SUDDEN CARDIAC DEATH DUE TO PHYSICAL EXERCISE IN CROATIA IN A 27-YEAR PERIOD

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Dedicated to the memory of our teacher professor Radovan Medved, MD, PhD, the founder of the Croatian Sports Medicine, a corresponding member of the Croatian Academy of Sciences and Arts.

Summary

In a period of 27 years: from January 1, 1984 to December 31, 2010 we noticed 69 sudden and unexpected cardiac deaths during physical exercise in Croatia. There were 6 sudden cardiac deaths in male athletes. At the autopsy, an athletic runner aged 21, had an acute myocardial infarction with normal coronaries and left ventricular wall thickened. A professional soccer player aged 17, had hypoplastic right coronary artery and narrowed ascending aorta, bacterial tonsillitis and subacute myocarditis. A student rugby player aged 29, and a school boy-basketball player aged 15, had hypertrophic cardiomyopathy. The arrhythmogenic right ventricular dysplasia was a cause of death in two athletes. The first was a short trails runner, and the second was a soccer player.

The hypertrophic cardiomyopathy was a cause of death in 3 athletes and 2 persons during recreational exercise. An acute myopericarditis was a cause of death in 2 professional soccer players, and in one who died during swimming. One had subacute diffuse myopericarditis, bacterial tonsillitis and narrowed ascending aorta of 10 mm. The other had chronic myopericarditis and cardiac aneurysm of the left ventricle. The third
had fibrinous pericarditis, hyperthropic cardiomyopathy, hypoplastic ascending aorta, bilateral bronchopneumonia and cerebral contusion with edema.

Sudden cardiac deaths appeared in 10 school boys. Two of them had been engaged in physical exercise at school, the third was a professional soccer player, the fourth was engaged in recreational swimming, and the fifth had just finished secondary school and was working at the site recreatively. In 3 of them congenital cardiovascular diseases was found: in 2 hypoplastic coronary arteries and in 2 hypertrophic cardiomyopathy. The fourth had normal heart findings including coronaries, but had bilateral pneumonia with a non-cardiogenic pulmonary edema. The fifth had a chronic myopericarditis with an aneurysm of the left ventricle.

Pneumonia was a cause of death in 3 male teenagers aged 18-19. The first was working at the site recreatively, the second was engaged in soccer recreatively and the third was professional soccer player. One died suddenly during physical exercise at the field and 2 died in the hospital. The first had bilateral bacterial pneumonia, non-cardiogenic pulmonary edema and cerebral edema. The second had bilateral bacterial pneumonia, adult respiratory distress syndrome, disseminated intravascular coagulation, cerebral edema, hypoplastic right coronary artery and myocardial fibrosis. The third had bilateral bacterial pneumonia, fibrinous pericarditis, and cerebral contusion with edema, thickening of the left ventricle 20 mm and hypoplastic ascending aorta.

We noticed 5 sudden death among Croatian male physicians, during or after recreational exercise: swimming, soccer, tennis and jogging; 3 were autopsied, and all had coronary heart disease. Two physicians who were not been autopsied, had possible an alcohol cardiomyopathy. Fifteen “elderly” men, died suddenly during exercise: 6 in swimming, 4 in playing tennis, one ridded, one was jogging, 2 bowling and one died during sexual act. At autopsy, 14 had coronary heart disease: 5 critical coronary artery stenosis, and 2 had signs of recent myocardial infarctions, 3 had left descending coronary artery occluded, and one with acute myocardial infarction of the anterior wall, 12 had myocardial scars due to previous myocardial infarctions and one signs of diffuse myocardial fibrosis. Twelve of them had left ventricular hypertrophy.

In Croatia the death rate among athletes reached 0.19/100 000 yearly (p=0.00005), in the male population aged 15-40 years, engaged in sports and recreational physical exercise: 0.71/100 000 (p=0.00001), in the total male population aged 15-64 engaged in sport and recreational exercise 0.96/100 000 (p=0.00001), in arrhythmogenic right ventricular dysplasia reached 0.06/100,000 (p=0.00000), in teenagers suffered of hypertrophic cardiomyopathy 0.10/100 000 (p=0.00000), in myopericarditis it was 0.11/100,000 (p=0.00000), in pneumonia 0.11/100 000 (p=0.00000), in teenagers 0.37/100 000 (p=0.00226), in physician-specialists reached 24.8/100 000 (p=0.00000), in elders reached 2.99/100 000 (p=0.00001).

**Keywords:** men; athletes; recreational physical exercise; sudden death
INTRODUCTION

In trained and functionally fit athletes with a healthy heart, acute cardiovascular incidents due to training are very scarce. Regular physical exercise, controlled and adapted to physical conditions has beneficial effects surpassing hazards. If the principles of physical training are considered, possible complications could be avoided. In persons aged under 30, who died during or immediately after physical exercise, the most common reasons for lethal events is a congenital heart disease: hypertrophic cardiomyopathy, congenital coronary artery anomalies (including an abnormal origin of coronary artery, hypoplastic coronary artery, myocardial bridging), aortic stenosis, aortic dissection (often associated with Marfan’s syndrome), myocarditis - higher than expected [1], an idiopathic dilated cardiomyopathy, mitral valve prolapsed, idiopathic cardiac hypertrophy, infiltrating cardiomiopathies (f.e. amyloidosis), right ventricular dysplasia [2,3], long Q-T interval syndrome, the Brugada syndrome: right bundle branch block, right precordial ST-segment elevation associated with ventricular tachycardia [4]. Risk factors for sudden death include positive familial history for conditions associated with sudden death, recurrent or exercise-induced non-neurogenic syncope, anginal chest pain, palpitations, dyspnea, cardiovascular diseases, seizure activity, and athletic competition [5]. That is why a medical check-up before exercise is essential, as well as medical controls in persons taking exercise [5-9].

Among cardiovascular disease leading to sudden death [8,10-15], arrhythmogenic right ventricular dysplasia or cardiopathy (ARVD/C) by some authors reached 2-5% of all sudden deaths in young persons, with a higher death rate during exercise [16]. ARVD is primary myocardial diseases caused of genetic defect of desmosomes, i.e. area in myocardium which links together the myocardial myofibrils. Those desmosomes are composed of several proteins. In this disease we are facing with changes of cell adhesion proteins: plakoglobin, plakophilin-2, desmoglein-2, desmoplakin [17,18]. Many of those proteins can have harmful mutations. ARVD is very often biventricular disease involving the right ventricular myocardium and typically the subepicardial region of the left ventricle characterized of myocyte vacuolization. It is replaced with fibro fatty tissue in the right ventricle leading to myocardial atrophy and with more fibrosis than fibro fatty infiltration in the left ventricle [16,19]. It could be connected with a high risk of ventricular instability and malignant ventricular arrhythmias. Sudden cardiac death could be the only manifestation of the disease, especially in young athletes [16].

Hypertrophic cardiomyopathy (HCM) is the most common cause of sudden cardiac death in young competitive athletes [20,21]. HCM means thickened and not
dilated left ventricular wall, without any other disease leading to left ventricular hypertrophy (LVH). The etiology of HCM is unknown. Teare [22] was the first who described it, as an asymmetrical heart hypertrophy. HCM is a primary myocardial disease [23] with 450 different mutations identified in 13 genes, in which three genes are responsible for more than 50% of all genotype cases: cardiac troponin-T (chromosome 1), myosin-binding protein C (chromosome 11) and beta-myosin heavy chain (chromosome 14). In HCM, LVH is asymmetric and could be concentric, localized in the basal anterior septum, posterior septum, anterolateral wall, posterior wall or in the apex. The consequence of HCM could be sudden cardiac death due to decrease of a cardiac output leading to dyspnea during exercise, postural arterial hypotension, palpitations or syncope, myocardial ischemia with typical or atypical angina chest pain during exercise and with a high risk of ventricular instability and a malignant ventricular arrhythmias. LVH increases the risk of sudden cardiac death by promoting subendocardial ischemia [24].

Myopericarditis is at the third place among the causes of sudden cardiac death in young persons engaged in sport and recreational exercise and the incidence being higher than expected [7,8,10,25-31]. Risk factors for sudden death associated with exercise include positive familial history for conditions associated with sudden death, cardiovascular diseases and symptoms suggesting them: syncope recurrent, exercise-induced, and non-neurogenic in nature, angina chest pain, palpitations, dyspnea, and seizure activity (5,32-34). Adolescents with these risk factors should be referred to a cardiologist for diagnostic and therapeutic interventions.

Acute respiratory tract infections are the most frequent reasons of morbidity and inability for exercise [7,8,11,17,31]. Their frequency reached over 65 per cent of all infections. Several epidemiological studies point out the increased risk of morbidity from the respiratory tract infections in athletes during the intensive endurance training sessions [13]. Athletes in periods of intensive training and after highly strenous training are more often affected by respiratory tract infections. But in those engaged in recreational physical exercise consider that such regular activities protect them from these infections [7,31,35].

Physicians engaged in physical exercise have specific problems because they very often underestimate their own physical condition. Complications, during or after exercise, might occur both in persons who suffer from heart disease and in those who are „free“ from disease as physician very often are [25,36-41].

Elderly people are specific group with regard to physical exercise, because numerous changes occur in the human organism with advancing age. Elderly people regularly engaged in recreational physical exercise have lower risk for cardiovascular complications than inactive persons [11,42-46].
The aim of this study is to analyze the prevalence of sudden cardiac death in males due or immediately after sports or recreational physical exercise in regard to reasons and other specific consequences in Croatia.

SAMPLE AND METHODS

According to the register of a population from 2001 year, from the Croatian Department of Statistics, in Croatia it were 4,437,460 inhabitants, of which it was 2,135,900 males. There were 1,484,625 males aged 15 to 65 years, and 265,108 males aged >65 years, and 7% of them practicing physical exercise (18,558). In a group of persons aged 15 to 65 years, there were 97,940 school boys and 116,056 athletes, who were practicing physical exercise (213,996). Of other males aged 15 to 65 years (1,270,629), 10% were practicing physical exercise (127,063). So we are coming to the number of 359,617 males older than 15 years, who were practicing physical exercise.

The data presented here are part of a large retrospective study dealing with 69 sudden and unexpected deaths during or immediately after physical exercise in men of all ages (and no women) in a 27-year period in Croatia: from January 1, 1984 to December 31, 2010. All of them were autopsied. The data of deceased persons were found from the Public Health Registry, Sports Clubs and Services of Forensic Medicine in Croatia. The cases of sudden cardiac death who were autopsied are presented according to the age of victims, type of physical activity-exercise-sport during which death occurs as well as in a subgroup of physically active physicians who died suddenly. Special attention is paid to the cases of fatal cardiac events in 6 athletes and 13 teenagers, 15 elderly persons engaged in physical exercise, and in a subgroup of 5 physicians. These cases are presented in detail as well as the cases with underlying HCM in 5 adolescents, ARVD in 2 athletes, myopericarditis in 3, and bronchopneumonia in 3 adolescents. The total prevalence as well as the prevalence of sudden cardiac death in males during or immediately after physical exercise in groups regarding chronological age and with regard to the possible cause of death was calculated as a number of cases per 100,000 persons engaged in such physical activities.

Statistical differences between deceased groups regarding the chronological age and underlying diseases in relation to all male inhabitants engaged in sports and recreational physical exercise in Croatia, was calculated using the Chi-square test and Poisson rates.

RESULTS

In a period of 27 years, 69 sudden cardiac death due to physical exercise were registered. The age of of victims was between 13 to 82 years and all of them were
males. The majority of victims (63.8%) were adults (20 to 65 years of age). Every fifth victim was 65 years of age or more (21.7%) and 14.5% of the total number of cases were adolescents, what is presented in Table 1.

**Table 1.** Number (n) and percent (%) of fatal cardiac incidents in a 27 year period (1984-2010) according to age and the engagement in sport vs. recreational exercise of victims

<table>
<thead>
<tr>
<th>Life period/physical exercise</th>
<th>Age range (in years)</th>
<th>N</th>
<th>Total n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>Athletes</td>
<td>Recreational physical exercise</td>
<td></td>
</tr>
<tr>
<td>Adolescents- sport</td>
<td>13-19</td>
<td>3</td>
<td>7</td>
<td>10</td>
</tr>
<tr>
<td>Adults</td>
<td>20-62</td>
<td>3</td>
<td>41</td>
<td>44</td>
</tr>
<tr>
<td>Elderly-recreational exercise</td>
<td>65-82</td>
<td>15</td>
<td>15</td>
<td></td>
</tr>
<tr>
<td>Total</td>
<td>13-82</td>
<td>6</td>
<td>63</td>
<td>69</td>
</tr>
</tbody>
</table>

The most prevalent sport in these victims was soccer (26.1%). It is followed by swimming, mostly recreational or in a free time: in 23.2% of cases. Jogging was fatal for 8 persons (one teenager, 6 adults- 2 of them were athletes who died suddenly during training, and one elderly person) or 11.6% of cases. Other sports as bowling, diving, cycling, hiking were reported in less than 10%. At the end of the list of sport, recreational physical exercise or physical activity during which death occurred, were basketball [1], exercising at school [1] and army [2], work at a site [1] and sexual activity (1 case).

The characteristics of the male athletes and adolescents (age, profession, sport-type of exercise, reported symptoms, physical findings – ECG, data and occasion of fatal incidents, resuscitation and forensic autopsy findings) who died suddenly during or immediately after training are presented in Table 2.

**Case 1.** An athletic runner, aged 21, had been without any health problems at rest or during physical exercise, and had normal physical findings of the heart and ECG, with normal length of a corrected Q-T interval. He was running on a hot summer afternoon in June 1988, collapsed and died. All reanimation efforts were unsuccessful, including the attempts of the Emergency Mobile Unit. At autopsy, finding showed an acute myocardial infarction of the front wall, normal coronaries, and an acute pulmonary and cerebral edema.

**Case 3.** A former rugby player and actually rugby referee, aged 29, had no symptoms of a heart disease. He did not sleep at least 48 hours before the incident. At that very day in June 1998, he was swimming in the pool (he was a good swimmer), he
Table 2. Male athletes and adolescents died suddenly during physical exercise

<table>
<thead>
<tr>
<th>CASE</th>
<th>AGE</th>
<th>PROFESSION/EXERCISE</th>
<th>SYMPTOMS</th>
<th>PHYSICAL FINDING/ ECG</th>
<th>LETHAL EVENT</th>
<th>RESUSCITATION</th>
<th>FORENSIC AUTOPSY</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>2</td>
<td>student / runner</td>
<td>no</td>
<td>normal finding acute</td>
<td>June 1988, during a run</td>
<td>yes</td>
<td>AMI of the anterior wall, normal CAA, pulmonary and cerebral edema</td>
</tr>
<tr>
<td>2</td>
<td>17</td>
<td>exercised 2-3 times a week</td>
<td>few days before the incident</td>
<td>acute</td>
<td>September 2010 while running on a street</td>
<td></td>
<td>heart enlargement: 390 g, LVH 19 mm, normal thickness of the RV: 3 mm; aortic diameter 5.1 cm, and narrowing of the descending part of the aortic arch</td>
</tr>
<tr>
<td>3</td>
<td>29</td>
<td>student/ rugby player / swimming</td>
<td>no</td>
<td>normal finding</td>
<td>June 1998 in the swimming pool</td>
<td>yes</td>
<td>LVH 25 mm, normal CAA, acute pulmonary edema</td>
</tr>
<tr>
<td>4</td>
<td>15</td>
<td>high-school student / basketball player</td>
<td>no</td>
<td>no data</td>
<td>May 2000 during a game</td>
<td>yes</td>
<td>LVH 17 mm, IV septum 40 mm, narrowed cavity of the LV; enlarged heart: 750 g, dilated aorta: 40 mm, normal CAA, myocardial fibrosis and scars up to 3 mm in diameters, acute pulmonary and cerebral edema</td>
</tr>
<tr>
<td>7</td>
<td>18</td>
<td>high-school student/ soccer player</td>
<td>no</td>
<td>no data</td>
<td>May 2006 kicked in a chest during a game, died in a regional hospital 2 days after admittance</td>
<td>yes</td>
<td>LVH 20 mm, normal cavities of the heart, hypoplastic ascending aorta, normal CAA, fibrinous pericarditis, bilateral broncho-pneumonia, chest contusion, cerebral contusion</td>
</tr>
<tr>
<td>10</td>
<td>17</td>
<td>professional soccer player</td>
<td>no</td>
<td>no data</td>
<td>February 1998 during a game</td>
<td>yes</td>
<td>enlarged heart RCA hypoplastic up to 1 mm, ascending aorta narrowed to 10 mm, subacute diffuse myocarditis, suppurant tonsillitis, acute pulmonary and cerebral edema</td>
</tr>
<tr>
<td>11</td>
<td>18</td>
<td>swimming recreationally</td>
<td>no</td>
<td>no data</td>
<td>June 2002 during swimming</td>
<td>yes</td>
<td>chronic myopericarditis, cardiac with left ventricular fibrosis and myocardial scars, aneurysm of the LV 2 cm in diameter, normal CAA, pulmonary edema</td>
</tr>
</tbody>
</table>
Table 2. Continued

<table>
<thead>
<tr>
<th>CASE</th>
<th>AGE</th>
<th>PROFESSION/ EXERCISE</th>
<th>SYMPTOMS</th>
<th>PHYSICAL FINDING/ ECG</th>
<th>LETHAL EVENT</th>
<th>RESUSCITATION</th>
<th>FORENSIC AUTOPSY</th>
</tr>
</thead>
<tbody>
<tr>
<td>ARVD</td>
<td>5</td>
<td>24</td>
<td>student/ runner</td>
<td>no</td>
<td>no data</td>
<td>January 2002 during training</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>6</td>
<td>13</td>
<td>school boy/ soccer player-cadet</td>
<td>no</td>
<td>no data</td>
<td>September 2007 during training</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>12</td>
<td>14</td>
<td>physical exercise in a school</td>
<td>no</td>
<td>no data</td>
<td>October 2001 during physical exercise</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>13</td>
<td>18</td>
<td>just finished secondary school – working at the site</td>
<td>General tiredness</td>
<td>no data</td>
<td>December 2001 caught a cold and felt exhausted, died suddenly during work</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>15</td>
<td>19</td>
<td>recreate soccer player</td>
<td>no</td>
<td>no data</td>
<td>January 2002 in a hospital 1 day after development of sepsis, shock, DIC, acute renal failure</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>14</td>
<td>19</td>
<td>swimming in the sea</td>
<td>no</td>
<td>no data</td>
<td>June 2009 during swimming</td>
<td>yes</td>
</tr>
</tbody>
</table>

Legend: CAA=coronary arteries; LV=left ventricle; RV=right ventricle; Lvh=left ventricular hypertrophy; Iv=interventricular; RCA=right coronary artery, ARVD=arrhythmogenic right ventricular dysplasia; AMI=acute myocardial infarction; ARDS=adult respiratory distress Syndrome; DIC=disseminated intravascular coagulation; HCM=hypertrophic cardiomyopathy.
felt increased tiredness, stopped to swim for a few minutes, then swimmmed once again, and then felt increased tiredness, collapsed and died. All reanimation efforts were unsuccessful including a Mobile Emergency Medical Unit. At the autopsy he had hypertrophic cardiomyopathy: by the criteria of gross pathological changes of cardiomegaly, left ventricular hypertrophy: 25 mm, with normal cavities of the whole heart and normal coronary arteries. Histological findings of myocardium showed diffuse interstitial and replacement fibrosis, dysplasia of the tunica media of small vessels and contraction band and coagulate necrosis. He had acute pulmonary and cerebral edema.

Case 4. A basketball player-school boy aged 15, during a game in May 2000, he felt exhausted, and was standing at the site, watching the others playing basketball. Then he resumed training, jumped suddenly, put the hand on his chest, collapsed and died. All reanimation attempts were unsuccessful including attempts at the Regional University Hospital. A forensic autopsy finding showed hypertrophic cardiomyopathy with interventricular septum of 40 mm and left ventricular wall of 17 mm. The left ventricle was not dilated. He had an acute pulmonary and cerebral edema.

Two young men: cases 5 and 6, suffered of arrhythmogenic right ventricular dysplasia.

Case 5. A male high degree short trails runner for years, aged 24, referred no cardiovascular or any other symptoms. During an athletic training in the morning in January 2002 he was running due to training and after short time suddenly collapsed and died. He was intubated and ventilated mechanically immediately by a physician at the field, and after that he was resuscitated by a medical team of the Re-animation Unit all the time during transport, to the Regional University Hospital. At the admittance he was unconscious, with anisocoric pupils, with no respirations, no heard heart sounds, with central cyanosis, and with flat line on an ECG monitor all the time during long unsuccessful resuscitation. The clinical diagnoses were: reanimatio facta, haemorrhagia cerebri suspecta, cardiomyopathia hypertrophica suspecta, sanguinatio ex ore, ventilatio mechanica, implantatio electrostimulatori cordis facta, dissociatio electromechanica. At the autopsy, the size of the whole heart was 11 x 11 x 6.5 cm, the left ventricle wall reached 15 mm; both ventricles were dilated, with normal mitral, aortal, tricuspid and pulmonary valves and no signs of coronary or aortal atherosclerosis. Histological finding of the right ventricle wall showed abundant subepicardial accumulation of an adipose tissue with infiltration between myocardial bundles. The finding of the lungs and of the brain showed acute edema. The autopsy diagnoses were: dysplasia arrhythmogenes ventriculi dextri...
cords, dilatatio cordis totius, oedema pulmonum et cerebri acutum, cyanosis universalis.

Case 6. A soccer player as a cadet, aged 13, without any previous physical discomfort, suddenly collapsed and died during soccer training in the middle of January 2002. He was resuscitated at the field by a coach and after that by a medical team of the Reanimation Unit, with no success. At autopsy, the size of the whole heart was 10 x 10 x 5 cm, the left ventricle wall reached 10 mm, the right ventricle wall 3 mm, both ventricles were dilated, with normal mitral, aortal, tricuspid and pulmonary valves and with no signs of coronary or aortal atherosclerosis. Histological finding of the myocardium showed abundant subepicardial accumulation of adipose and fibrotic tissues with segmental lymphocyte infiltration between bundles of both ventricles. The findings of the lungs and the brain showed acute edema. The autopsy diagnoses were: dysplasia arrhythmogenes ventriculi dextri et ventriculi sinistri cordis, oedema cerebri grave, oedema pulmonum, cyanosis universalis.

In athletes died during training or during to competitive game in Croatia, the death rate is 0.19/100 000 yearly, in male population aged 15-40 who practiced physical exercise 0.71/100 000, in total male population aged 15-64 who practice physical exercise reached 0.96/100 000.

The death rate of an arrhythmogenic right ventricular dysplasia what was a cause of death in two athletes reached 0.06/100 000.

Five young men died because of hypertrophic cardiomyopathy, of which cases 3, 4 and 7 are presented in the Table 2, and cases, 8 and 9 in the Table 3. The death rate reached 0.10/100 000.

Case 7 (Table 2). The third athlete aged 18 was a school boy and professional soccer player. During a soccer game in May 2006 one player kicked his head with a ball. He felt general weakness with a short breath and he felt down. He was transferred to the nearest University Hospital where he have been cured because of bilateral bacterial pneumonia, and died two days after admittance in a state of a septic shock, in spite of all therapeutic efforts. The autopsy showed hypoplastic ascending aorta and biventricular hypertrophy. The left ventricular wall reached 20 mm, and normal cavities of the heart. The heart weight was amounting to 450 g. He had a large bilateral bacterial pneumonia, fibrinous pericarditis, cerebral contusions with edema and pointed bleeding.

Case 8 (Table 3) The fourth was a physician, cardiovascular pathologist aged 34, who played soccer twice a week recreationally. He was without any physical discomfort during physical exercise. In the evening in August 1984 during a soccer game, he did not feel well, went to a car and drove few hundred meters, than sto-
Table 3. Male adults with hypertrophic cardiomyopathy and a group of physicians died suddenly during recreational physical exercise

<table>
<thead>
<tr>
<th>CASE</th>
<th>AGE</th>
<th>PROFESSION/EXERCISE</th>
<th>SYMPTOMS</th>
<th>PHYSICAL FINDING/ECG</th>
<th>LETHAL EVENT</th>
<th>RESUSCITATION</th>
<th>FORENSIC AUTOPSY</th>
</tr>
</thead>
<tbody>
<tr>
<td>9</td>
<td>25</td>
<td>soccer recreationally</td>
<td>no</td>
<td>no data</td>
<td>June 2002 during a game</td>
<td>yes</td>
<td>LVH 22 mm, ostium of the RCA narrowed to 1 mm, malposition of the ostium of the RCA, ostium of the LCA narrowed to 1 mm, normal coronary arteries, pulmonary edema</td>
</tr>
<tr>
<td>8</td>
<td>34</td>
<td>physician-pathologist/ soccer recreationally</td>
<td>no</td>
<td>no data</td>
<td>August 1984 during a game</td>
<td>yes</td>
<td>CHD, LVH 18 mm, normal cavities of the whole heart, LADCA narrowed, pulmonary and cerebral edema</td>
</tr>
<tr>
<td>16</td>
<td>42</td>
<td>internist-cardiologist/swimming</td>
<td>paroxysmal atrial fibrillation a year ago</td>
<td>no recent data</td>
<td>June 1982 drowned in a lake (he was a good swimmer)</td>
<td>no (found a day after)</td>
<td>not done</td>
</tr>
<tr>
<td>17</td>
<td>67</td>
<td>non-smoker surgeon/tennis</td>
<td>no</td>
<td>not recent data, normal ECG finding few years earlier</td>
<td>February 1992 during a game</td>
<td>yes</td>
<td>CHD, LADCA occluded, myocardial fibrosis, LV wall 15 mm</td>
</tr>
<tr>
<td>18</td>
<td>50</td>
<td>non-smoker immunologist/jogging</td>
<td>no</td>
<td>no data</td>
<td>July 1997 during jogging</td>
<td>no</td>
<td>CHD, LADCA occluded, LV wall 11 mm</td>
</tr>
<tr>
<td>19</td>
<td>60</td>
<td>smoker radiologist/swimming</td>
<td>general weakness in this day</td>
<td>LVH by ECG, arterial hypertension hyperlipoproteinemia</td>
<td>June 2002 during swimming</td>
<td>no</td>
<td>not done</td>
</tr>
</tbody>
</table>

Legend: LVH=left ventricular hypertrophy; RVH=right ventricular hypertrophy; IV=interventricular; RCA=right coronary artery; LADCA=left anterior descending coronary artery; HCM=hypertrophic cardiomyopathy; CHD=coronary heart disease.
pped, step out from a car, collapsed and died. All reanimation efforts were unsuccessful, including reanimation attempts in a nearest hospital – Regional Army Hospital. The forensic autopsy finding showed left ventricular hypertrophy of 18 mm, with no dilatation of the left ventricular cavity. Left descending coronary artery was narrowed. He had an acute pulmonary and cerebral edema.

Case 9 (Table 3). The fifth aged 25, played soccer recreationally. During an evening game in January 2005 in a sport hall, he suddenly collapsed and died. All reanimation attempts at the field were unsuccessful including a Mobile Emergency Medical Unit. The autopsy showed left ventricular hypertrophy of 22 mm and right ventricular hypertrophy of 7 mm, with normal cavities of the whole heart. The left and the right coronary arteries were narrowed at the ostial places to 1 mm (hypoplasia ostii arteriarum coronariarum cordis with malposition of the ostium of the right coronary artery). He had emphysema of the left pulmonary wing. He had an acute pulmonary and cerebral edema.

The data of three cases suffered of myopericarditis: two athletes and one engaged in recreative physical exercise are presented in Table 2 (case 7, case 10 and case 11).

Case 7. A school boy, aged 18, died suddenly during swimming is presented above as he had HCM, but also fibrinous pericarditis and large bilateral pneumonia.

Case 10. A professional soccer player, aged 17, was without symptoms. During regular training in February 1998 in a training hall, he suddenly collapsed and died. All reanimation attempts, including electrical defibrillation due to ventricular fibrillation repeated many times, were unsuccessful. An autopsy finding showed an enlargement of the whole heart hypoplastic right coronary artery, narrowed ascending aorta of 10 mm (normal finding at the ascending aorta reached 3.09 ± 0.41 cm), the thickness of the left ventricular wall was 18 mm, acute bacterial tonsillitis, and subacute diffuse myopericarditis.

Case 11. A school boy, aged 18 died suddenly during swimming. Resuscitation attempts were unsuccessful. An autopsy showed chronic myopericarditis and cardiac aneurysm of the left ventricle.

In this group of persons suffered of myopericarditis the death rate reached 0.10/100 000.

Ten teenagers-school boys who died during sports or recreative physical exercise are presented in the Table 2. Three of them presented before, died during sports activities: cases 4, 6 and 10. Seven teenagers died suddenly during recreational physical activities: cases 2, 7, 11, 12, 13, 14 and 15. The first was a school boy aged 17 (case 2) who was doing recreational physical exercise few times a week, without any com-
plaints. Few years before the incident, by the data of his family, a systolic murmur above the heart was discovered, with no written data about it. They told that the cardiac echosonography showed normal finding. Few days before the incident he has had a chest pain and a short breath. On a day of the incident in September 2010 he was running on a street, felt down and died. He was resuscitated on a street unsuccessfully. The forensic autopsy showed an enlargement of the heart reached 390 grams, thickening of the left ventricular wall of 19 mm (normal finding up to 12 mm), with normal thickness of the right ventricle: 3 mm, the aortic diameters were wide and reached 5.1 cm, and narrowing of the descending part of the aortic arch.

The second was a boy aged 14, who was engaged in regular physical activities in a school and died suddenly during exercise: a hypoplastic aorta and hypoplastic coronary arteries were found (case 12).

The third was 15 years old boy who died while playing basketball in school had hypertrophic cardiomyopathy, (case 4, presented above). The fourth victim, the 18 years old adolescent (case 13), had normal heart finding, bilateral pneumonia and possible high altitude noncardiogenic pulmonary edema. He was hospitalized in an Intensive Care Unit and dying after 24 hours. The fifth boy, aged 18, who died during recreational swimming, had signs of chronic myopericarditis with an aneurysm of the left ventricle (case 11). The mechanism for the lethal event in all five cases was probably malignant ventricular arrhythmia. The sixth boy – soccer player recreationally, aged 18, with hypoplastic ascending aorta, has been kicked in a chest during a game and died in a hospital due to bronchopneumonia (case 7). The seventh teenager aged 19, had cardiomegaly and sunk probably as a consequence of alcohol drink intake (case 14).

In male school boys in Croatia the death rate during physical exercise reached 0.37/100 000 yearly.

Three teenagers died during physical exercise of pneumonia. The first aged 18, smoker, just finished a secondary school and started to work recreatively at the site (case 13). The second aged 19 was a student who played soccer recreatively (case 15). In January 2002 he played soccer and few hours after a game he felt several pains in his left hip and leg and he developed high body temperature: 40°C. He left to the hospital at the other day. At the admittance he was septic and died one day later in spite of all therapeutic effort including antibiotics, inotropic drugs, fresh frozen plasma, coagulation factors, mechanic ventilation etc. The clinical diagnoses were: septic shock, adult respiratory distress syndrome, disseminated intravascular coagulation, acute renal failure. The forensic autopsy showed adult respiratory distress syndrome, disseminated intravascular coagulation, multiple pulmonary and supra-renal bleeding, hypoplastic right coronary artery, myocardial fibrosis and acute ce-
rebral edema. Histological finding showed bilateral bacterial pneumonia.

The third aged 18 was a school boy and professional soccer player. He didn’t complain of any discomfort during physical exercise (case 7).

In this period, five physicians specialists (one younger, three middle aged and one elderly) died suddenly during recreational physical exercise. In three of them a forensic autopsy was performed. All three had coronary heart disease; two of them left ventricular hypertrophy also. In all three left descending anterior coronary artery was atherosclerotic, with stenosis in one and occlusion in two. In one of them myocardial scars of two past myocardial infarctions were found. In no one acute myocardial infarction was found, so that obviously malignant ventricular arrhythmia was a cause of death. Two of them had a thickness of the left ventricle 15 and 18 mm. The main characteristics of them are presented in Table 3.

If 15% of male physicians specialists in Croatia are involved in recreate physical exercise (out of a total number of 4,957), the death rate reached 24.8/100,000.

Fifteen men aged 65 to 82 years, died due to physical exercise: 6 were engaged in swimming, 4 in tennis, one ridded, one was jogging, 2 bowling and one died during sexual act. One had symptoms of pectoral angina, 2 suffered from arterial hypertension, one accompanied by headache, and 2 had short breath during exercise due to congestive heart failure. Eleven were without symptoms. At autopsy, 14 had coronary heart disease, 5 had critical coronary artery stenosis, and 2 of them had signs of recent myocardial infarction, 3 had occluded left descending anterior coronary artery, one with acute myocardial infarction of the anterior wall, 12 had scars due to previous myocardial infarctions and one signs of diffuse myocardial fibrosis. In 12 of them left ventricular wall reached 15-23 mm. The characteristics of elderly men who died suddenly during or immediately after recreational physical exercise are presented in Table 4.

The mortality rates in elderly and younger men died suddenly during physical exercise are presented in Table 5. The reported deaths show that recreational physical exercise in elders reached 2.99/100,000, a higher incidence than the entire male population aged 15-64, engaged in recreational physical exercise: 0.96/100,000 (p=0.00001). That is a higher incidence than in the male population aged 15-40 engaged in physical exercise: 0.71/100,000 (p=0.00001). This incidence is higher than in young athletes: 0.19/100,000 (p=0.00000).

DISCUSSION

Sport activities have protecting effect on human organism. Sudden death in apparently healthy persons engaged in physical exercise is very rare. One subject was
Table 4. Elderly men died suddenly during physical exercise

<table>
<thead>
<tr>
<th>PHYSICAL ACTIVITY/EXERCISE/SPORT</th>
<th>CASE</th>
<th>AGE</th>
<th>SYMPTOMS</th>
<th>LETAL EVENT</th>
<th>CORONARY FINDING AT AUTOPSY</th>
<th>LVH (mm)</th>
<th>HEART ENLARGEMENT</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>20</td>
<td>65</td>
<td>pectoral angina ECG: ST, T changes</td>
<td>July 1988</td>
<td>CHD generalized, LADCA occluded, AMI anterior wall, biventricular hypertrophy</td>
<td>15</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>17</td>
<td>67</td>
<td>no</td>
<td>February 1992</td>
<td>CHD, diffuse myocardial fibrosis, LADCA occluded</td>
<td>15</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>21</td>
<td>65</td>
<td>no</td>
<td>June 1998</td>
<td>CHD, critical coronary stenoses, myocardial scar 6x3 cm anterior wall</td>
<td>25</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>32</td>
<td>66</td>
<td>no</td>
<td>May 2006</td>
<td>CHD, critical stenosis LADCA, AMI front wall, large myocardial scars front wall</td>
<td>21</td>
<td>680 g</td>
</tr>
<tr>
<td></td>
<td>23</td>
<td>69</td>
<td>no</td>
<td>September 2000 while swimming in the sea</td>
<td>CHD, myocardial scar anterior wall</td>
<td>18</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>25</td>
<td>82</td>
<td>no</td>
<td>July 2001 while swimming in the sea</td>
<td>CHD, myocardial scars, chronic pericarditis</td>
<td>23</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>26</td>
<td>82</td>
<td>no</td>
<td>July 2001 while swimming in the sea</td>
<td>CHD, myocardial fibrosis, pericardial adhesions, CAA with no changes</td>
<td>19</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>27</td>
<td>68</td>
<td>no</td>
<td>August 2001 while swimming in a swimming pool</td>
<td>CHD, critical stenosis LADCA, large scar of the LV</td>
<td>20</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>29</td>
<td>82</td>
<td>no</td>
<td>June 2002 while swimming in the sea</td>
<td>CHD, myocardial scar anterior wall, diffuse myocardial fibrosis</td>
<td>–</td>
<td>no</td>
</tr>
<tr>
<td></td>
<td>30</td>
<td>74</td>
<td>no</td>
<td>June 2002 while swimming in the sea</td>
<td>CHD generalized, myocardial scars posterior wall</td>
<td>–</td>
<td>no</td>
</tr>
</tbody>
</table>
a 15-year-old untrained boy who lost his way and run over a very rough terrain for miles, collapsed and died due to acute degeneration of numerous heart muscle fibers as the result of severe strain [47]. In young persons who died suddenly during physical exercise because of a cardiac arrest-ventricular fibrillation, in about 6% of the cases the cause is unexplained [2,3]. Beside the fact that his heart was not trained for such efforts, the role of coronary spasm and reperfusion perhaps could be taken into account in the pathogenesis of sudden death, as consequence of hyponatremia [48]. There is an open channel gene SCN5A as contributing to the risk for malignant ventricular arrhythmia.

The sudden and unexpected cardiac death rate among American athletes under the age of 30, amounts to 0.6/100 000 [49,50], which is higher than in our study. According to some authors, in competitive athletes the risk for sudden death in Minnesota reached 1:500 000 annually [7], or 1.5/1 000 000 [49], which is also higher than in our study. The relative risk of complications seems to be higher in exertion

Table 4. Continued

<table>
<thead>
<tr>
<th>PHYSICAL ACTIVITY/EXERCISE/SPORT</th>
<th>CASE</th>
<th>AGE</th>
<th>SYMPTOMS</th>
<th>LETAL EVENT</th>
<th>CORONARY FINDING AT AUTOPSY</th>
<th>LVH (mm)</th>
<th>HEART ENLARGEMENT</th>
</tr>
</thead>
<tbody>
<tr>
<td>JOGGING</td>
<td>24</td>
<td>65</td>
<td>headache, arterial hypertension</td>
<td>May 2001</td>
<td>CHD, biventricular hypertrophy</td>
<td>16</td>
<td>yes</td>
</tr>
<tr>
<td>RIDING A BICYCLE</td>
<td>31</td>
<td>72</td>
<td>short breath during exercise, arterial hypertension</td>
<td>August 2004</td>
<td>CHD, critical stenosis LADCA, myocardial scar LV: 40 mm hydropericardium: 500 ml</td>
<td>18</td>
<td>yes</td>
</tr>
<tr>
<td>BOWLING</td>
<td>28</td>
<td>73</td>
<td>no</td>
<td>December 2001</td>
<td>CHD, AMI posterior wall, myocardial scars anterior wall: 2 cm</td>
<td>21</td>
<td>410 g</td>
</tr>
<tr>
<td>SEXUAL ACTIVITY</td>
<td>33</td>
<td>80</td>
<td>short breath during exercise</td>
<td>May 2008</td>
<td>CHD, LADCA occluded, large myocardial scars front wall, diffuse myocardial fibrosis</td>
<td>-</td>
<td>yes</td>
</tr>
<tr>
<td></td>
<td>22</td>
<td>66</td>
<td>no</td>
<td>April 2000</td>
<td>CHD, critical LADCA stenosis AMI anterior wall, myocardial scar anterior wall 3x2.7 cm,</td>
<td>21</td>
<td>400 g</td>
</tr>
</tbody>
</table>

Legend: LV=left ventricle; RVH=right ventricular hypertrophy; LADCA=left anterior descending coronary artery; CHD=coronary heart disease; AMI=acute
than at rest [64]. It is estimated that in cross-country skiing the risk is 14.5 times higher than in other exercises. The analysis of the results shows, however, that the risk in all exercises is 4.5, in non-strenuous exercises it is 3.3, whereas it is 3 times higher in strenuous exercises, and in 9 in all population groups [51].

The young man in our study, aged 17, had multiple cardiovascular anomalies: hypoplastic right coronary artery narrowed ascending aorta and subacute myocarditis. In young persons who died suddenly in the Veneto Region in Italy, coronary anomalies reached 3.5% [7], and in USA 13.7% [7].

In athletes who died suddenly due to exercise, the most common reasons for such events are cardiomyopathies, coronary anomalies and myocarditis [3,49]. Among cardiomyopathies, ARVD is very often a disease with no symptoms and reached 1:2.500 to 1:5.000 of all sudden deaths during exercise in young athletes, which is lower than in our study: 2/7. Most of those persons suffering of ARVD have involved both ventricles. In the first of our cases, the right ventricle was involved, and in the second both ventricles were involved. The first symptoms could be ventricular arrhythmias and/or conduction disturbances: in about 75 per cent of those persons the first manifestation of the disease could be sudden cardiac death due to malignant ventricular arrhythmia [16,49,50-56], as were cases in two deceased athletes in our study. ARVD in Veneto Region in Italy was at the second place of the most common anomalies in young athletes who died suddenly: 12.5% [3], and is at the seventh place in the athletes in USA who died suddenly: 2.8% [7]. The disease has a familial occurrence, but the specific gene defect is not identified yet. In ECG the left bundle branch block could be present along with right axis deviation, inverted T-wave in

| Table 5. Mortality rates in elderly and younger men who died suddenly during physical exercise (1984-2010) |
|---|---|---|
| Age and type of physical exercise | Died during physical exercise | Died pro 100 000 |
| 1 | Athletes | 6 of 3 133 512 | 0.19 |
| 2 | School boys | 7 of 2 644 380 | 0.37 |
| 3 | Elderly men aged involved in recreational physical exercise | 15 of 501 066 | 2.99 |
| 4 | Physicians specialists | 5 of 20 128 | 24.84 |

Poisson Rates: 1 vs. 2: z = 0.306, p = 0.75935; 1 vs. 3: z = 7.346, p = 0.00000; 1 vs. 4: z = 16.772, p = 0.00000; 2 vs. 3: z = 6.406, p = 0.00000; 2 vs. 4: z = 14.701, p = 0.00000; 3 vs. 4: z = 4.326, p = 0.00002.
the right precordial leads and with ventricular premature beats, and after that with ventricular tachycardia. The right ventricular enlargement could be present, and the left ventricular enlargement is present in 50%. The right ventricular wall is relatively thin, and that is why in ARVD a subepicardial process is sometimes difficult to diagnose in vivo. Pathological anatomy finding depends on the stage of the disease and show changes in the myocardium including thinning in the some areas of the right ventricle with fibro-fatty changes [16,19,53,56-58]. Sometimes we are facing with an aneurysm of the right ventricle in a small number of these cases.

The etiology of ARVD is unknown. We are facing very often with a presence with inflammatory myocardial infiltrates and progression from myocarditis to ARVD, and also with the discovery of the mutation in genes related to desmosomes. There is an open question: could it be an inflammatory process modified by genetic influence in desmosome-related proteins [16]. This is an autosomal dominant hereditary disease in almost 50 per cent of all cases with variable penetrance [20], and could be located to various chromosomes [17,18], the first to chromosome 14q23-q24.

ARVD is uncommon under ten years of age, and rare appeared in the elderly [57,58]. By the Task Force criteria of this disease [32], the following criteria are necessary for definitive diagnosis of this disease: two major, or one major plus two minor, or 4 minor criteria; the borderline diagnosis consists of one major plus one minor, or three minor criteria; the possible diagnosis consists one major, or two minor criteria from different categories. The ECG showing an epsilon wave indicating ARVD, which is presented in only 33 per cent of patients suffering of ARVD [18,59,60]. Our two present young athletes we could not use the mentioned criteria, because they were without symptoms and died suddenly at the field during training.

Among cardiomyopathies by some data, HCM reached 0.2% of population [31]. HCM has variable thickening of the septum and the left ventricular wall and different clinical presentations. It could be obstructive forms of HCM (in about 30%) and non-obstructive forms (in about 60%), and in additional 10%, obstruction could be presented due to vigorous physical exercise [61]. There is no strong correlation between the degree of HCM and subjective symptoms [62-64]. Sometimes it is not easy to clinically distinguish physiological myocardial hypertrophy from HCM in athletes. Ventricular thickness more than 18 mm (sometimes 15), implies criteria for HCM.

The HCM reached 5% in the young in the Veneto Region in Italy [65], and it is at the first place of all causes of death during physical exercise in young athletes in USA: 26.4% [7]. It reached 0.1-0.2% in the population in USA, but it is a cause of sudden and unexpected cardiac death of competitive athletes in more than 50% [63]. The disease could be asymptomatic and the first manifestation could be sudden
cardiac death. These data were present in 2 athletes in our study and the death rate reached 0.06/100 000.

It is known that a magnitude of left ventricular or septal hypertrophy leads to high risk for sudden cardiac death and is a strong independent predictor to prognosis. The risk of sudden cardiac death raised progressively with left ventricular wall thickness: if LVH reached up to 19 mm, a risk for sudden death is low, but LVH of 30 mm or more doubled the risk of sudden cardiac death [66]. In those five presented cases of HCM in our study, a heavy obstructive form of HCM was seen in a school boy aged 15, with no previous symptoms, who died suddenly, i.v. septum reached 40 mm. It confirms the rule that extreme LVH are seen in the youngest with mild or no symptoms [66]. In other two athletes died suddenly, LVH reached 20 and 25 mm. In two of our persons engaged in physical activities recreationally and died suddenly, LVH reached 18-22 mm, in spite of the opinions that in persons with mild or moderate LVH the risk is low.

A sudden cardiac death due to physical exercise in young athletes in Croatia suffered of HCM reached 0.06/100.000 yearly (p=0.00000), in all young athletes suffered of other heart diseases reached 0.19/100 000 (p=0.00000), and in the total male population aged 15-40 engaged in sports and recreational physical exercise: 0.71/100.000. In preparticipation screening for HCM, a familial history data have to be included especially to sudden cardiac death (it could be negative), personal symptoms including recurrent exertion syncope, physical examination by physician (it could be normal finding in most of these patients), an ECG (it could be non-specific), LVH by echocardiography (<19 mm=low risk, >30 mm=high risk), and 24 h ECG- Holter (malignant ventricular arrhythmias).

The article deals with the aspects of interrelationship between physical exercise and different types of myopericarditis in young male athletes. In the Veneto Region, myocarditis was at the third place as the cause of death in athletes during physical exercise reaching up to 7.5% [65]. In our study the rate was 0.11/100 000 compared to 0.19/100 000 in young athletes [11] who died due to all cardiac reasons during exercise.

Myocarditis reached 7.5% in the Veneto Region [65], and was at the 3-rd place as a cause of death rate in athletes during physical exercise, in our study: 0.11 /100 000. In general Croatian population aged 15-40 who practice physical exercise, the death rate reached 0.71/100 000. This means that athletes obviously are protected from cardiovascular events.

Myopericarditis could lead to malignant ventricular arrhythmia and sudden cardiac death. Acute viral infections of the upper respiratory tract can diminish isometric myocardial strength for 15% [31,49]. In such cases, diminution of enzymatic
myocardial activity [46] and abnormality of myocardial structure could be detected by electronic microscopy in samples of repeated myocardial biopsies. In Croatia, Medved et al. [67] were the first who described a case report dealing with a sudden cardiac death due to chronic myopericarditis in athlete, aged 29, a professional soccer player who had clinical signs of heart enlargement and frequent left ventricular premature beats. He refused to follow the physicians’ advices to stop with competitive matches. During a game, he suddenly felt ill, suddenly collapsed and died in spring 1973. He was resuscitated, transferred to the nearest hospital but died in transit. Forensic autopsy showed chronic myopericarditis, LVH of 15 mm, the right ventricle wall reached 5 mm; the initial part of the aorta was narrowed, and with enlargement of the whole heart.

Physical exercise is contraindicated in different types of myopericarditis for at least six months. When to restart training depends on the disappearance of subjective symptoms, on the normalization of clinical state and biochemical parameters, i.e. double viral titers, on the normalization of electrocardiographic finding in a 24-hour electrocardiogram, and on electrocardiographic and echocardiography scans at rest and during exercise (stress). Sometimes in specific cases other procedures are needed: radionuclide studies, cardiac catheterization and magnetic resonance imaging.

Many possibilities exist in defining an exertion-related sudden cardiac death. The time period needed for the cardiovascular system to return to resting steady state varies with many factors such as the type, intensity and duration of the activity, and the health status and physical condition of the individual. Thus, it is not easy to define exactly what are an exertion-related death and especially a sudden death caused by exercise. Some researchers count only deaths during or immediately after the cessation of exercise as exercise-related deaths, while some authors include also deaths that had occurred 30 minutes, 60 minutes or even more after the exercise [51,68]. According to our data the rate of sudden and unexpected death during or after recreational physical exercise was 0.37/100 000 per year in boys in or immediately after secondary schools in Croatia. This is lower incidence than in other age groups in Croatia. For example the rate of sudden and unexpected death during or after physical exercise in male physicians-specialists was much higher than in adolescents: 24.8/100 000, and also higher than in men aged 15-40: 0.71/100 000 per year in Croatia, in men aged 15-64 years: 0.96/100 000 per year, and in the elderly: 2.99/100,000 per year [8]. Although the incidence of sudden unexpected death in the school boys was low, our findings bring up once again the question, whether some or all of these incidents could have been prevented. None of the boys had reported any noticeable symptoms at exertion. However, all of them suffered from chronic or acute cardiac or other diseases: two had congenital hypoplasia of the coronary
arteries, one together with narrowing of the ascending aorta and subacute myocarditis, one had HCM, one had a chronic myopericarditis with a left ventricular aneurysm. In one young man the heart findings were normal, but there was bilateral pneumonia and possible non-cardiogenic pulmonary and cerebral edema. This 18 years old adolescent was badly nourished, skinny, exhausted because of flu, and possibly dehydrated. The possible cause of death might have been hyponatremic encephalopathy with consecutive non-cardiogenic pulmonary edema during bilateral pneumonia. Non-cardiogenic pulmonary edema could be a clinical manifestation of hyponatremic encephalopathy [48]. The most common abnormalities of young persons who have died suddenly during or after physical exercise have been found to be hypertrophic cardiomyopathy (especially in young competitive athletes, 36%) or congenital coronary anomalies [6,69,70]. In Italy (Veneto region) in a 17-year period (1979–1996) among 49 competitive young athletes (44 male and 5 female) who died suddenly, hypertrophic cardiomyopathy was observed in only one subject or 2% in autopsy [71], and coronary anomalies in 18% of the victims. In a Swedish study [2], in a period of 14 years, 16 sudden and unexpected cardiac deaths among young orienteer’s aged 18–27 years (15 male and one female) were found. Six had acute myocarditis, four ARVD, and two had healed myocarditis. This data demonstrates that the causes of sudden death of young athletes at exertion can vary greatly in different countries and athletic populations. The most common mechanism for lethal event is ventricular fibrillation. The most difficult problem is that cardiac disease in young persons is usually not recognized prior to death, as was the case also in our study. An American study [7] suggests that a person can begin a gradual exercise program without consulting a physician, if the answer is no to the following questions: 1) history of heart trouble; 2) heart murmur, 3) heart attack, 4) arterial hypertension; 5) diabetes mellitus, 6) arthritis; 7) family history of premature coronary heart disease; 8) exercise-related shortness of breath, faintness, dizziness, or pain or pressure in the chest, neck, shoulder or arm. In order to give reliable answers to these questions, information from prior medical examinations are often needed [16]. Carefully collected anamnestic data with special attention to personal and familial (parental sudden death) cardiovascular history, personal physical examination, an electrocardiogram at rest, and especially an echocardiography study on the basis of other information are essential in every potential athlete or in suspicious cases.

Echocardiography is today a standard non-invasive cardiac investigation especially in screening of congenital heart diseases. A 12-lead electrocardiogram is the most valuable preparticipation cardiovascular modality of these three mentioned methods at the beginning, and an echocardiography is a superior method for detecting congenital cardiovascular anomalies. However, it cannot be considered as a large-scale
screening examination of symptomless individuals because of the equipment and expertness needed and also because of diagnostic uncertainties. Echocardiography can give a false positive finding e.g. in borderline thickness of the left ventricle or in dilated heart cavities. A false negative finding can occur in cases of HCM which might not be detectable until the adolescent period [7]. Detection of premonitory cardiac symptoms such as shortness of breath, history of exertion syncope or chest pain, has to lead to diagnostic procedures such as ergometry, thallium stress test, echocardiography and coronarography especially in competitive athletes [6]. Physical exercise is contraindicated in a case of acute infection, especially a respiratory one. In this report the cause of death was an acute respiratory infection, in one previously healthy adolescent. In another boy with hypoplastic aorta and coronaries and subacute myocarditis, a respiratory infection probably was a trigger. Thus, examination of the anamnestic and autopsy findings in the six cases of this data suggests that avoidance of heavy exertion at the time of respiratory infection (cases 7 and 10) and thorough preparticipation physical examination and indicated diagnostic tests (cases 3 and possibly 5) could have helped in avoiding the fatal event.

This study deals also with the aspects of the interrelationship between physical exercise and acute respiratory tract infections in young male athletes also. All three presented teenagers athletes suffered of bacterial pneumonia. Two of them had cardiovascular diseases: one aged 19 had hypoplastic right coronary artery and myocardial fibrosis and the other aged 18 had HCM and hypoplastic ascending aorta. Pneumonia as a cause of sudden cardiac death during physical exercise is very rare [11]. In Swedish study during 14 years, 15 young men and one young woman died during exercise [1]. By some authors HCM is the most frequent causes of death in younger subjects [49,72]. HCM was an uncommon cause of death in young competitive athletes: in Veneto region in Italy, 49 sudden death in athletes were registered from 1979-1996, of which 12.2% had anomalous origin of a coronary artery and only 2.0% had HCM [3]. In distinguishing physiological myocardial hypertrophy from HCM, we are facing with the next criteria: normal values of ventricular thickness reached 11 mm, borderline values between 11-13 mm and thickness for HCM above 15 mm. These data was present in one athlete in our study.

Two of presented cases had multiple cardiovascular anomalies and acute bacterial pneumonia at the same time. One had hypoplastic aorta with subacute diffuse myopericarditis. The other had hypoplastic right coronary artery. Coronary artery hypoplasia is an infrequent disease and could be associated with sudden cardiac death as could be narrowness of the ascending aorta or hypoplastic aorta. These result differ by different authors: in young persons who died suddenly in the Veneto Region in Italy, coronary anomalies reached 3.5% [7], and in 13.7% in USA [7].
Myocarditis possibly related to exercise during the febrile phase of a viral illness could be the reason for sudden death [49,72], especially when it is connected with the narrowing of the ascending aorta as was the case in one teenager in our study.

In Croatia the death rate including all reasons among athletes reached 0.19/100 000. The sudden and unexpected cardiac death rate among USA athletes below age of 30 amounts 1.6/100 000, which is higher than in our study. In Minnesota that rate reached 0.2/100 000 or 0.46/100 000 annually which is also higher than in our study [6,9].

From a study in USA [9], 122 sudden death due to physical exercise were described among 21.481 male physicians or 41/7.176 every four years, which is also higher than in our study.

The relative risk of cardiovascular complications is higher in exercise than at rest: cross-country skiing is 14.5 times higher than in other exercises. The risk in strenuous exercise is 4.5 times higher than in non-strenuous exercise [51]. In literature we were unable to find data of mortality rate during bacterial pneumonia in athletes during training. Physical exercise is contraindicated in acute respiratory tract infections. In all three present cases acute bacterial pneumonia had been a cause of death. When to start with physical training after pneumonia depends on disappearing of clinical signs of pneumonia, normalization of body temperature, disappearing of pulmonary infiltrate by chest X-ray or local signs of tonsillitis and regional physical finding i.e. an enlargement of lymphatic nodes, of C-reactive protein and erythrocyte sedimentation rate, and normalization of white cell count.

A medical checkup before recreational physical exercise is essential including a clinical examination, a serum concentration of risk factors and other risk factors, an electrocardiogram at rest, a stress test and echocardiography studies in clinical indication, and medical controls in persons taking exercise. It has to be done every three years in that aged 40 and more who are without symptoms. It has to be done every two years in those aged 35 and more with one or more risk factors for atherosclerosis – coronary heart disease, in persons with suspect cardiac, pulmonary or metabolic disease. A medical check-up has to be done once a year in those with known cardiac, pulmonary or metabolic disease or with positive stress test previously regardless of their age [41,66,69,71]. This study shows that medical evaluation is important because of the underlying problems such as sudden death during exercise. In non-trained persons and in the elderly a physical exercise should be recommended of a gradually intensity, which could not exceed 6 METs (1 MET = 3.5 ml O₂/kg/min).

In this study the aspects of the interrelationship between recreational physical exercise and cardiovascular risks in five physicians are presented. Three of them were autopsied and in all coronary heart disease was found, with no signs of an
acute myocardial infarction. In two a LVH was found which could be an important risk factor for a malignant arrhythmia. In two who were not been autopsied, an alcohol cardiomyopathy was probably a reason for malignant arrhythmia. One had atrial fibrillation one year before sudden death. The other had arterial hypertension also, an excessive hyperlipoproteinaemia, and probably a silent myocardial ischaemia due to coronary heart disease. Atherosclerosis is silent until critical stenosis, coronary thrombosis (observed in three physicians in this study); embolus or coronary dissection or aneurysm supervenes. Sudden cardiac death could sometimes be the first sign of coronary disease (in over 25 per cent), precipitated by excessive physical exercise, what was observed in five presented physicians. LVH, presented in two of three autopsied physicians, is often connected with arterial hypertension. In LVH there is also an imbalance between myocardial supply and requirements. This disease is very often a silent cardiovascular risk factor and could have important prognostic implications [25,41,51].

Sudden death in apparently healthy persons who are engaged in physical exercise is extremely rare. When cardiovascular incidents occur during exercise, the most frequent cause is an organic heart or vascular disease [25,36-40,51,73-76]. One should always bear in mind the fact that a great number of persons with coronary disease engage in physical exercise, and only a few have any discomfort. These data are supported by the analysis of the health-related condition of the Croatian population [25,36-41]: the so-called healthy persons of both genders aged 65–84 have 6 diagnoses on the range 0–17, including cardiovascular diseases. The relative risk of cardiovascular complications seems to be higher in exertion than at rest [25,36-41,51]. The relative risk of sudden death during up to 30 minutes of vigorous exercise reached 16.9, and absolute risk is low: 1/1,510,000.

Fifteen of 69 died persons from this study were aged 65 or over. Eleven of those persons did not have any previous cardiovascular symptoms, and they had not had any medical record in the recent years before the accident. In this group, in all but one coronary heart disease was the cause of death. In accordance with the literature, when cardiovascular incidents occur during physical exercise in elders, the most frequent cause of sudden death is coronary heart disease [11,42,75,76].

People engaged in physical exercise have a lower risk of cardiovascular complications than inactive persons. Exercise is thus an important component of healthy living for the senior citizens. It has been evidenced that physical exercise positively affects maintenance and promotion of health. Physical activity should be controlled, regular and adapted to the health condition [6]. The medical check-ups before exercise are essential, as well as medical control over persons taking exercise [76-80].
Physical exercise of an biologically elderly person may make a heart less stable; could actually increase chances for arrhythmia; it does bring protection against heart attack, but this protection is gained and lost rapidly. In elderly it is necessary to have an activity plan for preventive and therapeutic recommendations [11]. Most of the elderly could be active physically. In most of the elderly capable to exercise recommendations is moderate aerobic physical exercise for 30 minutes, 5 times a week, or vigorous activity (such as jogging) for 20 minutes three times a week [79,80].

In Croatia, about 7 per cent of the entire male population undertakes recreational physical exercise. The reported deaths reveal that low or moderate recreational physical exercise in elders reached 2.99/100 000, a higher incidence than the entire male population aged 15-64 engaged in recreational physical exercise: 0.96/100 000 yearly. That is a higher incidence than in the total male population aged 15-40, engaged in physical exercise in Croatia: 0.71/100.0000, with the significant difference (p=0.00001). It is also a higher incidence than in athletes suffering from acute bacterial pneumonia or suppurant tonsillitis, a higher incidence than in young athletes: 0.19/100 000 [17], and the difference is significant (p=0.00005). LVH of 15-25 mm was found in 12/15 indicating an underlying risk of untoward cardiovascular events in elderly persons without symptoms.

The prevalence of sudden death almost exclusively in males is probably consequence of two factors: males are doing heavier physical exercise than women, and coronary heart disease is more frequently presented in males.

By the recommendations of the American College of Sports Medicine and the American Heart Association [78], the exercise test is not necessary for asymptomatic persons before starting with a low or moderate level of physical exercise and there is no need for an examination of a physician. Persons with symptoms or with diagnosed cardiovascular disease(s), arterial hypertension, diabetes mellitus or some other chronic disease, have to consult a physician before physical exercise especially vigorous one. But our results does not go to this direction: by our experiences only 4 out of 15 elderly men who practiced low or moderate physical exercise had symptoms of cardiovascular disease. When cardiovascular incidents occur during physical exercise in elders, the most frequent cause of sudden death is coronary heart disease. In 14 out of 15 presented elderly men who died suddenly during or immediately after recreational physical exercise, coronary heart disease was found. In four of them a recent myocardial infarction was observed, five had critical coronary stenosis, three had occluded one coronary artery and thirteen had scars of previous myocardial infarction(s). That is why by our opinion, since preparticipation medical screening cannot ensure safety of exercise completely, physical condition check before exerci-
se is necessary to avoid possible harmful effect of physical exercise. Our data show that the usual underlying cause of sudden cardiac death during or immediately after physical exercise is coronary heart disease in elders. More than 50% of elders have at least one risk factor and about 10% of them have more than two risk factors for atherosclerosis. More efficient medical screening systems will be needed to cope with the increasing numbers of participants in exercise [56].

These findings suggest that medical examination of all elderly persons has to be done before recreational physical exercise and their health has to be monitored by clinical examination, searching for risk factors for atherosclerosis, performing ECG at rest, stress ECG, and echocardiography, because of the potential presence of an underlying disease that could result in acute cardiovascular complications during exercise.

In 37 deceased males who are not presented here in details, 32 of them suffered of coronary heart disease, 7 of them with signs of an acute myocardial infarction, 19 with signs of myocardial fibrosis or myocardial scars, and 18 with signs of left ventricular hypertrophy were observed.

References


Sažetak

Nagla kardijalna smrt tijekom tjelesne aktivnosti u Hrvatskoj u 27-godišnjem razdoblju


U sportaša je bilo šest naglih i neočekivanih smrtnih ishoda. Obdukcijom trkača u dobi 21 godine otkriven je akutni infarkt miokarda s normalnim vjenačnim arterijama i hipertrofiom lijeve klijetke. U drugog, u dobi 17 godina, vjenačne arterije bile su hipoplastične a silazna aorta sužena te su bili prisutni gnojni tonzilitis i subakutni miokarditis. Dva su sportaša umrla tijekom treninga zbog aritmogene displazije desne klijetke. Prvi je bio trkač na kratke pruge, a drugi nogometaš. U Hrvatskoj stopa nagle smrti u sportaša iznosi 0.19/100 000 godišnje (p = 0.00005); zbog aritmogene displazije desne klijetke ta stopa iznosi 0.06/100 000 (p = 0.00000), u svih mladih sportaša koji su bolevali od drugih bolesti srca ona iznosi 0.10/100 000 (p = 0.00000), u ukupnoj populaciji muškaraca u dobi 15 – 40 g. koji su uključeni u takmičarsku ili rekreacijsku tjelovježbu ta stopa iznosi 0.71/100 000 godišnje (p = 0.00001), a u svih muškaraca u dobi 15 – 64 g. 0.96/100 000 (p = 0.00000).

Pet sportaša preminulo je zbog hipertrofijske kardiomiopatije: trojica za vrijeme treninga i dvojica za vrijeme rekreacijske tjelovježbe. Jedan je imao opstrukcijsku hipertrofijsku kardiomiopatiju, a ostali su imali neopstrukcijski oblik. Prvi sportaš naglo je preminuo za vrijeme treninga trčanja, a drugi i treći sportaš bili su košarkaši. Dvojica su preminula za vrijeme nogometne igre. Nagla kardijalna smrt zbog hipertrofijske kardiomiopatije za vrijeme rekreacijske tjelovježbe u mladih sportaša u Hrvatskoj iznosi 0.10/100 000 (p = 0.00000).

Dogodila su se tri nagli i neočekivana smrtna ishoda zbog mioperikarditisa, i to kod dvojice nogometaša te jednog plivača. Obdukcijom je otkriveno da je prvi bolovao od subakutnog difuznog mioperikarditisa, gnojnog tonzilitisa i sužene uzlazne aorte, drugi od kroničnog mioperikarditisa i aneurizmatskog proširenja lijeve klijetke, a treći od fibrinoznog perikarditisa, zadebljanja lijeve klijetke od 20 mm, hipoplastične uzlazne aorte, obostrane bronhopneumonije i kontuzije mozga s edemom. U Hrvatskoj u tjelovježbača stopa smrtnog ishoda zbog mioperikarditisa iznosi 0.11/100 000 (p = 0.00000).

Stenoza aorte ustanovljena je u dva adolescenta. Prvi je bio učenik i košarkaš u dobi 17 g. Obdukcijom je otkriveno povećanje cijelog srca, suženje uzlazne aorte, zadeljana stijenka lijeve klijetke, bakterijski tonzilitis i subakutni difuzni miokarditis. Drugi učenik preminuo je naglo za vrijeme trčanja. Nalaz obdukcije upućivao je na povećanje cijelog srca, zadebljanje stijenke lijeve klijetke i suženje silazne aorte.

Malformacije vjenačnih arterija ustanovljene su u trojice mladića. U nogometaša koji je na-glo preminuo tijekom utakmice postojala je hipoplasija desne vjenačne arterije, suženje uzlazne aorte, akutni bakterijski tonzilitis i subakutni miokarditis. U dječaka koji je preminuo za vrijeme tjelovježbe ustanovljene su hipoplastične vjenačne arterije. U mladića koji je povremeno rekreacijski igrao nogomet i preminuo za vrijeme utakmice desna i lijeva vjenačna arterija bile su sužena ušća od 1 mm.

Među adolescentima bilo je 10 naglih i neočekivanih smrtni tijekom ili nakon tjelovježbe. Obdukcijski je kod četvorice ustanovljena prirođena bolest srca. Dvojica su imala hipoplastične koronarne arterije, jedan uz to i bakterijsku upalu tonzila, suženu aortu i subakutni miokarditis. Treći ima hiper-terofijsku kardiomiopatiju. Četvrti je bolovao od obostrane upale pluća s nekardiogenim edemom pluća. Peti je imao aneurizmu lijeve klijetke. Šesti je igrao nogomet.
rekreacijski i preminuo je zbog obostrate upale pluća, a imao je i fibrinozni perikarditis. Sedmi je imao kardiomegaliju, obostrani hidrotoraks i etanol u krvi (0.17%). Stopa smrti za vrijeme ili neposredno nakon tjelesnog vježbanja u adolescenata u Hrvatskoj iznosi 0.37/100 000 (p = 0.00226). Zbila se tri nagla i neočekivana smrtina ishoda zbog pneumonije u mladića. Jedan je radio rekreacijski na gradilištu, drugi se bavio rekreacijskim nogometom, treći je bio profesionalni nogometaš. Nalaz obdukcije u prvog upućivao je na obostranu bakterijsku upalu pluća, mogući nekardiogeni edem pluća i edem mozga. U drugog je nalaz upućivao na obostranu bakterijsku upalu pluća, respiracijski distresni sindrom odraslih, diseminiranu intravaskularnu koagulaciju, krvarenje u nadbubrežne žlijezde, hipoplastičnu desnu koronarnu arteriju i fibrozu miokarda. U trećeg je nalaz upućivao na obostranu bakterijsku upalu pluća, fibrinozni perikarditis, cerebralnu kontuziju s edemom, zadebljanje lijeve klijetke i hipoplastičnu uzlaznu aortu. Stopa smrtnosti u adolescenata zbog pneumonije u Hrvatskoj iznosi 0.11/100 000 (p = 0.00000).

Mladi nogometaš bez prethodnih zdravstvenih tegoba preminuo je naglo i neočekivano tijekom treninga. Obdukcijom je ustanovljena bakterijska upala tonzila, subakutni mioperikarditis i sužena uzlazna aorta.

Pet je liječnika specijalista naglo i neočekivano umrlo tijekom ili neposredno nakon rekreacijskog tjelesnog vježbanja: plivanja, nogometa, tenisa i trčanja. U trojice koja nisu imala prethodne simptome te su bila nepušači obdukcijom je ustanovljena koronarna bolest srca: u jednog je ustanovljena stenoza, u dvojico okluzija prednje silazne koronarne krvne žile, u jednog su ustanovljeni ožiljci miokarda od dva ranije preboljela infarkta miokarda, u dva hipertrofija lijeve klijetke. U dvojice koja nisu obducirana radilo se o mogućoj alkoholnoj kardiomiopatiji. Obojica su bila pušači. U jednog je zabilježena fibrilacija atrija godinu prije nagle smrti. U drugog se vjerojatno radilo o bolesti vjenčanih arterija, uz arterijsku hipertenziju i hiperlipoproteinemiiju. U Hrvatskoj je 4.957 liječnika specijalista muškaraca, a u njih stopa naglih smrti za vrijeme tjelesnog vježbanja iznosi 24.8/100 000 godišnje (p = 0.00000).

Petnaest „starijih“ muškaraca naglo je umrlo za vrijeme različitih oblika tjelesnog vježbanja: plivanja u moru, tenisa, kuglanja, rekreacijskog trčanja. Obdukcijom je otkriveno da ih je 14 imalo bolest vjenčanih krvnih žila: petorica kritično vjenačno suženje, dvojica uz recentni infarkt mišića srca prednje stijenke, trojica začepljenu prednju silaznu vjenačnu arteriju, jedan od njih uz to i akutni infarkt prednje stijenke srca, dvanaeostica ožiljke nakon preboljelih infarkta srca, jedan je imao difuznu fibrozu miokarda. Dvanaeostica su imala zadebljanje lijeve klijetke. U Hrvatskoj se oko 7% muškaraca starije dobi bavi rekreacijskim tjelesnim vježbom, a u njih stopa smrtnosti iznosi 2.99/100 000 (p = 0.00001).

Ključne riječi: muškarci; sportaši; rekreativna tjelesna aktivnost; iznenadna smrt