The Peripheral Arterial disease study (PERART/ARTPER): prevalence and risk factors in the general population

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Abstract

Background: The early diagnosis of atherosclerotic disease is essential for developing preventive strategies in populations at high risk and acting when the disease is still asymptomatic. A low ankle-arm index is a good marker of vascular events and may be diminished without presenting symptomatology (silent peripheral arterial disease). The aim of the study is to know the prevalence and associated risk factors of peripheral arterial disease in the general population.

Methods: We performed a cross-sectional, multicentre, population-based study in 3786 individuals >49 years, randomly selected in 28 primary care centres in Barcelona (Spain). Peripheral arterial disease was evaluated using the ankle-arm index. Values < 0.9 were considered as peripheral arterial disease.

Results: The prevalence (95% confidence interval) of peripheral arterial disease was 7.6% (6.7-8.4), (males 10.2% (9.2-11.2), females 5.3% (4.6-6.0); p < 0.001).

Multivariate analysis showed the following risk factors: male sex [odds ratio (OR) 1.62; 95% confidence interval 1.01-2.59]; age OR 2.00 per 10 years (1.64-2.44); inability to perform physical activity [OR 1.77 (1.17-2.68) for mild limitation to OR 7.08 (2.61-19.16) for breathless performing any activity]; smoking [OR 2.19 (1.34-3.58) for former smokers and OR 3.83 (2.23-6.58) for current smokers]; hypertension OR 1.85 (1.29-2.65); diabetes OR 2.01 (1.42-2.83); previous cardiovascular disease OR 2.19 (1.52-3.15); hypercholesterolemia OR 1.55 (1.11-2.18); hypertriglyceridemia OR 1.55 (1.10-2.19). Body mass index ≥ 25 Kg/m2 OR 0.57 (0.38-0.87) and walking >7 hours/week OR 0.67 (0.49-0.94) were found as protector factors.

Conclusions: The prevalence of peripheral arterial disease is low, higher in males and increases with age in both sexes. In addition to previously described risk factors we found a protector effect in physical exercise and overweight.

Background

The prevention and early diagnosis of atherosclerotic disease is one of the essential objectives in the field of cardiovascular disease since it is the main cause of mortality in developed countries. In the European Union these diseases represent approximately 40% of the deaths in both men and women [1]. Atherosclerosis is currently considered a chronic, progressive systemic disease of multifactorial aetiology [2] including arterial hypertension, hypercholesterolemia, diabetes mellitus and smoking as modifiable risk factors and age and sex as non modifiable factors [3]. These factors have been integrated in prediction tables based on regression models with the aim of detecting the population with a high risk of cardiovascular events [4]. However, the sensitivity and positive predictive value of these tables is low, and thus, most cardiovascular disease are produced in subjects without a high risk [4]. It is therefore important to develop markers of silent atherosclerotic disease which...
will help to better identify subjects at high risk of developing peripheral arterial disease in order to implement preventive measures. In addition, atherosclerotic disease remains clinically silent during most of the evolutionary process until phenomena of complications of atheroma plaques suddenly appear and lead to ischaemic vascular events. Therefore, from a primary prevention point of view, it is very important to develop strategies which allow identification of patients with atherosclerosis in subclinical stages, with primary care being the optimum setting for accessibility.

Peripheral arterial disease is a common manifestation of atherosclerosis and is characterized by increasing incidence of morbi-mortality [5]. Peripheral arterial disease is a powerful predictor of cardio and cerebrovascular events and is associated with an increase in mortality of up to 30% at 5 years and 50% at 10 years [5-7] thereby making early detection of this disease fundamental.

The prevalence of peripheral arterial disease varies greatly depending on the population studied, on the definition of peripheral arterial disease used (symptomatic or not), the diagnostic method, age, sex and the presence of other risk factors. Most studies have been carried out in populations at high risk, with few having been performed in countries of low risk [8-16]. The prevalence of peripheral arterial disease in countries with low cardiovascular risk is uncertain.

Underdiagnosis of peripheral arterial disease is very high since a very large proportion of these patients is asymptomatic [17]. The ankle-arm index is the most effective tool used to screen for peripheral arterial disease [18,19]. Compared with angiography, an ankle arm index <0.9 presents a sensitivity of 95% and a specificity of 99% for the detection of stenosis ≥50% [20], with the reliability of this test being good when performed by trained personnel [18-21].

The wide range of prevalence reported in the literature and the low number of studies in countries of low risk such as Spain led to the design of this study, the aim of which was to know the prevalence of peripheral arterial disease using the calculation of the ankle arm index in a general population over the age of 49 years and to determine the risk factors associated with a pathologic ankle arm index.

Methods

A detailed description of the methodology of the study has been published elsewhere [22].

Briefly, this was a cross-sectional, multicentre, descriptive, population-based study aim at determining the prevalence of symptomatic and asymptomatic peripheral arterial disease and related factors in a general population ascribed in primary care. Individuals were randomly selected (simple random sampling) from a database, which is more exhaustive and updated than the census, containing the population ascribed in the centers participating in the study. The randomly selected individuals were invited to participate in the study by phone. If the individuals were not found, up to 5 calls were made at different hours and days of the week to contact them. After agreement to participate subjects were given an appointment to perform an interview, blood sample extraction and anthropometric measurements, including ankle-arm index.

A total of 28 primary healthcare centers within the metropolitan area of the city of Barcelona and the county of Barcelonès Nord-Maresme, including urban and semi-rural centers, participated in the study. Finally, from September 2006 to June 2008, a total of 3786 patients over the age of 49 years were included in the study.

Data collection

The ankle arm index examination of the subjects was carried out in the participating centres by two healthcare professionals trained in the technique, under standardized conditions. A standardized Doppler Ultrasonic device was used (Mini-Dopplex D 900-P, Huntleigh Healthcare, 8 MHz). Ankle-arm index was performed in the two paramalleolar arteries of both lower extremities. For each leg the ankle-arm index was the ratio of the higher of the two systolic pressures (tibial posterior and anterior artery) and arm systolic pressure using the systolic pressure of the highest arm. If the ankle-arm index was < 0.9 the technique was performed by the other professional. In cases in which the second professional found an ankle arm index ≥ 0.9 the first repeated the test and the latter value was considered as the final result.

The following variables were collected: demographic and lifestyle, including smoking and physical activity with a validated questionnaire, self-reported and clinical history of previous cardiovascular disease (acute myocardial infarction, angina, stroke and transient ischaemic attack), intermittent claudication, hypercholesterolemia, diabetes mellitus, arterial hypertension; anthropometric variables (height, weight, and waist circumference), blood analysis (total cholesterol, HDL-cholesterol, triglycerides, and glycaemia); cardiovascular risk using the Framingham-Wilson equations, Framingham calibrated by the REGICOR and SCORE groups; metabolic syndrome (NCEP criteria) [23]; and the Edinburgh vascular questionnaire [8,15].

Statistical analysis

The prevalence of peripheral arterial disease was computed in the whole sample. However, patients with
Mönckeberg sclerosis, indicated as an ankle arm index >1.4 (arterial calcification) were excluded from further analyses involving potential associated risk factors with peripheral arterial disease. Comparison of categorical variables was performed using chi squared tests and for continuous variables the t-test was used. In addition, non parametric tests were used when distribution requirements of the variables analyzed were not fulfilled. Logistic regression models (LRM) were adjusted using peripheral arterial disease as the dependent variable. The individual effect, measured with the odds ratios [OR, (95% confidence interval)] of the potential risk factors, was studied in age- and sex-adjusted LRM. Multivariate LRM were thereafter constructed to assess the independent effects of each variable adjusted for the other in the model. Best models were selected taking into account the correlation between variables and the Akaike Information Criteria (AIC) [24]. Only variables with p < 0.05 remained in the model. Significance was defined with a p value less than 0.05. All tests were bilateral. Analysis was performed using Stata version 10 (StataCorp, College Station, TX, USA, 2007).

Ethics
This study was approved by the local Ethics Committee (IDIAP Jordi Gol Foundation of Investigation in Primary Care and Instituto de Salud Carlos III). Informed written consent was obtained from all the participants. Likewise, the recommendations of the World Medical Association Declaration of Helsinki were followed.

Ethical considerations
The authors of this manuscript have certified that they comply with the Principles of Ethical Publishing in BMC Public Health.

Results
The percentage of participation in the study was of 63%, with 3786 participants >49 years [1746 men (46.1%) and 2040 women (53.9%)] with a mean age of 64.9 years ± 8.9 being included. Of these, 286 patients had an ankle-arm index < 0.9. The prevalence of peripheral arterial disease in the total population was 7.6% (95% confidence interval 6.7-8.4) (men 10.2% (9.2-11.2), women 5.3% (4.6-6.0); p < 0.001). For each 10 years of increased age, the prevalence of peripheral arterial disease doubled. The results of the present study indicate that the prevalence of peripheral arterial disease using the ankle-arm index < 0.9 was 7.6% (95% confidence interval 6.7-8.4) (men 10.2% (9.2-11.2), women 5.3% (4.6-6.0); p < 0.001). For each 10 years of increased age, the prevalence of peripheral arterial disease doubled.

Risk factors for the presence of peripheral arterial disease according to logistic regression analysis
Table 2 shows the results of the individual analysis of the risk factors associated with the presence of peripheral arterial disease according to the LRM adjusted for age, sex and centre and table 3 describes the results of the multivariate analysis according to gender. Walking more than 7 hours weekly (OR 0.67) and overweight/obesity (OR 0.57) were protector factors on multivariate analysis, although on stratification by sex, they were only statistically significant in women. Male gender (OR 1.62), age (OR 2.00 per 10 years), difficulty in performing physical exercise (OR 7.08 breathlessness performing any activity) and smoking (OR 3.83 for current smokers) were positively associated with peripheral arterial disease in both sexes. Hypertension (OR 1.85), diabetes (OR 2.01) and previous CDV (OR 2.19) were associated with peripheral arterial disease overall and among men, and hypercholesterolemia (OR 1.55) and high triglycerides (OR 1.55) were associated with peripheral arterial disease overall and among women only. Variables only significant for one gender in the multivariate analysis showed a similar trend in both sexes. Although education, occupation and IC were associated with peripheral arterial disease in the age- and sex-adjusted models, these factors had no significant effect in the multivariate models. Stroke, transient ischaemic attack, angina or myocardial infarction were related to peripheral arterial disease in the age- and sex-adjusted model but were collapsed into a single cardiovascular disease variable in the multivariate model providing better AIC indexes. Information regarding cardiovascular disease risk tables or metabolic syndrome was not included in the multivariate model since most of their components were already included. Nevertheless, a significant association (table 2) was observed between the metabolic syndrome and peripheral arterial disease [OR 1.79 CI 95% (1.36-2.36)] and these variables showed a positive and significant association with peripheral arterial disease in the age and sex adjusted models.

Discussion
The results of the present study indicate that the prevalence of peripheral arterial disease using the ankle-arm
index (7.6%) is lower in a country of low cardiovascular risk such as Spain in relation to other countries with a higher risk [8-13]. Thus, our results coincide with studies which have reported lower values of cardiovascular disease such as ischaemic heart disease [25]. In another Spanish study [15], the prevalence of peripheral arterial disease was lower (4.5%). Nonetheless, the age of the subjects studied was from 35 to 79 years which could explain the difference with our results considering that it is well known that the prevalence of cardiovascular disease increases with age [7].

The prevalence of peripheral arterial disease in the general population is not well known in countries of low risk. In Spain, all the previous studies have been undertaken in populations at high risk or well selected for different factors: male sex, smokers, diabetics, metabolic syndrome or previous cardiovascular disease. In these studies the prevalence ranged from 3.9% to 26.2% according to the population studied [25-30], with the high prevalence among hospitalized (26.2%) [27] and diabetic patients (21.4%) [30] being of note. The low prevalence of peripheral arterial disease in countries with low cardiovascular risk [25] but a high prevalence of cardiovascular risk factors (French paradox) could be explained by protective factors in Mediterranean countries, such as dietetic factors. Studies in Spain have confirmed this low prevalence of peripheral arterial disease [15,16], coronary heart disease [31,32] and stroke [33,34] compared with studies conducted in Northern Europe.

One alarming data of our study was the under diagnosis of peripheral arterial disease of 81.0% which, according to some authors, may be attributed to the large number of patients (80-90%) who remain asymptomatic and that the ankle arm index is not routinely evaluated in primary care offices. It is important to note that in the present study 33.8% of the patients with a pathologic ankle arm index presented clinical manifestations of IC, as defined by the Edinburgh questionnaire [8,35,36]. The patients may attribute this symptomatology to other diseases or pathological processes. In most of the studies published, the IC values among patients with peripheral arterial disease were between 5% and 29.6% [10,12], with the exception of the study by Coni et al. [37] in which the prevalence of IC was very high (37.5%) and similar to that of our study, although, in contrast with the previous study, we used the Edinburgh questionnaire.

The proportion of patients with arterial calcification was of note (6.2%). Its clinical significance has been
### Table 1 Sample characteristics by peripheral arterial disease

| Category                                                        | PAD      | Control*                     | p-valueb |
|----------------------------------------------------------------|----------|------------------------------|----------|
|                                                                | n = 286  | n = 3265                     |          |
| Gender                                                         |          |                              | < 0.001  |
| Men                                                           | 11.1 (178)| 88.9 (1419)                  |          |
| Women                                                          | 5.5 (108)| 94.5 (1846)                  |          |
| Age (years) (mean ± SD)                                       | 70.70 ± 9.26| 64.22 ± 8.60                | < 0.001  |
| Education                                                      |          |                              | < 0.001  |
| Illiterate                                                     | 14.2 (30)| 85.8 (182)                   |          |
| Primary school                                                | 8.1 (196)| 91.9 (2229)                  |          |
| Secondary school                                              | 5.4 (27)| 94.6 (477)                   |          |
| High school                                                   | 5.7 (7)| 94.3 (116)                   |          |
| University                                                    | 3.4 (4)| 96.6 (112)                   |          |
| Occupation status                                             |          |                              | < 0.001  |
| Currently working                                             | 3.6 (30)| 96.4 (813)                   |          |
| Housewife                                                     | 4.7 (34)| 95.3 (689)                   |          |
| Retired                                                       | 12.2 (192)| 87.8 (1380)                 |          |
| Unemployed                                                    | 3.9 (6)| 96.1 (148)                   |          |
| Disabled                                                      | 11.1 (15)| 88.9 (120)                  |          |
| Other                                                         | 6.3 (3)| 93.8 (45)                    |          |
| Ability to perform physical activity                          |          |                              | < 0.001  |
| No limitation                                                 | 3.8 (51)| 96.2 (1291)                  |          |
| Mild limitation                                               | 8.5 (150)| 91.5 (1618)                 |          |
| Only able to do light activity                                | 19.9 (72)| 80.1 (289)                   |          |
| Breathless with any activity                                  | 29.3 (12)| 70.7 (29)                    |          |
| Walking (hours/week)                                          |          |                              | 0.376    |
| 0-3                                                           | 7.2 (65)| 92.8 (844)                   |          |
| >3-7                                                          | 8.0 (61)| 92.0 (697)                   |          |
| >7                                                            | 6.4 (86)| 93.6 (1252)                  |          |
| Tobacco smoking                                               |          |                              | < 0.001  |
| Never smoker                                                  | 5.3 (104)| 94.7 (1871)                 |          |
| Former smoker                                                 | 11.3 (108)| 88.7 (844)                  |          |
| Current smoker                                                | 11.9 (74)| 88.1 (550)                  |          |
| Claudication (people with leg pain when walking and no pain when stop) |          |                              | < 0.001  |
| No                                                            | 5.9 (184)| 94.1 (2952)                  |          |
| Yes                                                           | 26.9 (94)| 73.1 (255)                   |          |
| Obesity                                                       |          |                              | 0.164    |
| Underweight/Average (BMI<25)                                  | 9.7 (61)| 90.3 (566)                   |          |
| Overweight (25≤BMI<30)                                        | 7.3 (118)| 92.7 (1496)                  |          |
| Obese (BMI ≥ 30)                                              | 8.2 (107)| 91.8 (1198)                  |          |
| Waist circumference                                           |          |                              | 0.035    |
| 1st tertile<sup>c</sup>                                       | 6.9 (78)| 93.1 (1060)                  |          |
| 2nd tertile                                                  | 7.6 (89)| 92.4 (1076)                  |          |
| 3rd tertile                                                  | 9.7 (118)| 90.3 (1104)                  |          |
| Hypertension (medical record)                                 |          |                              | < 0.001  |
| No                                                            | 4.7 (88)| 95.3 (1791)                  |          |
| Yes                                                           | 11.9 (193)| 88.1 (1431)                 |          |
| Hypercholesterolemia (medical record)                         |          |                              | < 0.001  |
| No                                                            | 6.0 (108)| 94.0 (1691)                  |          |
| Yes                                                           | 10.2 (170)| 89.8 (1501)                  |          |
little studied. Nonetheless, it has recently been shown to be associated with a greater risk of morbi-mortality although not as important as an ankle arm index < 0.9. These patients were excluded from the analysis of the associated factors to allow better comparability among healthy patients and those with peripheral arterial disease. This will be studied in the future.

Similar to other studies, the present series found a higher prevalence of peripheral arterial disease in men which progressively rose with age in both sexes. In women, a protector effect was found with physical exercise (walking >7 hours per week) and overweight, although this was not statistically significant in men. This protector effect of physical exercise may be justified by a greater development of collateral circulation, which is currently one of the therapeutic recommendations in these patients. The protector effect of obesity and overweight independent of physical exercise and other risk factors may be surprising, but a better prognosis has also been described in patients with obesity in

### Table 1: Sample characteristics by peripheral arterial disease (Continued)

| Blood analysis (mean ± SD) | 210.03 ± 42.10 | 216.53 ± 38.48 | 0.007 |
|---------------------------|----------------|----------------|-------|
| Total cholesterol (mg/dL) | 52.30 ± 14.09  | 56.03 ± 14.48  | < 0.001 |
| Diabetes (medical record) | Normal 6.9 (181) | 93.1 (2447) | < 0.001 |
| Stroke (medical record)   | No 64.0 (191)   | 93.6 (2802)    | < 0.001 |
| Angor (medical record)    | No 7.7 (266)    | 92.3 (3167)    | < 0.001 |
| Myocardial infarction (medical record) | No 7.2 (245) | 92.8 (3148) | < 0.001 |
| Framingham index (aged≤74) | <10 2.9 (32)    | 97.1 (1081)    | < 0.001 |
| REGICOR index (aged≤74)   | <5 3.1 (41)     | 96.9 (1295)    | < 0.001 |
| SCORE index (aged<65)     | <2.5 2.4 (27)   | 97.6 (1105)    | < 0.001 |
| <5 7.0 (89)               | 93.0 (1177)    | 0.001 |
| >5 15.0 (48)              | 85.0 (271)     | 0.001 |
| Metabolic syndrome (NCEP) | No 7.2 (198)    | 92.8 (2559)    | < 0.001 |
| Yes 11.4 (88)             | 88.6 (685)     | 0.001 |

All values are presented as prevalence of healthy and peripheral arterial disease as percent (n) unless otherwise specified. SD: standard deviation. BMI: Body mass index expressed in Kg/m².

a Patients with calcification excluded.

b p value for a chi-squared test (categorical variable) and t-test (continuous variable). Fisher exact test and rank sum tests have also been performed for categorical and continuous variables respectively leading to similar p values.

c Sex-specific tertiles. Cut points are 97 cm and 105 cm for men and 90 cm and 101 cm for women.
Table 2 Association between peripheral arterial disease and potential risk factors. Logistic regression models adjusted by age and sex

| Risk Factor                                | OR (95%CI)          | p-value for trend |
|--------------------------------------------|---------------------|------------------|
| **Men**                                    |                     |                  |
| Age (per 10 years)                         | 2.25 (1.96-2.60)    |                  |
| Education (reference = Illiterate)         |                     |                  |
| Primary school                             | 0.65 (0.42-1.00)    |                  |
| Secondary school                           | 0.55 (0.31-0.99)    |                  |
| High school                                | 0.48 (0.20-1.16)    |                  |
| University                                 | 0.29 (0.10-0.87)    |                  |
| Occupation status (reference = Currently working) |         |                  |
| Housewife                                  | 1.02 (0.57-1.83)    |                  |
| Retired                                    | 1.14 (0.70-1.86)    |                  |
| Unemployed                                 | 1.10 (0.45-2.71)    |                  |
| Disabled                                   | 2.50 (1.29-4.88)    |                  |
| Other                                      | 1.76 (0.51-6.05)    |                  |
| Ability to perform physical activity (reference = No limitation) | | < 0.001 |
| Mild limitation                            | 2.07 (1.47-2.92)    |                  |
| Only able to do light activity             | 4.84 (3.19-7.36)    |                  |
| Breathless with any activity               | 6.70 (3.10-14.45)   |                  |
| Walking (hours/week) (reference = 0-3)     |                     | 0.002            |
| >3-7                                       | 0.88 (0.60-1.29)    |                  |
| >7                                         | 0.58 (0.40-0.82)    |                  |
| Tobacco smoking (reference = Never smoker) |                     | < 0.001          |
| Former smoker                              | 2.14 (1.45-3.17)    |                  |
| Current smoker                             | 3.91 (2.58-5.93)    |                  |
| Claudication                               | 5.00 (3.73-6.71)    |                  |
| Obesity (reference = BMI<25)               |                     | 0.685            |
| Overweight (25≤BMI<30)                     | 0.67 (0.48-0.94)    |                  |
| Obese (BMI ≥30)                            | 0.85 (0.60-1.20)    |                  |
| Waist circumference                        |                     | 0.174            |
| 2nd tertile                                | 1.02 (0.73-1.41)    |                  |
| 3rd tertile                                | 1.23 (0.90-1.67)    |                  |
| Hypertension                               | 1.99 (1.51-2.62)    |                  |
| Hypercholesterolemia                       | 1.86 (1.43-2.41)    |                  |
| Triglycerides ≥ 150 mg/dL                  | 2.05 (1.58-2.66)    |                  |
| Diabetes                                   | 2.39 (1.82-3.15)    |                  |
| Stroke                                     | 2.92 (1.74-4.90)    |                  |
| Transient ischaemic attack                 | 1.73 (0.93-3.20)    |                  |
| Angor                                      | 1.90 (1.23-2.93)    |                  |
| Myocardial infarction                      | 3.81 (2.50-5.82)    |                  |
| Framingham index (aged≤74)                 |                     | < 0.001          |
| 10-20                                      | 1.42 (0.88-2.29)    |                  |
| >20                                        | 2.57 (1.49-4.43)    |                  |
| REGICOR index (aged≤74)                    |                     | < 0.001          |
| 5-10                                       | 1.60 (1.05-2.43)    |                  |
| >10                                        | 2.61 (1.55-4.38)    |                  |
heath failure and a lower incidence of acute myocardial infarction [39,40]. However, we did not find any relationship with waist circumference, although on adjusting for the body mass index (BMI) we did find a relationship with the second [OR 1.25 (0.87-1.81)] and the third tertile [OR 1.56 (1.02-2.37)] compared to the first tertile, which disappeared on adjustment for the remaining variables in the multivariate model.

With regard to studies evaluating the association with different factors, our results coincide in age [8-13], male gender [8-13], hypertension [9-11], diabetes [11,12], smoking [10-12], and history of cardiovascular disease [9,10]. In the present study we found an association with hypercholesterolemia and hypertriglyceridemia in women. In other studies [11,12], an association has also been reported with hypercholesterolemia, while the results related to triglycerides in the literature are contradictory [9].

With respect to limitations of the study, as usually occurs in population-based studies in Spain, women were slightly over-represented [14]. Although the data from the census were not used, in Spain the use of the population assigned to the primary care centres is preferable since it is more exhaustive and updated than that of the census [41]. On the other hand, the cross-sectional design of the study did not allow determination of the causal effect of the different factors associated with peripheral arterial disease.

The technical ease of the ankle-arm index and its ready adoption in the primary care offices has replaced the use of carotid echo-Doppler for detecting patients with high cardiovascular risk. However, a large proportion of the population has an intermediate risk, thus it remains to be defined in which patients ankle-arm index should be a priority. According to the present study these patients would include: males, aged over 60 years, smokers or

| Table 3 Association between peripheral arterial disease and potential risk factors. Multivariate logistic regression models by sex. |
|-----------------------------------------------|
| **Total** | **Men** | **Women** |
| **OR (95%CI)** | **OR (95%CI)** | **OR (95%CI)** |
| Men | 1.62 (1.01-2.59) | 1.83 (1.42-2.36) | 2.17 (1.52-3.09) |
| Age (×10 years) | 2.00 (1.64-2.44) | 1.77 (1.17-2.68) | 1.36 (1.11-2.19) |
| PA Mild limitation | 3.64 (2.16-6.13) | 4.09 (2.12-7.91) | 2.93 (1.97-5.90) |
| PA Only able light activity | 7.08 (2.61-19.2) | 7.47 (1.87-29.8) | 5.40 (1.14-25.7) |
| >7 h/w walking | 0.67 (0.49-0.94) | 0.82 (0.55-1.23) | 0.43 (0.22-0.82) |
| Former smoker | 2.19 (1.34-3.58) | 2.48 (1.26-4.90) | 1.47 (0.52-4.11) |
| Current smoker | 3.83 (2.23-6.58) | 4.18 (2.01-8.69) | 4.00 (1.59-10.1) |
| BMI ≥25 Kg/m² | 0.57 (0.38-0.87) | 0.65 (0.39-1.08) | 0.43 (0.20-0.92) |
| Hypertension | 1.85 (1.29-2.65) | 1.78 (1.16-2.72) | 2.03 (0.99-4.14) |
| Hypercholesterolemia | 1.55 (1.11-2.18) | 1.36 (0.90-2.07) | 1.87 (1.03-3.40) |
| High triglycerides | 1.55 (1.10-2.19) | 1.27 (0.82-1.98) | 2.29 (1.29-4.05) |
| Diabetes | 2.01 (1.42-2.83) | 2.27 (1.50-3.44) | 1.59 (0.85-2.99) |
| Cardiovascular disease | 2.19 (1.52-3.15) | 2.54 (1.66-3.90) | 1.43 (0.67-3.02) |

Patients with calcification excluded.
All models adjusted for the variables included in the table and centre.
PA: Physical activity (reference = No limitation), BMI: Body mass index
a p for trend are < 0.001 (total and men) and 0.009 (women) for physical activity.
bReference = less than or equal to 7 hours walking per week.
cSex-specific tertiles. Cut points are 97 cm and 105 cm for men and 90 cm and 101 cm for women.

| Table 2: Association between peripheral arterial disease and potential risk factors. Logistic regression models adjusted by age and sex (Continued) |
|-----------------------------------------------|
| **SCORE index (aged<65)** | < 0.001 |
| 2.5-5 | 1.43 (0.71-2.89) |
| >5 | 3.99 (2.00-7.95) |
| Metabolic syndrome (NCEP) | 1.79 (1.36-2.36) |

Patients with calcification excluded.
All models adjusted for age, sex and centre.
BMI: Body mass index.
Sex-specific tertiles. Cut points are 97 cm and 105 cm for men and 90 cm and 101 cm for women.
ex-smokers, those with clinical manifestations of IC or difficulty in performing physical activity, patients with hypertension, diabetes or hypertriglyceridemia. The addition of ankle-arm index in prediction tables of cardiovascular risk can improve the sensitivity and positive predictive values, especially in patients with intermediate or low cardiovascular risk.

Conclusions
In summary, the prevalence of peripheral arterial disease in our population of low-medium risk is lower compared to studies carried out in countries of high risk. Our results coincide with most of the studies published on risk factors for peripheral arterial disease, but the protector effect of physical exercise and overweight is of note. It is necessary to determine the prognostic value of peripheral arterial disease compared to cardiovascular disease in a longitudinal study and to define the population in whom the use of the ankle arm index should be a priority to thereby correctly stratify the cardiovascular risk of our population beyond the use of the tables for risk of cardiovascular disease. Further studies are required to clarify the clinical significance and the association with cardiovascular risk of patients with arterial calcification and those in whom the ankle arm index cannot be determined.

Abbreviations
(CVD): Cardiovascular disease; (PAD): Peripheral arterial disease; (AAl): Ankle-arm index; (IC): Intermittent claudication; (LRM): Logistic regression models; (AIC): Akaike Information Criteria; (BMI): Body mass index.

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MTA, JMVB, MS, RF, PT, MV, MDR, JL and MB participated in the design of the study; MTA, JMVB, MS, RF, PT, CA, JL and AS contributed to the coordination study; GP participated in the statistical calculations. All the authors have read and approved the final manuscript.

Competing interests
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