ORIGINAL ARTICLE

Evolution of Early Repolarization Patterns after 5 Years in a Military Population at Low Cardiovascular Risk and Practical Implications in Military Medical Expertise

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Background: The French military population is a young and athletic population with a high prevalence of early repolarization patterns (ERP) compared to the general population. Screening of military officers at risk of sudden death (SD) is a priority in military medical expertise. The aim of our study was to evaluate the prevalence of ERP and its evolution over a period of 5 years in a specific asymptomatic population, free of heart disease and cardiovascular risk factors.

Methods: From March to December 2008, we prospectively collected the electrocardiograms (ECGs) of military officers enlisted into the submarine forces of France. For 5 years (until 2012), the military officers included in the study underwent a clinical examination twice per year. A new ECG was carried out 5 years after the initial one.

Results: A total of 250 male subjects were included (mean age 22.87 ± 0.5 years). The prevalence of ERP was 19.2%. The most common appearance was an elevation of a slurring type J point (31/48 or 64.5%) in inferolateral leads (18/48 or 37.8%). After 5 years, the prevalence of ERP was identical to that of the previous period (19.2%) with no change in appearance, distribution, and amplitude in 47 subjects (97.9%). Repolarization changed in two patients (one occurrence of ERP and one disappearance). During the 5 years of follow-up, the subjects remained asymptomatic

Conclusion: ERP is common in our population of young, athletic, and asymptomatic military officers and changes little in 5 years.

Ann Noninvasive Electrocardiol 2015;20(5):420-425

early repolarization; sudden death; young subjects

The French military population is a young, predominantly male population subject to specific physical and environmental constraints. The incidence of sudden death (SD) in young military officers under the age of 35 years is estimated at 2.5 deaths/100,000 per annum.¹ One of the main missions of military cardiologists is the detection and treatment of acquired or congenital heart disease responsible for SD. To detect these heart diseases, all French military officers undergo a clinical examination and a routine electrocardiogram (ECG) during their induction and regularly throughout their career. In the presence of suggestive clinical and ECG abnormalities, the patient is addressed to the cardiology department of one of the military hospitals for further advice.

In 2008, Haissaguere² defined early repolarization patterns (ERP) as a J point elevation (QRS-ST

Funding: We did not receive any funding for this study.

© 2014 Wiley Periodicals, Inc. DOI:10.1111/anec.12231

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Figure 1. Slurring and notching type appearance of early repolarization pattern.

junction), which can take the form of slurring or notching greater than or equal to 0.1 mV in at least two lower and/or lateral leads (II, III, aVF, I, aVL, V_4-V_6) excluding leads V_1-V_3 and associated with a risk of SD by ventricular fibrillation (VF; Fig. 1). The screening of subjects with ERP at risk of SD is difficult because this pattern is common in the general population (3–13%) unlike the occurrence of a VF in a healthy heart.²⁻⁵ Thus, this pattern is not sufficient in itself to identify patients at high risk of SD. Risk stratification happens in practice through investigation of clinical variables and ECG. Clinical evidence of malignancy is represented by a family history of SD before the age of 45 and unexplained syncope.⁶ Derogatory ECG elements combine the dynamic character and a J point amplitude greater than 0.2 mV, the appearance and distribution of a I point in the form of notching in the lower and inferolateral leads and the appearance of a horizontal or descending ST segment.³⁻⁸ The prevalence of ERP in a population of young, athletic military officers has been estimated at 13% and until 2008 its discovery did not lead to further cardiovascular explorations or restriction of employment or physical activity.⁹ Since then, our practical attitude comes down to a risk stratification of SD among these young military officers based on clinical variables and ECGs. The continuation of the military officer in his professional activity depends on the results of this evaluation.

METHODS

Main Objective

The main objective of the study is to estimate the prevalence, appearance, distribution, and evolution of ERP after 5 years according to Haissaguere's criteria² in a specific population for whom we are medically liable, to optimize our decisions around fitness to work within the framework of medical expertise.

Population Studied and Study Conduct

From March to December 2008, we prospectively collected the ECGs of military officers enlisted into the submarine forces of France under the following inclusion criteria: subjects aged over 17 years, Caucasians and practicing at least two hours of sport per week. They had to be asymptomatic in cardiovascular terms (no unexplained syncope) and not have a family history of SD before the age of 45 years. They could not present cardiovascular risk factors such as hypertension (blood pressure >140/90 mmHg), dyslipidemia (total cholesterol >200 mg/dL), diabetes (fasting glucose >126 mg/dL), active smoking, or body mass index (BMI) greater than 30 kg/m.²

The exclusion criteria are as follows: all symptomatic or asymptomatic subjects with ECG abnormalities such as a complete or incomplete branch block, an atrioventricular block, or an atrial fibrillation were excluded from the study.

For 5 years (until 2012), the military officers included in the study underwent a clinical examination twice per year. A new ECG was carried out 5 years after the initial one. The first series of ECGs carried out in 2008 is denoted P1; the second series of ECGs carried out 5 years later (2012) is denoted P2.

ECG Reading Phase

For each period P1 and P2, the ECGs were carried out at rest, lying down, with 12 standard leads, and a rate of 25 mm/s. The ECGs were anonymous and read separately by two cardiologists in the first instance. The presence of ERP was denoted RP+, the absence of ERP was denoted RP-, and a borderline appearance denoted as "borderline." In the second instance, the ECGs were reread together. The results were validated by the agreement of three readings. In the case of "borderline" appearance, discrepancy and in the absence of consensus, the ECGs were classified as RP-. The appearance and distribution of the J point were specified as well as the appearance of ST elevation.

Statistical Analysis

Descriptive statistics are presented after calculation of means and percentages and determination of confidence intervals at 95% (binomial formula). In terms of analytical statistics, the percentages were compared using the Fisher's exact test or chi-square test, after verification of performance conditions. The comparisons of means were carried out by the Student's *t*-test, after verification of performance conditions (equality of variances). The comparisons of means on paired series were also carried out using the Student's *t*-test. The level of statistical significance was set at 5% (P < 0.05).

RESULTS

During the study period, 250 male subjects were included. The mean age of the subjects was 22.87 \pm 0.5 years. The general characteristics of the population are summarized in Table 1. During the 5 years of follow-up, the population remained free of cardiovascular risk factors and individuals

continued regular exercise >2 hours per week. All subjects remained asymptomatic. No SD occurred.

Period P1: Analysis of ECGs

The prevalence of RP+ was 19.2% or 48 cases. There was no statistically significant difference between the RP+ and RP- groups in terms of age. RP+ subjects had a statistically significantly lower heart rate, a shorter QTc, and a higher Sokolow index (Table 2). The most common appearance of a J point was the slurring type (31 of 48 or 64.5%) in the inferolateral branches (18/48 or 37.8%). The appearance of a notching type J point was only found in 35.5% of cases. ST elevation was ascending in 60.4% of cases. ST segment elevation greater than 2 mm was not found. These results are summarized in Tables 2 and 3.

Period P2: Analysis of ECGs

Five years later, the prevalence of ERP was identical to the previous period (19.2%). RP+ subjects also had a statistically significant shorter QTc and higher Sokolow index. There was no significant difference in the HR. The predominant appearance of a J point was slurring for 32 cases (66.7%) and inferolateral localization (19 cases or 39.5%). The ST segment elevation was less than 2 mm in all cases. It was ascendant in 58.3% and horizontal in 41.7%. The results are summarized in Tables 2 and 3.

Evolution of ERP

There was identical persistence in terms of distribution, appearance, and amplitude of RP+ in 47 subjects (97.9%). Heart rate is generally higher for ECG P1 than for ECG P2 without any statistically significant difference (Fig. 1). There was no significant difference between the ECG ERP+ P1 and P2 in terms of the QTc and Sokolow index. Repolarization changed in two patients. We noted the disappearance of ERP in P2 with an almost identical HR in the ECGs of P1 and P2 $(HR_{P1} = 61 \text{ bpm}, HR_{P2} = 62 \text{ bpm})$. We found an appearance of ERP in one subject in P2 with a lower HR reading in P1 compared to P2 ($HR_{P1} =$ 75 bpm; $HR_{P2} = 100$ bpm). The appearance of a J point was inferolateral slurring associated with an ST elevation of >1 mm horizontal. The results are summarized in Table 4.

Table 1. General Characteristics of the Study Population

Total	N = 250
Age (years)	22.87 ± 0.5
Weight (kg)	76.7 ± 1.5
Height (m)	1.77 ± 0.1
Heredity/hypertension/diabetes/dyslipidemia/smoking	0
Family history of sudden death/unexplained syncope	0
Sport practice >2 hours/week	250 (100%)

Population	Total, n = 250	ARP±, n = 48 (19.2%)	ARP-, n = 202 (80.8%)	Р
		Period P1		
Age	22.87 (22.4–23.35)	22.65 (22.3-23.4)	23.7 (22.9–24)	0.07
EČG characteristics	. ,			
FC (BPM)	68.5 (±12.7)	66 (±13.5)	71.7 (±13)	0.008
OTc Bazett (ms)	388 (±29.2)	382.7 (±33)	394.3(±29)	0.008
HVG Sokolow (mm)	29 (±0.7)	31.04 (±0.6)	27.6(±1.6)	0.019
		Period P2		
Age	27.87 (27.4–28.35)	27.68 (27.3–28.4)	28.78 (27.9–29)	0.1
ECG characteristics				
FC (BPM)	66.8 (±12.7)	65.8 (±13.5)	67.6 (±13)	0.03
QTc Bazett (ms)	392.2 (±30.2)	388.1 (±33)	396.4 (±31)	0.01
HVG Sokolow (mm)	28.5 (±0.6)	30.4 (±1.2)	26.97 (±0.7)	0.019

 Table 2. Clinical and ECG Characteristics in the Periods 1 and 2 (P1–P2)

Table 3. Appearance-Distribution of J Point and Segment ST Elevation (P1 and P2)

	RP+ P1 (n = 48/19.2%)	RP+ P2 (n = 48/19.2%)	Р
HR (bpm)	66.0	65.8	>0.05
QTc Bazett (ms)	382.7	387.9	>0.05
HVG Sokolow (mm)	31.0	30.5	>0.05
Appearance-distribution of point J			
Inferolateral slurring (D2D3Vf-V5V6)	18 (37.8%)	19 (39.5%)	>0.05
Lateral slurring (DIaVI-V5V6)	7 (14.5%)	7 (14.5%)	>0.05
Lower slurring (D2D3Vf)	6 (12.5%)	6 (12.5%)	>0.05
Inferolateral notching (D2D3Vf-V5V6)	8 (16.6%)	8 (16.6%)	>0.05
Lateral notching (DIaVI-V5V6)	5 (10.5%)	4 (8.3%)	>0.05
Lower notching (D2D3Vf)	4 (8.3%)	4 (8.3%)	>0.05
Appearance of ST segment elevation			
Ascending ST elevation	29 (60,4%)	28 (58,4%)	>0.05
Horizontal or descending ST elevation	19 (39.6%)	20 (41.6%)	>0.05

DISCUSSION:

Prevalence of ERP

In our selected healthy population, we found an ERP prevalence of 19.2%, while this has been estimated between 3.3% and 13.1% in the general population. In 2008, Haissaguere determined a prevalence of 5% in a control population of 412 patients whose mean age was 36.5 ± 12 years². Tikkanen in a cohort of 10,864 patients found a prevalence of ERP of 5.8% (7.1% in men and 4.6% in women) with a mean age of 44 ± 8.5 years.³ The MONICA/KORA study found a prevalence of 13.1% in a population of 6213 patients with a mean age of 52 ± 10 years.⁴ In 2011, Noseworthyl also showed in the study of two large American

and European cohorts comprising 3995 and 5489 patients, respectively, an ERP prevalence of 4.5% across both populations. The prevalence was 6.1% in the U.S. population and 3.3% in the European population.⁵ Our young and athletic population therefore does not strictly represent the general population described in the literature. Nor can we compare with a population of elite athletes as studies on this population show a higher prevalence of ERP at 20%. Noseworthy, in a study including 879 athletes, showed an ERP prevalence of 25.1% (male athletes: 33% female athletes: 11%).10 Tikkanen conducted a study involving 62 Finnish athletes aged 13-15 years and 503 Americans aged 17-24 years.⁷ Among Finnish athletes, the prevalence of ERP was 44%, while it was estimated at 30% among Americans. These different clinical studies on the general population and athletes show that the presence of ERP is influenced by genetic and clinical factors. Thus, the male predominance and geographical distribution of ERP would support a genetic contribution.¹¹ Age, physical activity, and vagal tone are the factors influencing the presence of an ERP with a decline in prevalence as the patient's age increases. In practice, this translates into a more common ERP in athletic males whose ECG is characterized by a lower HR and a shorter QT interval than subjects without ERP. This translates in our population as an increased prevalence of ERP.

Persistence of ERP

Elevation of the J point defining the appearance of ERP is dynamic and variable over time.² An elevated I point was therefore shown shortly before the occurrence of an electrical storm in patients with an implantable cardioverter defibrillator (ICD) as secondary prevention of SD in a healthy heart.¹² The dynamic nature of ERP was also demonstrated by Noseworthy in a healthy population of elite athletes. His study showed an increased prevalence of ERP (37.2%) in a population of 146 male athletes (soccer and rowing) and a significant increase in this (52.7%, P = 0.003) after a training period of 90 days (\approx 20 hours/week).¹⁰ In the general population, ERP is variable over time and regresses with age. Indeed, Stein and Adhikarla showed a regression in ERP in more than 50% of cases in 244 patients (mean age 42 \pm 13 years) from a cohort of 29,281 subjects after 10 years of follow-up.13,14 In our

study, ERP persisted in 97.9% of patients without any change in the appearance, amplitude, and distribution of point J. These differing results in the literature can be explained by the lack of influence of aging in our population over 5 years. They ultimately remained young (<40 years) and statistically identical at the two periods (sport and HR).

ERP Prognosis after 5 Years

Over the period of 5 years, no clinical event such as death, syncope, or SD occurred. Clinically, our population at baseline had no clinical risk criteria of SD. The most common ECG appearance of repolarization was a slurring type point J elevation <2 mm (64.5%) in the lower, lateral, and inferolateral leads, associated with an ascending ST segment elevation. This appearance is suggests a mild form. Indeed, Tykkanen showed that patients with a point J elevation higher than 2 mm have three times the risk of death from arrhythmias compared to a control population.³ The notching appearance of the point I in the lower and inferolateral branches is more often present in patients with a VF.8 Finally, RP+ patients associated with a horizontal or descending ST segment elevation had a poor prognosis.⁷

Implication in Medical Expertise

The discovery of an ERP in our military population requires us to have a pragmatic approach in medical prevention and expertise. Screening patients at risk of SD associated with ERP is achieved through a specific examination and a rigorous interpretation of the ECG, in particular investigating the previously described malignancy criteria. Upon the discovery of an ERP in an asymptomatic military officer and in the absence of malignancy criteria, there will be no restriction in terms of fitness to work or physical activity. In the presence of SD in the family and/or unexplained syncope, officers undergo further investigations in a military hospital cardiology service (pharmacological tests, programmed ventricular stimulation). The symptomatic RP+ military officer may be limited in his professional practice if it presents specific risks (pilot, paratrooper, commando). The symptomatic RP+ military officer in secondary prevention of SD has an ICD, but can no longer carry out his professional activity.¹²

Study Limitations

This is a single-center study of a specific population. The findings may not fully apply to the general population. The design of our study does not provide the continuous appearance of repolarization because it involves two ECG recordings carried out 5 years apart (ECG P1 and P2). The information is therefore incomplete. We provided an observational ECG study of ERP in military recruits, which lacks prognostic value and long-term follow-up data. The surface and implantable Holters allow continuous analysis thereof as well as specific information, in particular vagal tone information.

ERP is commonly found in young, athletic military officers, free of heart disease and cardiovascular risk factors. This ERP persists over 5 years and has a good prognosis. In our professional practice, the different periodic medical visits carried out on the military officers include evaluation of the risk of SD associated with ERP by examination and regular ECGs.

Acknowledgments: This study was approved by the army medical service ethics committee.

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