

## ORIGINAL

# Are risk factors and cardiovascular risk scales controlled in hypertensive patients under treatment?

*¿Están controlados los factores de riesgo y las escalas de riesgo cardiovascular en los pacientes hipertensos en tratamiento?*

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## Summary

**Introduction:** High blood pressure is considered one of the main cardiovascular risk factors; therefore, it is essential to exercise adequate control over it.

**Hypothesis:** The blood pressure control in people undergoing treatment is not very accurate and, in addition, the values of the different scales related to cardiovascular risk are high in this group of treated people.

**Material and methods:** Descriptive cross-sectional study conducted in 34,595 hypertensive patients under treatment for both hypertension and associated comorbidities. We assessed the influence of treatments on the degree of control of arterial hypertension and on the values of different scales related to cardiovascular risk.

**Results:** 49.31% of our total population exhibited normal values of blood pressure normal (59.25% in women and 44.54% in men). Almost all the scales analyzed (overweight and obesity, atherogenic indices, metabolic syndrome, cardiovascular risk scales and fatty liver scales) presented worse values in the group that was receiving treatment for arterial hypertension, for dyslipidemia and for diabetes simultaneously.

**Discussion:** Half of the people who were receiving drug treatment for hypertension show high blood pressure levels, so it is necessary discuss about what strategies are necessary to improve this situation.

**Keywords:** Hypertension, cardiovascular diseases, obesity, dyslipidemia, metabolic syndrome.

## Resumen

**Introducción:** La hipertensión arterial se considera uno de los principales factores de riesgo cardiovascular, por lo que es fundamental ejercer un adecuado control sobre ella.

**Hipótesis:** El control de la presión arterial en personas en tratamiento es poco preciso y, además, los valores de las diferentes escalas relacionadas con el riesgo cardiovascular son elevados en este grupo de personas tratadas.

**Material y métodos:** Estudio descriptivo transversal realizado en 34.595 pacientes hipertensos en tratamiento tanto de la hipertensión como de las comorbilidades asociadas. Se evaluó la influencia de los tratamientos en el grado de control de la hipertensión arterial y en los valores de diferentes escalas relacionadas con el riesgo cardiovascular.

**Resultados:** El 49,31% de nuestra población total presentaba valores normales de presión arterial (59,25% en mujeres y 44,54% en hombres). Casi todas las escalas analizadas (sobrepeso y obesidad, índices aterogénicos, síndrome metabólico, escalas de riesgo cardiovascular y escalas de hígado graso) presentaron peores valores en el grupo que recibía tratamiento para la hipertensión arterial, para la dislipidemia y para la diabetes simultáneamente.

**Discusión:** La mitad de las personas que estaban recibiendo tratamiento farmacológico para la hipertensión arterial presentan niveles elevados de presión arterial, por lo que es necesario discutir sobre qué estrategias son necesarias para mejorar esta situación.

**Palabras clave:** Hipertensión, enfermedades cardiovasculares, obesidad, dislipidemia, síndrome metabólico.

## Introduction

Systolic blood pressure is the force exerted by the blood on the arteries when the heart contracts, while the diastolic reflects the pressure in the arteries when the heart rests. It is a remarkable feature that arterial hypertension is a disease that can be controlled, which decreases the quality and life expectancy since it increases the risk of cardiovascular diseases. The modification of the lifestyle and the treatment with specific medication allows controlling the arterial hypertension.

Among the main risk factors, we find age (over 50 years), male gender (although women in post-menopause have increased risk of hypertension), family history, obesity, sedentary lifestyle, alcohol, tobacco, stress, high salt consumption and some contraceptives (especially, in smoking women), among others.

Hypertension treatment is based on drugs and lifestyle changes and low salt diet. It is necessary to avoid the consumption of processed foods, sausages, cold meat, hard cheeses; select low-sodium foods and waters; and encourage the consumption of fruits and vegetables as a source of potassium (which helps control blood pressure). In case of overweight, a reduction in body weight is indicated. Quitting smoking is also recommended. Regular physical activity might be patterned according to the age of each patient, but the recommendation is 30 minutes of daily moderate physical activity.

Currently, antihypertensive are classified according to their mechanism of action as:

- Diuretics. Drugs that reduce the cardiovascular morbidity and mortality associated with HT<sup>1</sup>. There are three different subgroups of diuretics: thiazides, loop of Henle diuretics and potassium sparing agents.
- Beta-blockers. Highly effective, both in monotherapy and in association, in the treatment of mild-to-moderate HT, in the prevention of cardiovascular complications<sup>2</sup>, in the clear prevention of reinfarction in patients with ischemic heart disease<sup>3</sup> and in increasing survival in patients with heart failure<sup>4</sup>.
- Calcium channel blockers are characterized by a very fast onset of action, which can lead to hypotension in vulnerable individuals, especially the elderly<sup>5</sup>. For this reason, these drugs are currently generally prescribed in extended-release presentations.
- Angiotensin-converting enzyme inhibitors not only reduce blood pressure, but also reduce the vascular damage caused by hypertension (renal failure or heart failure<sup>6</sup>).
- Angiotensin II receptor antagonist. Like ACEI, the decrease in BP with these drugs is not accompanied by reflex tachycardia and, unlike those, they do not cause coughing or angioedema.

## Study purposes

- To determine the prevalence of cardiovascular risk factors in hypertensive patients receiving pharmacological treatment.
- To determine the values of the cardiovascular risk scales in hypertensive patients receiving pharmacological treatment.
- To determine the prevalence of altered values of cardiovascular risk scales in hypertensive patients receiving pharmacological treatment.

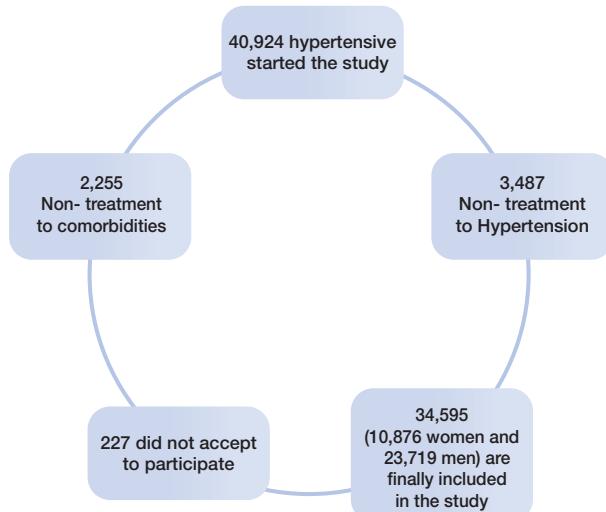
## Hypothesis

Hypertensive individuals, with or without associated comorbidity, who are receiving antihypertensive treatment, in addition to treatment for the associated comorbidity, have low control of cardiovascular risk factors and high values on the cardiovascular risk scales.

## Material and methods

Descriptive, cross-sectional, and observational study of 40,924 hypertensive patients. Of these, 2,255 were excluded because they were not receiving treatment for comorbidity, 3,847 were not taking their antihypertensive treatment and 227 did not agree to participate in the study. The final number of participants who met all the inclusion criteria was 34,595 (10,876 women and 23,719 men) with mean age of 51.61 years (51.38 years in women and 51.72 years in men). (See flow chart in figure 1).

**Figure 1:** Flowchart of participants.



**Inclusion criteria:**

- Being diagnosed hypertensive and on active antihypertensive treatment. In case of associated comorbidity, it must also be under active treatment.
- Age between 18 and 70 years.
- Agree to participate in the study.

**Parameters related to CVD risk included in the assessment:**

Anthropometric and clinical parameters. Anthropometric and clinical measurements and blood sampling to determine the analytical parameters were performed by the health personnel of the occupational health units involved in the study. For the collection of the different anthropometric parameters, international recommendations were followed. All measurements were performed by trained health personnel to minimize interobserver bias.

Weight (in kg) and height (in cm) were determined with a SECA 700 professional measuring scale. Abdominal waist circumference (cm) was calculated with a model 20 SECA Measuring Tape.

**Overweight and obesity scales:**

Body mass index (BMI) was obtained using the Quetelet index: weight in kilograms divided by the square of the height expressed in meters. Based on the BMI, four categories were established according to the classification of the Spanish Society for the Study of Malnutrition and Obesity (SEEDO)<sup>7</sup>: Underweight: BMI<18.5; Normal weight: BMI 18.5-24.9; Overweight: BMI 25-29.9; Obesity: BMI ≥ 30.

The waist / height ratio (ICALT) was considered high from 0.50.

CUN BAE. -44.988 + (0.503 × age) + (10.689 × sex) + (3.172 × BMI) - (0.026 × BMI<sup>2</sup>) + (0.181 × BMI × sex) - (0.02 × BMI × age) --- (0.005 × BMI<sup>2</sup> × sex) + (0.00021 × BMI<sup>2</sup> × age), where men = 0 and women = 1 with respect to sex, measuring age in years<sup>8</sup>.

The following classifications were used to stratify into the different categories of overweight and obesity:

- Male population: < 20% normal weight, 20-25% overweight, > 25% obesity
- Female population: < 30% normal weight, 30-35% overweight, > 35% obesity.

ECORE-BF<sup>9</sup> -97.102 + 0.123 (age) + 11.9 (sex) + 35.959 (LnIMC) where male is equal to 0 and female is equal to 1. The authors propose the same cut-off points as CUN BAE.

Relative fat mass<sup>10</sup>.

Women: 76- (20 × (height/w waist))

Men: 64- (20 × (height/w waist))

Suggested cut-off points are 40% in women and 30% in men.

Palafox formula<sup>11</sup>. Men = ([BMI/BP] \*10) + BMI. Women = ([BMI/BP] \*10) + BMI + 10. The authors propose the same cut-off points as CUN BAE.

Deurenberg formula<sup>12</sup>. 1.2 x (BMI) + 0.23 x (Age in years) - 10.8 x (sex) - 5.4 where female is equal to 0 and male is equal to 1. It is considered obesity from 25.5% in men and 32% in women.

Body roundness index<sup>13</sup>

$$BRI = 364,2 - 365,5 \times \sqrt{1 - \left( \frac{(WC/(2v))^2}{(0,5 \text{ height})^2} \right)}$$

Visceral adiposity index<sup>14</sup>

*Females:*

$$VAI = \left( \frac{WC}{36,58 + (1,89 \times BMI)} \right) \times \left( \frac{TG}{0,81} \right) \times \left( \frac{1,52}{HDL} \right)$$

*Males:*

$$VAI = \left( \frac{WC}{39,68 + (1,88 \times BMI)} \right) \times \left( \frac{TG}{1,03} \right) \times \left( \frac{1,31}{HDL} \right)$$

The waist circumference is expressed in cm and LDL and triglycerides in mmol/L. Obesity cut-off points vary with age.

Normalized weight-adjusted index (NWAI)<sup>15</sup>.

NWAI= [(weight/10) – (10 × height) + 10] weight in kg and height in meters

Blood pressure was determined after a resting period of about 10 minutes in the supine position, using a calibrated OMRON M3 automatic sphygmomanometer. Three consecutive readings at one-minute intervals were taken and the mean value of the three measurements was calculated. The measurements were classified according to the ESH/ESC criteria<sup>16</sup>: Normal BP: SBP < 130 mmHg and/or DBP < 85 mmHg; Pre-HT: SBP 130-139 mmHg and/or DBP 85-89 mmHg; HT 1: SBP 140-159 mmHg and/or DBP 90-99 mmHg; HT 2: SBP > 160 mmHg and/or DBP > 100 mmHg.

Analytical parameters. Blood extraction to determine the analytical parameters included in the study was performed by peripheral venipuncture in the same session and at the same place as the anamnesis and physical examination, after a minimum overnight fast of 12 hours. Values were expressed in milligrams/deciliter (mg/dL). The values to define the presence or absence of diabetes were defined based on the 2013 American Diabetes Association recommendations. Fasting plasma glucose: Normal < 100 mg/dL. Altered basal blood glucose: 100-125 mg/dL. Diabetes > 125 mg/dL. The cut-off values for lipids

have been defined based on those included in the 2010 SEMERGEN-SEA Consensus Document for the approach to dyslipemic patients<sup>17</sup>.

Atherogenic indices. For the different atherogenic indices, the formulas accepted by the scientific community were used. For each index different cut-off points were defined to establish the atherogenic risk categories according to the existing data in the literature<sup>18</sup>.

### **Metabolic Syndrome.**

- NCEP ATP III criteria<sup>19</sup> (Adult Treatment Panel III National Cholesterol Educational Program). At least 3 of the following risk factors are required: Waist circumference >102 cm in men and >88 cm in women. Serum triglycerides ≥ 150 mg/dL or being on specific treatment for this lipid abnormality. Blood pressure ≥ 130/85 mm Hg or being on specific treatment with antihypertensive drugs. cHDL < 40 mg/dL in men and < 50 mg/dL in women or being on specific treatment for this lipid abnormality. Fasting blood glucose > 100 mg/dL or being on specific treatment with antidiabetic drugs.
- International Diabetes Federation (IDF) criteria<sup>20</sup>. Central obesity defined as waist circumference ≥ 94 cm for Caucasian men and ≥ 80 cm and at least two of the 4 factors mentioned above in NCEP-ATPIII are compulsory.
- JIS criteria: presence of at least three of the following factors: abdominal obesity (waist circumference ≥ 94 cm in men and ≥ 80 cm in women); triglycerides ≥ 150 mg/dl or treatment; HDL cholesterol <40 mg/dl in men and <50 mg/dl in women or treatment; blood pressure ≥ 130/85 mmHg or treatment; fasting glucose ≥ 100 mg/dl or treatment<sup>21</sup>.

### **Cardiovascular risk scales**

- REGICOR scale (*Registro Gironí del Cor*): Framingham model calibrated for the Spanish population. The REGICOR tables allow estimation of the risk of suffering a coronary event (angina, symptomatic or silent myocardial infarction and/or death of coronary origin) in the following 10 years. The REGICOR tables are only applicable to subjects aged 35 to 74 years old. Each subject is classified as: < 5%: low risk, 5-9.9% moderate risk, 10 -14.9% high risk and ≥15% very high risk<sup>22</sup>.
- DORICA scale<sup>23</sup>. It also indicates the risk of suffering a coronary event within the following 10 years. It is considered low risk < 5, slight risk 5-9, moderate risk 10 -19, high risk 20 -39 and very high risk ≥ 40.
- SCORE (Systematic Coronary Risk Evaluation) scale presents differential tables according to the country's risk level<sup>24</sup>. It estimates the risk of cardiovascular and cerebrovascular death within 10 years. The version for low-risk countries, recommended for Spain<sup>25</sup>, was used. It is considered low risk < 3%, moderate risk 4-5% and high risk > 5%<sup>26</sup>.
- ERICE scale (Spanish Cardiovascular Risk Equation). Estimates the risk of suffering a fatal or non-fatal

cerebrovascular event over a 10-year period. The tables apply to people between 30 and 80 years old. Low risk <5%: low risk, mild risk 5-9%, moderate risk 10-14%, moderate-high risk 15-19%, high risk 20-39% and very high risk over 30%<sup>27</sup>.

- Framingham categories<sup>28</sup>.
- Framingham vascular age<sup>29</sup>. This tool has its origin in the Framingham cardiovascular risk scale, which can be calculated from the age of 30 years.
- SCORE vascular age<sup>30</sup>. This tool has its origin in the SCORE cardiovascular risk scale. As with the scale from which it derives, it can be calculated in persons between 40 and 65 years.

### **Nonalcoholic fatty liver scales:**

- Fatty liver index<sup>31</sup>

$$\text{FLI} = \left( e^{0.953 \log_e(\text{triglycerides}) + 0.139 \text{BMI} + 0.718 \log_e(\text{ggt}) + 0.053 \text{waist circumference} - 15.745} \right) / \left( 1 + e^{0.953 \log_e(\text{triglycerides}) + 0.139 \text{BMI} + 0.718 \log_e(\text{ggt}) + 0.053 \text{waist circumference} - 15.745} \right) \times 100$$

A FLI with values over 60 is considered high risk.

- Hepatic steatosis index<sup>32</sup>

HSI = 8 x ALT / AST + BMI (+ 2 if diabetes 2, + 2 if female)  
Values over 36 are considered high risk.

- ZJU index<sup>33</sup>

BMI (kg / m<sup>2</sup>) + Blood glucose (mmol / L) + TG (mmol / L) + 3 \* ALT (IU / L) / AST (IU / L) ratio (+ 2 if female). Values over 38 are considered high risk.

- Framingham steatosis index<sup>34</sup>

- 7.981 + 0.011 × age(years) - 0.146 × sex (female = 1, male = 0) + 0.173 × BMI(kg/m<sup>2</sup>) + 0.007 × triglycerides(mg/dl) + 0.593 × hypertension(yes = 1, no = 0) + 0.789 × diabetes(yes = 1, no = 0) + 1.1 × ALT/AST ratio ≥ 1.33(yes = 1, no = 0).

Values over 28 are considered high risk.

- Lipid accumulation product<sup>35</sup>.

- In men: (waist circumference (cm) - 65) × (triglyceride concentration (mMol)).  
- In women: (waist circumference (cm) - 58) × (triglyceride concentration (mMol))

Other parameters related to cardiovascular risk:

- Triglyceride glucose index triglyceride glucose index-IMC, triglyceride glucose index-p waist TyGindex = LN (TG [mg/dl] × blood glucose [mg/dl]/2). TyGindex-IMC = TyGindex × IMC TyGindex-p waist = TyGindex × p waist<sup>36-39</sup>.

- Waist weight index. This indicator is calculated by applying the formula. WWI = waist circumference / √weight

- Cardiometabolic index<sup>40</sup> is calculated by multiplying the waist-height index by the atherogenic triglycerides / HDL-c index.

- Atherogenic dyslipidemia and lipid triad<sup>41</sup>. Atherogenic dyslipidemia is characterized by elevated triglyceride levels ( $> 150$  mg / dl), low HDL ( $<40$  mg / dl in men and  $<50$  mg / dl in women), and normal or mildly elevated LDL. If LDL values are high ( $> 160$  mg / dl) we speak of lipid triad.

### Statistical analysis.

The SPSS 27.0 package was used for the statistical study. For the initial descriptive analysis, once the normal distribution had been verified and following the Kolmogorov-Smirnov method, the mean and standard deviation values were used. In subsequent analyses, when the variable was continuous, the comparison of means was performed using Student's t test if the variable followed a normal distribution, or the nonparametric Mann-Whitney U test if it did not meet the normality criterion. When the variable was qualitative, Pearson's chi-squared test was used to compare proportions, with a confidence level of 95%. ROC curves were used to calculate the cut-off points. P values less than 0.05 were considered statistically significant.

## Results

The anthropometric, clinical, analytical, and sociodemographic characteristics of the 24,595 people who entered the study are presented in **table II**.

The mean values of the different scales that assess overweight and obesity are presented in **table III**. The results are divided in each sex according to treatment. The highest values appear in almost all cases and in both sexes in the group of people being treated for HT, dyslipidemia, and diabetes.

The values of blood pressure and analytical parameters do not show a homogeneous pattern according to the treatment being received in either sex. The complete data are shown in **table IV**.

Almost all the cardiovascular risk scales studied show higher values in the group of people being treated for the three conditions (HT, dyslipidemia, and diabetes), and this is true for both sexes. The complete data are presented in **table V**.

As shown in **table VI**, the fatty liver scales show higher values in the group receiving the three treatments, in both sexes. The same occurs with the atherogenic indices and with the other metabolic indicators.

When the prevalence of high values of the overweight and obesity scales is evaluated, it is found that in both sexes the highest prevalences are found in the group that receives hypertensive treatment in addition to treatment for dyslipidemia and diabetes. The complete data can be found in **table VII**.

**Table II:** Anthropometric, clinical, analytical, and sociodemographic characteristics of the population.

	Men n=10,876 Mean (sd)	Women n=23,719 Mean (sd)	Total n=34,595 Mean (sd)	p
Age (years)	51.38 (8.23)	51.72 (7.78)	51.61 (7.93)	<0.0001
Height (cm)	159.61 (6.58)	172.68 (6.97)	168.57 (9.15)	<0.0001
Weight (kg)	73.92 (15.50)	88.86 (15.93)	84.16 (17.25)	<0.0001
Waist (cm)	79.15 (12.02)	90.52 (11.52)	86.95 (12.81)	<0.0001
SBP (mmHg)	132.35 (18.54)	139.04 (18.10)	136.94 (18.50)	<0.0001
DBP (mmHg)	80.90 (11.19)	85.29 (11.16)	83.91 (11.36)	<0.0001
Cholesterol (mg/dl)	207.47 (36.68)	200.20 (38.36)	202.49 (37.99)	<0.0001
HDL (mg/dl)	53.39 (7.96)	46.07 (8.18)	48.37 (8.79)	<0.0001
LDL (mg/dl)	130.77 (35.73)	124.36 (36.81)	126.41 (36.59)	<0.0001
Triglycerides (mg/dl)	117.50 (63.98)	154.14 (99.67)	142.62 (91.58)	<0.0001
Blood glucose (mg/dl)	97.49 (25.06)	105.73 (32.37)	103.14 (30.50)	<0.0001
GPT (U/L)	24.05 (13.84)	33.95 (19.00)	30.86 (18.15)	<0.0001
GOT (U/L)	20.00 (8.68)	25.64 (11.95)	23.77 (11.29)	<0.0001
GGT (U/L)	27.46 (27.26)	48.04 (53.48)	41.63 (47.86)	<0.0001
	%	%	%	p
20-49 years	36.63	34.73	35.33	<0.0001
50-70 years	63.37	65.27	64.67	
Social Class I-II	19.44	19.63	19.57	<0.0001
Social Class III	80.56	80.37	80.43	
Non-smokers	68.46	67.84	68.04	0.476
Smokers	31.54	32.16	31.96	
HT	53.82	43.12	46.48	<0.0001
Diabetes + HT	18.82	22.26	21.18	<0.0001
Dyslipidemia + HT	16.96	18.86	18.27	<0.0001
Diabetes + dyslipidemia + HT	10.40	15.76	14.07	<0.0001

**Table III:** Mean values of overweight-obesity indicators according to drug consumption by sex.

	Women					Men				
	HT n=5,853 Mean (sd)	HT+Diab n=2,047 Mean (sd)	HT+DLP n=1,845 Mean (sd)	HT+DLP+Diab n=1,131 Mean (sd)	p	HT n=10,228 Mean (sd)	HT+Diab n=5,280 media (sd)	HT+DLP n=4,474 media (sd)	HT+DLP+Diab n=3,737 media (sd)	p
Age	49.47 (8.59)	51.85 (7.70)	54.13 (6.89)	55.91 (5.76)	<0.0001	49.88 (8.28)	52.40 (7.52)	52.75 (7.06)	54.57 (6.13)	<0.0001
BMI	28.81 (5.87)	29.93 (5.91)	28.05 (5.11)	29.85 (5.16)	<0.0001	29.56 (4.80)	30.17 (5.00)	29.19 (4.30)	30.39 (4.63)	<0.0001
Waist/Height	0.49 (0.08)	0.50 (0.07)	0.48 (0.07)	0.50 (0.07)	<0.0001	0.52 (0.06)	0.53 (0.06)	0.52 (0.06)	0.53 (0.06)	<0.0001
CUN BAE	40.85 (6.51)	42.38 (6.29)	40.65 (5.64)	42.88 (5.25)	<0.0001	30.45 (5.84)	31.43 (5.81)	30.35 (5.24)	31.97 (5.33)	<0.0001
ECORE-BF	41.04 (7.13)	42.72 (7.01)	40.78 (6.33)	43.28 (6.01)	<0.0001	30.35 (5.73)	31.39 (5.74)	30.33 (5.24)	31.98 (5.33)	<0.0001
RFM	34.86 (5.90)	35.36 (5.74)	33.83 (5.46)	35.54 (5.11)	<0.0001	25.31 (4.73)	25.43 (4.61)	24.83 (4.56)	25.66 (4.44)	<0.0001
Palafolls	42.43 (6.13)	43.66 (6.18)	41.71 (5.34)	43.60 (5.38)	<0.0001	32.81 (5.00)	33.49 (5.22)	32.48 (4.49)	33.74 (4.83)	<0.0001
IMG	40.56 (7.31)	42.45 (7.26)	40.71 (6.29)	43.28 (6.19)	<0.0001	30.74 (6.02)	32.06 (6.16)	30.96 (5.37)	32.82 (5.60)	<0.0001
NWAI	1.38 (1.52)	1.66 (1.52)	1.20 (1.30)	1.66 (1.31)	<0.0001	1.56 (1.45)	1.74 (1.50)	1.45 (1.28)	1.80 (1.38)	<0.0001
BRI	3.41 (1.52)	3.51 (1.46)	3.12 (1.26)	3.48 (1.28)	<0.0001	3.92 (1.33)	3.94 (1.31)	3.76 (1.19)	3.98 (1.22)	<0.0001
VAI	3.50 (2.06)	3.70 (2.17)	3.94 (2.93)	4.51 (2.62)	<0.0001	9.65 (7.29)	10.27 (8.02)	10.88 (7.87)	12.29 (10.02)	<0.0001

BMI. Body Mass Index. CUN BAE. Body Adiposity Estimator of the Clínica Universitaria de Navarra. ECORE-BF. Cordoba-Body Fat Equation. RFM Relative fat mass. IMG Fat mass index, NWAI Normalized weight adjusted index. BRI Body Roundness Index, VAI Visceral Adiposity Index.

**Table IV:** Mean values of blood pressure and analytical parameters according to drug consumption by sex.

	Women					Men				
	HT n=5,853 Mean (sd)	HT+Diab n=2,047 Mean (sd)	HT+DLP n=1,845 Mean (sd)	HT+DLP+Diab n=1,131 Mean (sd)	p	HT n=10,228 Mean (sd)	HT+Diab n=5,280 media (sd)	HT+DLP n=4,474 media (sd)	HT+DLP+Diab n=3,737 media (sd)	p
TSBP	131.50 (18.41)	134.60 (18.79)	130.91 (18.65)	135.07 (17.99)	<0.0001	140.31 (18.40)	136.42 (17.10)	139.04 (18.10)	140.51 (17.97)	<0.0001
DBP	81.24 (11.39)	81.67 (11.09)	79.31 (10.75)	80.33 (10.72)	<0.0001	83.80 (10.66)	85.29 (11.16)	83.80 (10.66)	85.76 (11.07)	<0.0001
Cholesterol	208.03 (35.53)	207.42 (35.09)	209.75 (40.10)	200.91 (38.84)	<0.0001	204.04 (36.21)	198.72 (35.50)	199.89 (41.27)	192.17 (42.72)	<0.0001
HDL	54.09 (8.49)	53.02 (7.67)	52.93 (7.21)	51.13 (6.07)	<0.0001	47.13 (8.42)	45.41 (7.98)	45.86 (7.85)	44.37 (7.73)	<0.0001
LDL	131.89 (34.41)	130.88 (33.90)	132.31 (38.99)	122.33 (38.89)	<0.0001	128.79 (34.67)	124.35 (34.85)	122.33 (39.09)	114.44 (4.21)	<0.0001
Triglycerides (mg/dl)	110.93 (55.79)	117.66 (60.69)	125.14 (81.65)	138.76 (70.41)	<0.0001	144.64 (87.45)	149.30 (99.43)	164.74 (103.09)	174.28 (120.90)	<0.0001
Blood glucose (mg/dl)	92.37 (16.12)	104.15 (33.26)	95.33 (17.77)	115.45 (40.42)	<0.0001	97.32 (19.92)	113.63 (40.15)	99.63 (19.93)	124.85 (46.06)	<0.0001
GPT	23.11 (12.73)	23.48 (14.97)	25.34 (13.21)	27.21 (16.42)	<0.0001	32.82 (18.53)	33.15 (18.48)	35.77 (19.27)	35.57 (20.20)	<0.0001
GOT	20.01 (8.49)	19.76 (10.32)	20.09 (6.88)	20.40 (7.94)	<0.0001	24.77 (9.29)	25.20 (11.59)	27.23 (15.31)	26.57 (13.27)	<0.0001
GGT	25.95 (24.34)	26.91 (23.99)	28.16 (22.80)	34.05 (44.67)	<0.0001	45.60 (52.84)	45.27 (49.41)	51.44 (51.51)	53.69 (61.34)	<0.0001

**Table V:** Mean values of cardiovascular risk scales according to drug use by sex.

	Women					Men				
	HT n=5,853 Mean (sd)	HT+Diab n=2,047 Mean (sd)	HT+DLP n=1,845 Mean (sd)	HT+DLP+Diab n=1,131 Mean (sd)	p	HT n=10,228 Mean (sd)	HT+Diab n=5,280 media (sd)	HT+DLP n=4,474 media (sd)	HT+DLP+Diab n=3,737 media (sd)	p
ALLY EV SCORE	6.69 (5.38)	7.66 (5.60)	7.56 (5.22)	8.42 (5.10)	<0.0001	10.47 (7.26)	11.29 (7.27)	10.39 (7.20)	11.38 (7.16)	<0.0001
SCORE	0.85 (1.28)	1.21 (1.54)	1.32 (1.55)	1.67 (1.60)	<0.0001	2.65 (2.77)	3.18 (3.06)	2.94 (2.83)	3.37 (3.00)	<0.0001
ALLY EV FRAMINGHAM	14.99 (12.92)	29.46 (10.21)	14.89 (12.56)	28.85 (9.46)	<0.0001	16.03 (9.97)	26.53 (8.85)	15.97 (10.17)	26.62 (8.63)	<0.0001
REGICOR	3.30 (2.19)	3.93 (2.19)	3.41 (2.18)	3.81 (2.72)	<0.0001	3.75 (2.44)	4.08 (3.00)	3.65 (2.42)	4.12 (3.06)	<0.0001
ERICE	5.36 (4.62)	9.17 (7.29)	7.65 (4.52)	12.50 (6.67)	<0.0001	10.31 (6.69)	15.73 (9.14)	12.14 (6.94)	17.60 (8.62)	<0.0001
DORICA	4.98 (3.48)	6.51 (4.28)	6.27 (3.76)	8.14 (4.64)	<0.0001	9.14 (5.61)	10.71 (6.42)	9.51 (5.53)	11.46 (7.12)	<0.0001
CVD RISK	9.76 (6.83)	12.11 (8.11)	11.19 (6.94)	14.47 (8.63)	<0.0001	19.22 (10.16)	22.62 (10.54)	20.95 (10.14)	24.93 (10.01)	<0.0001
Framingham categories	6.00 (4.24)	7.80 (5.36)	7.46 (4.45)	10.03 (5.97)	<0.0001	11.61 (7.32)	13.95 (8.75)	12.20 (7.43)	15.05 (9.33)	<0.0001
Framingham categories duro	3.07 (2.97)	4.31 (4.13)	3.94 (3.35)	5.92 (4.92)	<0.0001	8.87 (6.31)	10.87 (7.83)	9.35 (6.38)	11.84 (8.59)	<0.0001

ALLY EV. Lost years of vascular age. CVD RISK. Cardiovascular

**Table VI:** Mean values of fatty liver scales and atherogenic indices according to drug consumption by sex.

	Women					Men				
	HT n=5,853 Mean (sd)	HT+Diab n=2,047 Mean (sd)	HT+DLP n=1,845 Mean (sd)	HT+DLP+Diab n=1,131 Mean (sd)	p	HT n=10,228 Mean (sd)	HT+Diab n=5,280 media (sd)	HT+DLP n=4,474 media (sd)	HT+DLP+Diab n=3,737 media (sd)	p
Fatty liver index	32.74 (27.64)	37.55 (28.24)	30.99 (25.29)	41.28 (27.85)	<0.0001	54.60 (27.46)	56.43 (26.93)	56.11 (26.09)	61.58 (26.47)	<0.0001
Hepatic steatosis index	40.22 (7.38)	42.67 (7.32)	39.51 (6.59)	43.42 (6.58)	<0.0001	39.54 (6.90)	41.72 (6.90)	39.20 (6.24)	42.28 (6.32)	<0.0001
ZJU index	40.95 (6.65)	42.22 (7.30)	40.28 (5.56)	43.79 (6.66)	<0.0001	40.14 (5.59)	41.16 (6.31)	40.18 (5.46)	42.72 (6.46)	<0.0001
Fatty liver disease index	33.90 (6.44)	34.69 (6.69)	33.12 (5.32)	35.98 (6.14)	<0.0001	34.89 (5.40)	35.49 (5.80)	34.80 (5.10)	36.44 (5.48)	<0.0001
Lipid accumulation product	28.45 (24.86)	30.29 (24.60)	27.66 (25.93)	35.59 (27.66)	<0.0001	44.67 (38.91)	45.27 (40.14)	45.98 (38.59)	53.12 (48.19)	<0.0001
IA cholesterol/HDL	3.94 (0.91)	3.98 (0.85)	4.04 (0.96)	3.98 (0.90)	<0.0001	4.47 (1.15)	4.52 (1.17)	4.48 (1.21)	4.47 (1.31)	<0.0001
IA Triglycerids/HDL	2.13 (1.21)	2.30 (1.33)	2.45 (1.83)	2.80 (1.62)	<0.0001	3.24 (2.26)	3.47 (2.54)	3.77 (2.61)	4.15 (3.19)	<0.0001
IA LDL/HDL	2.51 (0.82)	2.52 (0.76)	2.55 (0.86)	2.43 (0.83)	<0.0001	2.84 (0.98)	2.84 (1.00)	2.75 (1.03)	2.67 (1.11)	<0.0001
IA HDL/LDL+VLDL	0.38 (0.14)	0.37 (0.12)	0.37 (0.12)	0.37 (0.12)	<0.0001	0.33 (0.12)	0.32 (0.13)	0.33 (0.13)	0.34 (0.14)	<0.0001
IA cholesterol-HDL	153.94 (36.64)	154.40 (35.84)	156.83 (40.81)	149.77 (39.56)	<0.0001	156.91 (37.38)	153.32 (36.97)	154.03 (42.05)	147.80 (43.89)	<0.0001
Cardiometabolic index	1.08 (0.69)	1.17 (0.74)	1.20 (0.91)	1.43 (0.89)	<0.0001	1.74 (1.31)	1.86 (1.45)	1.98 (1.42)	2.23 (1.81)	<0.0001
Triglyceride glucose index	8.43 (0.49)	8.59 (0.56)	8.57 (0.50)	8.84 (0.61)	<0.0001	8.71 (0.55)	8.86 (0.64)	8.83 (0.60)	9.09 (0.67)	<0.0001
Waist triglyceride index	101.15 (56.99)	107.40 (59.71)	109.68 (74.25)	126.53 (69.41)	<0.0001	150.91 (98.50)	154.93 (108.39)	166.98 (108.63)	181.13 (131.67)	<0.0001
Waist weight index	9.28 (0.78)	9.20 (0.67)	9.13 (0.70)	9.21 (0.65)	<0.0001	9.67 (0.74)	9.59 (0.68)	9.59 (0.70)	9.60 (0.68)	<0.0001

**Table VII:** Prevalence of overweight-obesity with different scales according to drug consumption by sex.

	Women					Men				
	HT n=5,853 Mean (sd) %	HT+Diab n=2,047 Mean (sd) %	HT+DLP n=1,845 Mean (sd) %	HT+DLP+Diab n=1,131 Mean (sd) %	p	HT n=10,228 Mean (sd) %	HT+Diab n=5,280 media (sd) %	HT+DLP n=4,474 media (sd) %	HT+DLP+Diab n=3,737 media (sd) %	p
Waist/Height >0,50	40.80	45.43	34.09	47.30	<0.0001	61.61	62.71	59.50	64.97	<0.0001
BMI obesity	37.50	44.70	29.76	42.79	<0.0001	41.13	45.72	38.44	49.69	<0.0001
BMI overweight	34.00	34.49	40.81	41.82		43.56	41.99	46.69	40.49	
CUN BAE obesity	81.12	87.40	85.31	94.08	<0.0001	83.54	87.56	85.49	91.73	<0.0001
CUN BAE overweight	14.37	10.26	11.82	5.31		13.69	10.81	12.43	7.55	
ECORE-BF obesity	79.51	85.69	83.09	93.37	<0.0001	83.52	87.56	85.38	91.30	<0.0001
ECORE-BF overweight	15.27	11.33	12.95	5.13		13.54	10.70	12.41	7.76	
RFM obesity	55.37	59.94	49.92	62.69	<0.0001	71.04	71.82	69.18	73.67	<0.0001

BMI. Body Mass Index. CUN BAE. Clínica Universitaria de Navarra Body Adiposity Estimator. ECORE-BF. Equation Córdoba-Body Fat. RFM Relative Fat Mass.

**Table VIII:** Prevalence of altered blood pressure values and analytical parameters according to drug use by sex.

	Women					Men				
	HT n=5,853 Mean (sd) %	HT+Diab n=2,047 Mean (sd) %	HT+DLP n=1,845 Mean (sd) %	HT+DLP+Diab n=1,131 Mean (sd) %	p	HT n=10,228 Mean (sd) %	HT+Diab n=5,280 media (sd) %	HT+DLP n=4,474 media (sd) %	HT+DLP+Diab n=3,737 media (sd) %	p
HT	38.97	44.41	35.34	44.30	<0.0001	56.20	58.24	48.39	59.03	<0.0001
Cholesterol ≥ 200	57.66	57.16	55.72	44.47	<0.0001	53.09	47.27	46.58	37.97	<0.0001
LDL > 130	50.86	49.73	48.08	36.25	<0.0001	47.93	43.81	40.93	32.99	<0.0001
Triglycerides > 150	17.17	20.66	23.96	32.54	<0.0001	34.74	36.63	44.23	46.96	<0.0001
Blood glucosa 100-125	19.78	26.18	27.05	29.44	<0.0001	29.26	32.23	35.83	32.70	<0.0001
Blood glucosa > 125	2.00	12.85	3.36	24.49	<0.0001	4.61	21.33	5.30	32.00	<0.0001

**Table IX:** Percentage of patients with controlled blood pressure figures according to treatments received.

	Women n=10,876 % HT controlled	Men n=23,719 % HT controlled	Total n=34,595 % HT controlled
HT + Dislipidemia + diabetes	55.70	40.97	44.39
HT	61.03	43.80	50.07
HT + diabetes	55.59	41.76	45.63
HT + dislipidemia	64.66	51.61	55.42
Mean	59.25	44.54	49.31

**Table VIII** shows the prevalence of HT and analytical abnormalities in treated patients. **Table IX** shows that women overall have controlled blood pressure in 59.25% of cases, while in men the figure drops to 44.54%.

**Table X** shows that most of the scales for cardiovascular risk, metabolic syndrome and atherogenic dyslipidemia have higher prevalence of altered values in the group receiving antihypertensive treatment, as opposed to dyslipidemia and diabetes, which is the same in both sexes.

## Discussion

No prior study has been found in the literature that assesses the degree of blood pressure control in hypertensive patients with comorbidities (dyslipidemia and diabetes), nor any previous work that focuses on the control of cardiovascular risk parameters or scales in hypertensive patients under treatment; therefore, we cannot compare our results with those of other preceding authors.

For the discussion we will focus on the degree of control of blood pressure in hypertensive patients under treatment.

Remarkably, the overall degree of control of blood pressure in our study was 49.31% (59.25% in women and 44.54% in men).

In a study of 124 patients with an average age of 68 years in ten community pharmacies in different areas of the Valencian Community (Spain), 46.80% of these patients were not under control. Almost half of the patients treated with antihypertensive drugs presented out-of-range blood pressure values according to the 2018 European Guideline for the management of hypertension<sup>42</sup>. Another study in 265 hypertensive individuals on treatment showed control of blood pressure figures in 33% of males versus 49% of females. These studies rank an intermediate-low position compared to other previous studies on Spanish and European populations<sup>43</sup>. Despite this, a significant margin for improvement remains, as shown by the more favorable results presented by studies in Denmark<sup>44</sup>, Canada and the United States and other European countries<sup>45</sup> with a control percentage of more than 50% in treated subjects.

An important Spanish study, Di@bet.es, showed that only 26.6% have controlled blood pressure, higher in women (24.90%) than in men (16%)<sup>46</sup>. Di@bet.es is a national study designed to estimate the prevalence of diabetes mellitus and other cardiovascular risk factors in

the Spanish adult population. It presents the prevalence of arterial hypertension and the degree to which it is recognized, treated, and controlled. A sample of the Spanish population with 5,048 adults aged  $\geq 18$  years was included. A clinical interrogation and examination including three resting and sitting blood pressure readings were performed to calculate the mean. Hypertension is defined as systolic blood pressure  $\geq 140$  mmHg and/or diastolic blood pressure  $\geq 90$  mmHg

and/or on antihypertensive drug treatment. The data are lower than those obtained in our study, perhaps because all the people in our study received treatment. The factors associated with poorer control were male sex, overweight or obesity, and an associated diagnosis of diabetes mellitus.

Prior to this study, the data on HT control in our country were more encouraging, and even previous registries

**Table X:** Prevalence of altered values of CVR scales and metabolic scales with different scales according to drug consumption by sex.

	Women					Men				
	HT n=5,853 Mean (sd) %	HT+Diab n=2,047 Mean (sd) %	HT+DLP n=1,845 Mean (sd) %	HT+DLP+Diab n=1,131 Mean (sd) %	p	HT n=10,228 Mean (sd) %	HT+Diab n=5,280 media (sd) %	HT+DLP n=4,474 media (sd) %	HT+DLP+Diab n=3,737 media (sd) %	p
Moderate SCORE	7.56	11.01	11.88	16.09	<0.0001	23.33	25.95	25.57	28.91	<0.0001
High SCORE	2.84	4.74	5.04	5.15		20.43	26.62	24.30	29.13	
Moderate ALLY EV SCORE	14.31	14.38	15.22	18.54	<0.0001	11.79	11.40	12.11	11.67	<0.0001
High ALLY EV SCORE	51.05	40.97	39.05	36.53		55.87	60.88	55.83	62.04	
Moderate REGICOR	20.95	26.00	23.95	25.96	<0.0001	26.67	28.65	26.07	27.90	<0.0001
High-very high REGICOR	1.74	5.31	1.64	3.64		2.94	5.12	2.49	5.23	
Moderate ALLY EV Framingham	18.82	8.25	17.40	5.66	<0.0001	29.18	11.87	26.82	10.34	<0.0001
High ALLY EV Framingham	45.41	86.32	47.30	89.38		42.51	83.94	43.89	85.56	
Moderate-moderately high ERICE	11.63	26.62	23.46	41.15	<0.0001	45.51	48.10	54.88	55.18	<0.0001
High-very high ERICE	0.19	5.93	0.00	12.39		11.38	20.50	16.51	25.05	
Moderate DORICA	12.23	20.89	19.51	31.72	<0.0001	31.73	39.46	34.52	40.69	<0.0001
High-very high DORICA	0.30	1.25	0.67	2.88		6.05	9.16	6.76	11.90	
Sd MTB ATP III	30.33	55.54	50.51	100.00	<0.0001	33.15	56.16	68.10	100.00	<0.0001
Sd MTB ATP IDF	30.31	46.80	35.34	47.39	<0.0001	29.34	38.71	32.32	39.84	<0.0001
Sd MTB ATP JIS	35.01	60.33	58.43	99.12	<0.0001	54.28	82.12	72.42	99.36	<0.0001
Sd MTB ATP SZABO	47.02	94.87	91.33	99.91	<0.0001	56.03	97.14	92.38	99.89	<0.0001
Atherogenic dyslipidemia	10.25	13.53	14.58	22.99	<0.0001	14.73	18.64	18.26	25.26	<0.0001
Lipid Triad	2.82	2.88	3.69	3.98	<0.0001	4.01	3.88	4.29	3.98	<0.0001
Hypertriglyceridemic waistline	4.89	4.79	4.28	8.66	<0.0001	15.16	16.34	17.21	22.10	<0.0001

ALLY EV. Lost years of vascular age.

**Table XI:** Prevalence of altered values of fatty liver scales and metabolic indices according to drug consumption by sex.

	Women					Men				
	HT n=5,853 Mean (sd) %	HT+Diab n=2,047 Mean (sd) %	HT+DLP n=1,845 Mean (sd) %	HT+DLP+Diab n=1,131 Mean (sd) %	p	HT n=10,228 Mean (sd) %	HT+Diab n=5,280 media (sd) %	HT+DLP n=4,474 media (sd) %	HT+DLP+Diab n=3,737 media (sd) %	p
High Fatty liver index	19.83	24.97	16.07	27.94	<0.0001	45.48	47.49	47.12	55.88	<0.0001
High Hepatic steatosis index	69.25	82.10	71.09	90.08	<0.0001	68.60	81.23	68.51	85.06	<0.0001
High ZJU index	61.32	68.67	66.41	77.10	<0.0001	62.13	66.04	63.20	75.50	<0.0001
High Fatty liver disease index	53.12	50.26	64.58	57.63	<0.0001	60.57	59.19	67.40	52.76	<0.0001
Moderate IA cholesterol/HDL	23.71	24.82	26.94	25.73	<0.0001	28.47	30.91	29.28	25.73	<0.0001
High IA cholesterol/HDL	0.41	0.39	0.65	0.35		0.15	0.28	0.34	0.35	
High IA triglyceridos/HDL	16.88	19.83	28.30	32.98	<0.0001	40.99	44.77	52.06	56.81	<0.0001
High IA LDL/HDL	25.44	25.11	27.21	22.46	<0.0001	40.02	41.72	38.69	52.59	<0.0001
High IA cholesterol-HDL	74.83	75.57	73.66	69.76	<0.0001	76.96	74.28	71.28	64.52	<0.0001

ALLY EV. Lost years of vascular age.

carried out in series, with similar methodologies, such as the PRESCAP<sup>47</sup> 2002, 2006 and 2010 or CARDIOTENS<sup>48</sup> 1999 and 2009 registries, had shown a temporary trend towards a continuous improvement in the degree of control. The PRESCAP studies showed how the percentage of controlled hypertensive patients progressively improved in recent years, from 36% in 2002 to 41.50% in 2006 and 47% in 2010. Similarly, the CARDIOTENS registry showed how the degree of control increased from 40% in 1999 to 55% in 2009, representing a relative increase of 38.50%. In the CARDIOTENS study, lack of blood pressure control was associated with lifestyle and diet-related factors, specifically obesity and smoking, both of which were associated with worse control. The data from our study are more in agreement with those obtained in the Cardiotens study. In a Peruvian study, 53.80% of hypertensive patients showed blood pressure figures within normal values<sup>49</sup>.

The fact that the 'poor control' factor is based on a single measurement of blood pressure may have resulted in an overestimation of prevalence and an underestimation of control in some studies<sup>50</sup>.

The degree of control in people already being treated is difficult to improve. Despite the existence of treatments that have proven to be very effective and efficient, therapeutic objectives are not met in the 50% of the cases. On the one hand, there is the possibility of influencing on the so-called "therapeutic inertia", and with the change or prescription of combined therapy as soon as necessary, proving that the patient tolerate it. On the

other hand, through health education it is necessary to influence greater therapeutic adherence, probably an important pillar for improving the data obtained in this study<sup>51-53</sup>.

An Ecuadorian study<sup>54</sup> showed that the prevalence of obesity determined by BMI was 27.70% among hypertensive patients, which coincides with a study carried out in Costa Rica<sup>55</sup> showing a prevalence of 30%. In our study the prevalence of obesity was much higher than that obtained in these studies.

### **Limitations of the study**

The study was carried out in the working and non-working population, aged between 18 and 70 years, and in specific geographical areas, so it cannot be extrapolated to the general population and to the entire national territory.

### **Strengths of the study**

The sample size is very large, one of the largest carried out to date. In addition, it should be noted that the population has been segregated according to possible comorbidities and, in addition, not only blood pressure but also many scales related to cardiovascular risk have been taken into account to assess the usefulness of pharmacological treatments.

### **Conflict of interests**

The authors have no conflict of interest.

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