1 Atrial fibrillation in female endurance athletes

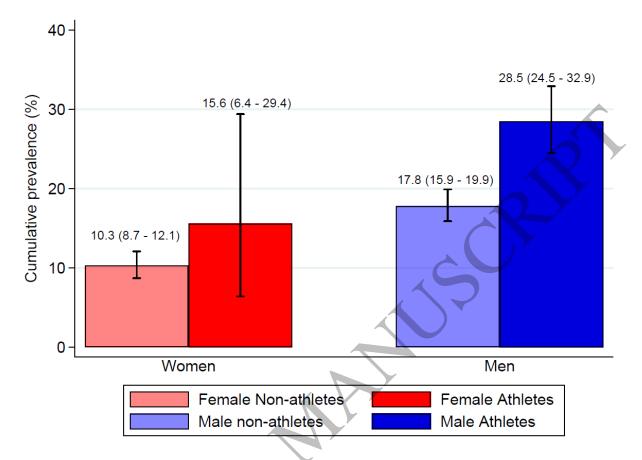
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- 17 Part of the work (the cumulative prevalence of atrial fibrillation in the female athletes) was presented
- 18 during the oral abstract session 'Sports Cardiology and Exercise 2', taking place at the open stage of the
- 19 European Society of Cardiology congress in Malaga, Spain, on 14 April 2023.
- The work has been supported with a PhD-grant from the Northern Norway Regional Health Authority(grand number HNF-1568-21).
- 22 Word count: 995 (including references and figure legend, excluding statements and table).
- 23 Key words: Atrial fibrillation; Female athletes; Exercise; Endurance sports; Arrhythmias; Sports
- 24 cardiology; Physical activity; Athlete; Women
- 25 Physical activity (PA) and exercise have been associated with the risk of atrial fibrillation (AF) in a U-
- 26 shaped manner (1). While moderate PA may reduce the risk, the highest prevalence of AF is observed in
- 27 sedentary individuals, and amongst middle-aged male endurance athletes (2). Characteristic for these
- athletes is that AF occurs in the absence of structural heart diseases and other established AF risk
- 29 factors, suggesting prolonged and vigorous endurance exercise may be a causal factor for AF, and the
- 30 'athlete's heart' is a proarrhythmic condition (3). However, the pathophysiological mechanisms for AF in
- 31 athletes are not fully understood (4). Female athletes have been underrepresented in previous studies
- 32 and the role of female sex in the association between PA, exercise and AF remains unclear. We aimed to

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- 1 study the prevalence of AF amongst older female competitive athletes with a history of prolonged
- 2 exposure to endurance sports.
- 3 We conducted a prospective cohort study comprising Norwegian non-elite participants aged ≥65 years in
- 4 the cross-country (XC) ski competition 'The Birkebeiner race', and individuals of the same age group (65-
- 5 74 years) attending the population-based Tromsø Study. We have described the methods of the study
- 6 and reported the risk of AF amongst the male participants previously (2). AF and all reported covariates
- 7 were self-reported by questionnaires at baseline (2009-2010) and during follow-up surveys (2014, 2020).
- 8 We report cumulative prevalence of AF, defined as self-reported AF at least once during the study period
- 9 among participants still alive at follow-up, divided by all participants who attended the baseline survey
- 10 and were still alive at follow-up. We report continuous variables as means (standard deviations) and
- 11 categorical variables as percentages. The study was approved by the Health Research Ethics Committee
- 12 (REK: 2020-175586). All participants gave informed consent to participate.
- 13 Out of 51 invited female XC-skiers and 1799 non-athletes, 46 (90%) and 1375 (74%) participated in the
- 14 study. Table 1 shows baseline characteristics. Female athletes reported a median exposure to endurance
- 15 exercise of 26 years, and had completed the Birkebeiner XC-race for a median of 9 years. One athlete
- 16 (2%) and 97 non-athletes (7%) died during follow-up. During the 10-year follow-up period, 7 out of 46
- athletes reported AF. The cumulative prevalence of AF was 15.6% (95% CI 6.4-29.4) and 10.3% (8.7-12.1)
- 18 in female athletes and non-athletes, respectively. Cumulative prevalence for AF in female and male
- 19 athletes and non-athletes are shown in Figure 1. Three athletes reported other cardiac conditions than
- 20 AF, and another three suffered a stroke.
- 21 Despite a low prevalence of other cardiac conditions and established AF risk factors, the cumulative
- 22 prevalence of AF in older female athletes was relatively high. We did not observe a statistically significant
- 23 difference in AF prevalence between athletes and non-athletes. Notably, none of the athletes with AF
- suffered a stroke and the mortality was low. Although not fully explained, exercise-induced cardiac
- 25 remodeling (EICR), such as left atrial enlargement, is amongst the suspected underlying mechanisms for
- 26 exercise-induced AF (5). For unknown reasons, EICR appears to have different characteristics in female
- 27 compared to male athletes of younger ages (6). AF in female endurance athletes have been only sparsely
- studied (7), and future studies should aim to address how EICR and potential sex differences may be
- associated with the risk of AF in female and male athletes. The low number of female athletes in our
- 30 study is a main limitation, but reflects female underrepresentation in endurance sports competitions.
- 31 The self-reporting of AF and other covariates is another limitation.
- 32 In conclusion, we found a relatively high prevalence of AF in older female athletes with prolonged
- 33 exposure to endurance sport, but the prevalence of AF did not differ between athletes and non-athletes.
- 34 Studies with better representation of female athletes are needed to address sex differences in
- 35 pathophysiological mechanisms underlying EICR and exercise-associated AF.
- 36 Figure 1.



1

2 Figure legend:

- 3 Cumulative prevalence of atrial fibrillation with 95% confidence intervals in older female and male cross -
- 4 country skiers during 10-year follow up, and in non-athletes of the same age participating in the Tromsø
- 5 Study, based on current and previous analyses (2).
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1 **Table 1.**

2 Baseline characteristics of female athletes and non-athletes.

3

Baseline characteristics	Athletes (n=46)	Non-athletes (n=1375)
Age, years	67.5 (2.2)	68.8 (2.8)
BMI, kg·m ⁻²	22.0 (1.7)	27.1 (4.7)
Overweight participants	4 (8.9)	571 (41.6)
Obese participants	0 (0.0)	317 (23.1)
Coronary heart disease	1 (2.3)	140 (10.3)
Previous stroke	0 (0.0)	52 (3.9)
Diabetes mellitus	1 (2.4)	104 (7.7)
Currently or previously	9 (21.4)	557 (41.2)
antihypertensive medication		
Currently or previously lipid	6 (14.6)	378 (28.3)
lowering medication		
University educated	25 (55.6)	247 (18.4)
Current smoker	0 (0.0)	221 (16.5)
Frequency of alcohol consumption		
>Once per month	31 (73.8)	559 (41.9)
Leisure time physical activity		
Sedentary	0 (0.0)	219 (19.4)
Light activity	6 (14.3)	809 (71.5)
Moderate	23 (54.8)	103 (9.1)
Vigorous	13 (31.0)	1 (0.1)

4 Means (standard deviation) for continuous and as numbers (percentage) for categorical variables. Except

5 from coronary heart disease, previous stroke, diabetes mellitus and lipid-lowering drugs, all differences

6 were statistically significant (p<0.05, parametric t-tests or $\chi 2$ test).

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1 Funding

- 2 This work has been supported with a PhD-grant from the Northern Norway Regional Health Authority
- 3 (grand number HNF-1568-21).
- 4

5 Data Availability Statement

- 6 Data from The Birkebeiner Study may by shared upon reasonable request to the corresponding author.
- 7 Data underlying this article were provided by the Tromsø study by permission and may be shared with
- 8 permission.
- 9

10 Author's contributions

- 11 Authorship: MM and AHR contributed to the conception, design and acquisition of the work. All authors
- 12 contributed to the analysis, or interpretation of data for the work. MM and KRJ drafted the manuscript.
- 13 All critically revised the manuscript, gave final approval, and agree to be accountable for all aspects of
- 14 work ensuring integrity and accuracy.
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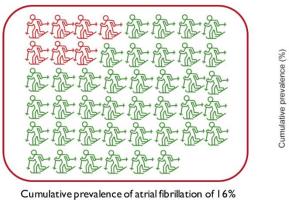
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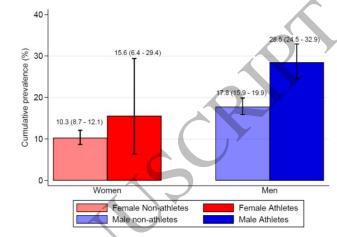
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Prolonged vigorous endurance exercise is associated with increased risk of atrial fibrillation in middle-aged and older male athletes, but few studies have reported prevalence of atrial fibrillation in female endurance athletes



during 10 years follow-up of older recrational

female cross-country skiers



The cumulative prevalence of atrial fibrillation among older recreational female endurance athletes was high compared to less active women of the same age-group, but lower than in male endurance athletes. The study suggests that vigorous endurance sport practice may also be a risk factor for AF in female athletes.

Graphical Abstract 160x97 mm (x DPI)