heart failure or severe LV dysfunction to reduce mortality and hospitalization. | | A | 411 | |
| In patients with heart failure and preserved EF, there is no evidence that antihypertensive therapy per se or any particular drug, is beneficial. However, in these patients, as well as in patients with hypertension and systolic dysfunction, lowering SBP to around 140 mmHg should be considered. Treatment guided by relief of symptoms (congestion with diuretics, high heart rate with beta-blockers, etc.) should also be considered | IIa | CG 2013 ESH/ESC G | uidelines for themanagement of ar | rerial hypertension |

SUMMARY OF RECOMMENDATIONS ON THERAPEUTIC STRATEGIES IN HYPERTENSIVE PATIENTS WITH HEART

| DISEASE | | | |
|--|-----|---|-----|
| ACE inhibitors and angiotensin receptor blockers (and beta-blockers and mineralocorticoid receptor antagonists if heart failure coexists) should be considered as antihypertensive agents in patients at risk of new or recurrent atrial fibrillation. | IIa | c | - |
| It is recommended that all patients with LVH receive antihypertensive agents. | | в | 458 |
| In patients with LVH, initiation of treatment with one of the agents that have shown a greater ability to regress LVH should be considered, i.e. ACE inhibitors, angiotensin receptor blockers and calcium antagonists. | lla | в | 580 |

ACE = angiotensin-converting enzyme; CHD = coronary heart disease; EF = ejection fraction; LV = left ventricle; LVH = left ventricular hypertrophy; SBP = systolic blood pressure.

^aClass of recommendation.

^bLevel of evidence.

^cReference(s) supporting recommendation(s).

ORAL ANTIARRYTHMIC DRUGS USED FOR MAINTANING SINUS RHYTHM

| Drug | Dose | Main contra-indications and precautions | Warning signs warranting discontinuation | AV nodal slowing | Suggested ECG monitoring during initiation |
|--|--|--|--|-------------------------------------|--|
| Amiodarone | 600 mg in divided doses for 4 weeks, 400 mg for 4 weeks, then 200 mg once daily | Caution when using concomitant therapy with QT- prolonging drugs and in patients with SAN or AV node and conduction disease. The dose of VKAs and of digitalis should be reduced. Increased risk of myopathy with statins. Caution in patients with pre-existing liver disease. | QT prolongation >500 ms | 10–12 bpm in AF | Baseline, I week, 4 weeks |
| Dronedarone | 400 mg twice daily | Contra-indicated in NYHA Class III or IV or unstable heart failure, during concomitant therapy with QT-prolonging drugs, or powerful CYP3A4 inhibitors (e.g. verapamil, diltiazem, azole antifungal agents), and when CrCl <30 mg/mL. The dose of digitalis, beta-blockers, and of some statins should be reduced. Elevations in serum creatinine of 0.1–0.2 mg/dL are common and do not reflect a decline in renal function. Caution in patients with pre-existing liver disease. | QT prolongation >500 ms | 10–12 bpm in AF | Baseline, I week. |
| Flecainide Flecainide slow release | 100–150 mg twice daily 200 mg once daily | Contra-indicated if CrCl <50 mg/mL, liver disease, IHD or reduced LV ejection fraction. Caution in the presence of SAN or AV node or conduction disease. CYP2D6 inhibitors (e.g. fluoxetine or tricyclic antidepressants) increase plasma concentration. | QRS duration increases >25% above baseline | None | Baseline, day 1, day 2–3 |
| Propafenone Propafenone SR | 150–300 mg three times daily 225–425 mg twice daily | Contra-indicated in IHD or reduced LV ejection fraction. Caution in the presence of SAN or AV node and conduction disease, renal or liver impairment, and asthma. Increases concentration of digitalis and warfarin. | QRS duration increase >25% above baseline | Slight | Baseline, day 1, day 2–3 |
| d,l sotalol | 80–160 mg twice daily | Contra-indicated in the presence of significant LV hypertrophy, systolic heart failure, asthma, pre-existing QT prolongation, hypokalaemia, CrCl<50 mg/mL Moderate renal dysfunction requires careful adaptation of dose. | QT interval >500 ms, QT prolongation by >60 ms upon therapy initiation | Similar to high dose blockers | Baseline, day 1, day 2–3 |

PATIENT'S MEDICAL TREATMEN FOR LAST 6 MONTH

- Bisoprolol 5 mg per day
- Propafenon 150 mg 2 times per day (without this drug –recurrence of AF paroxysms); episodically additionally 300 with/without procainamide 500 mg
- Valsartan 80 mg per day
- Atorvastatin 10 mg (do not intake regularly)
- Aspirin 75 mg per day

OUR RECOMMENDED TREATMENT

- B-blocker CARVEDILOL 12,5 mg 2 times p/day under control of ECG
- AAD PROPAFENONE 150 mg 3 times per day under control of ECG; additional 300 mg of propafenon in case of paroxysm of AF
- ► <u>ARBs</u> VALSARTAN 160 mg in the morning
- Anticoagulant RIVAROXABAN 15 mg p/day
- Statin-ROSUVASTATIN 20 mg in the evening
- Consulting with other subspecials to change treatment strategy (repeat catheter ablation?)

ADDITIONAL RECOMMENDED EXAMINATIONS

- Repeat 24h ECG monitoring in a month
- T4, T3, Anti-TPO
- Biochemical blood test (liver (ALT, AST, AP) and renal function tests (BUN), coagulogram
- Blood electrolytes (K, Na)
- Chest X-Ray
- Ultrasound of thyroid gland and abdomen
- Consultation with an endocrinologist

PROGNOSIS

 Prognosis for life - non-compliance to doctor's appointments - <u>non-satisfactory</u>
 The prognosis for recovery - <u>an</u> unfavorable

CONCLUSION 1.2.

- According to recent studies it has been demonstrated that pulmonary vein <u>CA has</u> <u>favourable outcomes at 6-12 months post-ablation</u>, but there are only few studies with a long-term follow-up and, as we see on our clinical case, after 2 years patient present with current deterioration of AF
- The vast majority of very longstanding paroxysmal/persistent AF patients maintained sinus rhythm at a mean follow-up time of 5 years following CA, associated with a significant improvement in symptom scores and, as we see on our clinical case, after 2 years patient maintained sinus rhythm, but with recurrence paroxysms of AF for last year with mild/moderate of symptom scores
- Often this procedure is not a radical solution of the problem, and <u>most patients</u> (as it also was shown on the example of our clinical case) are <u>require adjunctive therapies including</u> <u>antiarrhythmics, DC cardioversions and re-ablation</u>

CONCLUSION 2.2.

- Also our patient needs correction of the treatment of arterial hypertension and more properly diagnosis (and treatment) of thyroid disorder, and improvement the regulation at all levels - from the daily rhythm of the HR up to relations in the activity of the vagal activity branches, first of all, interventions in the lifestyle and searching for the optimum time drug administration
- Of course, consider the presence of multiple syndromes on presented clinical case, we must not forget about the problem of polypharmacy and try to avoid it (many studies in ambulatory care define polypharmacy as a medication count of five or more medications, but it is practically impossible to investigate the biochemical compatibility in vivo of more than 4 drugs)

THANK YOU FOR ATTENTION!