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# Prevalence of hypertension and possible risk factors of hypertension unawareness among individuals aged 30-75 years from two Panamanian provinces: Results from population-based cross-sectional studies, 2010 and 2019 

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#### Abstract

\section*{Background}

Recent estimates of hypertension in Panama remain unknown. We aim to describe the variation in prevalence and unawareness of hypertension in two Panamanian provinces using two different cross-sectional population-based studies and to investigate risk factors associated with hypertension unawareness.

\section*{Methods}

Data were derived from a sub-national study conducted in the provinces of Panama and Colon (PREFREC-2010 [2,733 participants]) and from a nationally representative study (ENSPA-2019), in which we restricted our analyses to the same provinces (4,653 participants). Individuals aged 30-75 years who had (a) self-reported history of hypertension or (b) blood pressure (BP) $\geq 140 / 90 \mathrm{mmHg}$ or (c) a combination or both were classified as hypertensive. Participants with $B P \geq 140 / 90 \mathrm{mmHg}$ who denied a history of hypertension were considered unaware of the condition. Multivariable logistic regression models were used to estimate the association between risk factors and unawareness, expressed as odds ratios (OR) and $95 \%$ confidence interval (CI).


## Findings

In 2010, the prevalence and unawareness of hypertension in men were $51.6 \%(95 \% \mathrm{Cl}$ : $45.7-57.5$ ) and $32.3 \%$ (25.4-40.1), respectively, and in women $46.0 \%$ (42.1-49.9) and $16.1 \%$ (12.6-20.4), respectively. In 2019, the prevalence and unawareness of hypertension in men were $46.5 \%$ (42.1-51.0) and $52.3 \%$ (45.9-58.6), and in women $42.1 \%$ (39.6-44.7)

Data Availability Statement: The datasets used and analyzed in this study contain potentially identifying information and sensitive information that is protected via Law 89/2019 as well as a mandate from the Ethics Committee. They are available on request via email to Planning directorate of the Ministry of Health at cnino@gorgas.gob.pa.

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and 33.3\% (29.8-37.0). Men (2010 and 2019), age <50 years (2010 and 2019), having no/ primary education (2010), and living in a non-urban region (2019) were positively associated with hypertension unawareness, whereas obesity (2010), physical inactivity (2010), family history of hypertension (2019), and BP assessment in the year before study enrollment (2010 and 2019) were inversely associated with hypertension unawareness.

## Interpretation

Benefits of a decrease in the prevalence of hypertension are being undermined by an increase in hypertension unawareness. Actions should be encouraged to strengthen the implementation of the existing healthcare program for cardiovascular risk factor control.

## Introduction

Cardiovascular disease (CVD) is a major global health concern, causing more deaths than all other causes combined [1]. CVD was responsible for $77.0 \%$ of deaths in the Americas in 2000, a percentage that increased to $81.0 \%$ in 2016 [2]. Due to this burden of disease, strategies targeting social, economic, and political determinants, as well as control of major cardiovascular risk factors, need to be implemented.

Hypertension is one of the most important modifiable risk factors for premature CVD, with most of the increase occurring in low- and middle-income countries (LMICs) [3, 4]. Variation in hypertension risk factors, such as obesity, high-sodium diet, excessive alcohol consumption, and physical inactivity, may explain some of the CVD heterogeneity between countries [3]. In addition, most people with hypertension do not have symptoms, making screening for its detection necessary [5]. Studies from 2000-2010 have shown a substantial increase in hypertension awareness in high-income countries (HIC), whereas in LMICs the increase has been slight [6]. However, since 2010, a plateau and even a decrease in hypertension awareness has been identified in HIC [7-9], as well as in LMICs [10, 11].

Panama, an upper-middle-income country [12] and the second fastest-growing country in the Americas region in terms of gross domestic product (GDP) per capita [13], faces great disparities in the distribution of wealth (Gini Index of 49.8) [14]. These disparities are also observed in health conditions among ethnic groups, as well as in different geographic areas [15-17]. The country has experienced a demographic and epidemiological transition associated with a double burden of disease (non-communicable diseases (NCDs) and infectious diseases) [18]. However, NCD-related risk factors such as hypertension predominates with a prevalence among adults of $42.3 \%$ (according to the National Health Survey of Panama (ENSPA) in 2019) [19]. Furthermore, stroke and ischemic heart disease (IHD) were the leading causes of mortality in 2019 ( 41.9 per 100,000 population and 41.8 per 100,000 population, respectively) [20].

Starting in 2014, a healthcare program was implemented, which focused on a multipronged approach to NCDs control, including promotion, prevention, and early detection of hypertension and other cardiovascular risk factors [21]. However, there are no recent data on the country's prevalence of hypertension to estimate the effectiveness of this healthcare program.

We aim (1) to describe the variation in the prevalence and unawareness of hypertension in two Panamanian provinces in two different population-based cross-sectional studies, conducted in 2010 and in 2019, and (2) to investigate the possible risk factors associated with hypertension unawareness.

## Materials and methods

## Study population

The PREFREC (2010) study. The PREFREC study (Spanish language for "Prevalence of Cardiovascular Risk Factors associated with CVD"), $\mathrm{n}=3,590$, is a sub-national, cross-sectional, descriptive study conducted between October 2010 and January 2011, designed to estimate the prevalence of well-known risk factors associated with CVDs in the provinces of Panama and Colon, where $57.4 \%$ of the total country population resided when the survey was implemented. The study included citizens older than 18 years who lived permanently in private housing of urban, rural, and indigenous areas. A complex sampling technique (threestages, stratified, and randomized) was used. Further details regarding PREFREC have been described earlier [22]. In the present study, we included participants aged between 30 and 75 years ( $\mathrm{n}=2,733$ ), out of them $67.5 \%$ were women ( Fig 1 ).

The ENSPA (2019) study. The ENSPA study (Spanish language for "National Health Survey of Panama"), $\mathrm{n}=28,483$, is a cross-sectional, nationwide population-based study conducted between June and December 2019, designed to investigate the population's general health status and disease conditions. The study included individuals of all ages who resided permanently in private housing in rural, urban, and indigenous areas ( $0-14$ years: $\mathrm{n}=10,486$; $\geq 15$ years: $\mathrm{n}=17997$ ). The sample design was a three-stage, stratified, and by conglomerates. The representativeness of the results is at the district level (second-level administrative division) in the entire country, except in the Panama and San Miguelito districts (province of Panama) where it has representativeness of corregimiento (third-level administrative division). Further details regarding ENSPA are described on its website in the Spanish language [23]. For the present study, we included participants living in the provinces of Panama and Colon whose age was between 30 and 75 years, $(\mathrm{n}=4,653)(\mathrm{Fig} 1)$.

Both studies were conducted by the Gorgas Memorial Institute of Health Research, the Ministry of Health of Panama (MoH), and the INEC (Spanish language for "National Institute of Statistics and Census"). The sampling design was calculated using population projections


Fig 1. Flow chart of selected participants. SBP = systolic blood pressure. DBP = diastolic blood pressure. *Denominator for the prevalence of hypertension. ${ }^{* *}$ Numerator for the prevalence of hypertension and the denominator for the prevalence of hypertension unawareness.
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from the two latest National Censuses (2000 for the PREFREC (2010) study and 2010 for the ENSPA (2019) study) for both studies when the data collection took place [24].

## Data collection

Participants answered a questionnaire in Spanish through in-person interviews, which collected information on demographics, socioeconomic, medical and family history, lifestyle, and anthropometric measurements (including blood pressure). Weight, height, and blood pressure (BP) were undertaken using standardized instruments.

## Outcome assessment

Outcome variables were assessed using self-reported medical history of hypertension and BP measurements.

Participants' self-reported medical history was based on the question (presented in both surveys): have you ever been told by a physician that you have hypertension, also called high blood pressure? (The fragment, also called high blood pressure was solely present in the PREFREC study).

Participants' BP measurements were considered to be the average of the second and third BP measurement. In the PREFREC (2010) study, BP measurements have been previously described [25]. In brief, systolic blood pressure (SBP) and diastolic blood pressure (DBP) were measured three times in a sitting position after a minimum of five minutes of rest and recorded with a five-minute interval between each other, using an electronic device (American Diagnostic Corporation model 6013). Measurements were performed in the right arm. In the ENSPA (2019) study, following the WHO Stepwise Approach to Surveillance (STEPS) [26], SBP and DBP were measured three times in a sitting position after fifteen minutes of rest and recorded with a three-minute interval between each other, using an electronic device (OMRON model HEM-7120). Measurements were performed in the left arm.

Hypertension was defined as (a) having a self-reported medical history of hypertension or (b) having a mean SBP $\geq 140$ and/or DBP $\geq 90 \mathrm{mmHg}$ (regardless of their self-reported medical history of hypertension) or (c) a combination of both [27]. This sample constituted our study population for estimating the prevalence of hypertension unawareness. Unawareness of hypertension was defined as having a mean SBP $\geq 140$ and/or DBP $\geq 90 \mathrm{mmHg}$ and no selfreported medical history of hypertension. We excluded participants who denied self-reported medical history of hypertension and were also missing the second and/or third BP measure$\operatorname{ment}(\mathrm{s})(\mathrm{n}=120$ in the PREFREC study; $\mathrm{n}=487$ in the ENSPA study) (Fig 1).

## Exposure variables

Demographic and socioeconomic characteristics included age (years), sex (men/women), ethnicity (Afro-Panamanian, Caucasian, Mestizo, Indigenous, or others), region (urban, nonurban), education (no/primary education, secondary education, and higher education), and monthly family income (MFI) ( $<250$ Panamanian balboas (PAB), 250-999 PAB, and $\geq 1,000$ PAB).

A wide range of established risk factors for hypertension was evaluated. Smoking tobacco consumption was classified into current smokers, ex-smoker, or non-smokers. Body mass index (BMI) was obtained by dividing the participant's weight measurement in kilograms by the square of their height in meters squared and categorized into underweight ( $<18.5 \mathrm{~kg} / \mathrm{m}^{2}$ ), normal weight ( 18.5 to $24.9 \mathrm{~kg} / \mathrm{m}^{2}$ ), overweight ( 25 to $29.9 \mathrm{~kg} / \mathrm{m}^{2}$ ), and obesity ( $\geq 30 \mathrm{~kg} / \mathrm{m}^{2}$ ) [28]. Physical inactivity was assessed in the PREFREC (2010) study when they reported doing less than 150 minutes of certain activities per week. In the ENSPA (2019) study, physical
inactivity among study participants was defined if the metabolic equivalents per minute (METs-minute) per week calculated using the GPAQ (General Practice Assessment of Quality) were less than 600 [29]. Family history of hypertension was recorded when first- and/or sec-ond-degree relatives were reported to have the condition. Self-reported medical history of diabetes was defined if the participant answered positively to the question, have you been told by a physician that you have diabetes?

Healthcare-related factors like BP assessment in the year before study enrollment (assessed in both studies) and health check-up in the year before study enrollment (assessed in the ENSPA (2019) study) were evaluated. The S1 Table shows the definitions of the exposure variables used in the present study.

## Statistical analysis

Continuous variables are presented as median and interquartile range (IQR). Categorical variables are presented as percentages with the corresponding $95 \%$ confidence intervals (CIs). Both categorical and continuous variables assessed the total population of the provinces of Panama and Colon when the respective study took place using the expansion weights calculated and provided by the INEC. Results are then presented weighted (N). Missing data were excluded from the analysis.

To examine the association of each exposure variable with hypertension unawareness, its respective odds ratios (OR) with their $95 \%$ CIs were calculated by applying unconditional logistic regression models. Crude and multivariable logistic regression models were performed to address potential confounding bias. The first adjusted model (model A) accounted for the demographic factors (sex, age, and ethnicity), established risk factors for hypertension (tobacco consumption, BMI categories, physical inactivity, family history of hypertension, and self-reported medical history of diabetes), and BP assessment in the year before study enrollment. Then, in a second model (model B) the socioeconomic determinants (region, MFI, and education) were added to model A. In addition, we included in the (S2 and S3 Tables) crude and multivariable logistic regression models using merged data from both studies (both PREFREC and ENSPA) in which the prevalence of hypertension (S2 Table) and the prevalence of hypertension unawareness (S3 Table) were the outcome variables, respectively, with the aforementioned predictor variables together with the study period as an additional predictor (being PREFREC the reference level).

Prevalence tables are presented stratified by sex. Data were analyzed using SPSS V.20.0 software and R program version 4.0 .0 with the survey package version 4.0.

## Ethical statement

Studies were approved by the National Bioethics Committee of the Gorgas Memorial Institute of Health Studies and conducted following the Declaration of Helsinki. All participants signed an informed consent to be enrolled in the study.

## Results

## Demographic and socioeconomic characteristics

Overall, the predominant ethnicity was mestizo (men: 60.4\% in PREFREC (2010) and 47.5\% in ENSPA (2019), women: 60.7\% in PREFREC (2010) and 49.4\% in ENSPA (2019)), and most individuals lived in an urban region (Table 1). According to the education levels, fewer participants reported having no education/primary education in ENSPA (2019), compared to PREFREC (2010) ( $26.8 \%$ vs $20.3 \%$ for men; $32.1 \%$ vs $23.0 \%$ for women). In contrast, more

Table 1. Distribution of baseline characteristics in individuals aged between 30 and 75 years by sex and the study year (PREFREC 2010 and ENSPA 2019) in the provinces of Panama and Colon.

| Demographic and socioeconomic characteristics | Men |  | Women |  |
| :---: | :---: | :---: | :---: | :---: |
|  | PREFREC 2010 | ENSPA 2019 | PREFREC 2010 | ENSPA 2019 |
|  | ( $\mathrm{N}=255481$ ) | ( $\mathrm{N}=666 \mathrm{752}$ ) | ( $\mathrm{N}=531$ 325) | ( $\mathrm{N}=677$ 584) |
| Age, years |  |  |  |  |
| Median (IQR) | 52.0 (40.0-62.0) | 50.0 (40.0-61.0) | 48.0 (39.0-58.0) | 48.0 (38.0-60.0) |
| Ethnicity-\% (95\% CI) |  |  |  |  |
| Mestizo | 60.4 (54.7-65.9) | 47.5 (42.4-51.6) | 60.7 (57.0-64.2) | 49.4 (46.9-51.8) |
| Afro-Panamanian | 17.9 (13.9-22.6) | 24.8 (21.2-28.9) | 20.4 (17.7-23.3) | 20.5 (18.6-22.5) |
| Caucasian | 12.7 (9.3-17.0) | 21.2 (17.9-24.9) | 14.0 (11.5-16.8) | 21.1 (19.2-23.3) |
| Indigenous | 2.8 (1.3-6.2) | 4.3 (3.0-6.0) | 3.1 (2.0-4.8) | 5.3 (4.2-6.7) |
| Others | 6.3 (3.9-10.0) | 2.2 (1.3-3.7) | 1.9 (1.1-3.5) | 3.6 (2.8-4.8) |
| Region-\% (95\% CI) |  |  |  |  |
| Urban | 85.3 (82.9-87.4) | 79.5 (76.5-82.2) | 87.6 (86.2-88.9) | 82.0 (80.3-83.5) |
| Non-urban | 14.7 (12.6-17.1) | 20.5 (17.8-23.5) | 12.4 (11.1-13.8) | 18.0 (16.5-19.7) |
| Education-\% (95\% CI) |  |  |  |  |
| No/primary education | 26.8 (22.4-31.8) | 20.3 (17.4-23.4) | 32.1 (28.7-35.7) | 23.0 (21.0-25.0) |
| Secondary education | 44.5 (38.8-50.2) | 58.4 (54.2-62.4) | 46.7 (42.9-50.5) | 55.0 (52.5-57.4) |
| Higher education | 28.7 (23.6-34.5) | 21.4 (18.0-25.1) | 21.3 (18.4-24.5) | 22.1 (20.1-24.2) |
| Monthly Family Income-\% (95\% CI) |  |  |  |  |
| $<250$ PAB | 19.3 (15.9-23.8) | 19.2 (16.3-22.5) | 31.7 (28.4-35.3) | 23.9 (22.0-26.0) |
| 250-999 PAB | 64.3 (58.8-69.3) | 58.6 (54.4-62.6) | 55.5 (51.7-59.3) | 61.0 (58.6-63.4) |
| $\geq 1,000 \mathrm{PAB}$ | 16.2 (12.4-20.8) | 22.2 (18.8-26.0) | 12.7 (10.3-15.7) | 15.1 (13.3-17.0) |
| Established risk factors for hypertension-\% (95\% CI) |  |  |  |  |
| Tobacco consumption |  |  |  |  |
| Current smoker | 11.7 (8.6-15.8) | 7.6 (5.7-10.0) | 3.9 (2.6-5.7) | 2.4 (1.7-3.3) |
| Ex-smoker | 47.0 (41.3-52.7) | 4.2 (2.9-6.2) | 15.7 (13.1-18.6) | 1.0 (0.6-1.6) |
| Non-smoker | 41.3 (35.8-47.0) | 88.2 (85.3-90.6) | 80.5 (77.3-83.3) | 96.6 (95.6-97.4) |
| BMI categories ${ }^{1}$ |  |  |  |  |
| Underweight | 1.4 (0.5-3.7) | 1.4 (0.7-2.6) | 1.1 (0.6-1.9) | 1.5 (1.0-2.3) |
| Normal weight | 40.7 (35.2-46.5) | 23.6 (20.2-27.4) | 26.3 (23.1-29.7) | 18.3 (16.4-20.4) |
| Overweight | 32.7 (27.7-38.1) | 41.1 (36.8-45.6) | 37.3 (33.7-41.1) | 31.1 (28.8-33.5) |
| Obesity | 25.2 (20.5-30.5) | 33.8 (29.7-38.2) | 35.3 (31.8-39.0) | 49.1 (46.5-51.7) |
| Physical inactivity | 20.4 (16.4-25.1) | 52.5 (48.2-56.7) | 12.3 (10.1-15.0) | 66.9 (64.4-69.3) |
| Family history of hypertension | 64.0 (58.2-69.4) | 37.1 (33.2-41.1) | 74.4 (71.0-77.5) | 46.0 (43.5-48.4) |
| Self-reported medical history of diabetes | 10.1 (7.3-13.8) | 4.9 (3.5-6.8) | 9.5 (7.5-11.9) | 8.6 (7.3-10.2) |
| Hypertension-\% (95\% CI) |  |  |  |  |
| Yes | 51.6 (45.7-57.5) | 46.5 (42.1-51.0) | 46.0 (42.1-49.9) | 42.1 (39.6-44.7) |
| Hypertension unawareness ${ }^{2}$ | 32.3 (25.4-40.1) | 52.3 (45.9-58.6) | 16.1 (12.6-20.4) | 33.3 (29.8-37.0) |
| Healthcare-related factors-\% (95\% CI) |  |  |  |  |
| BP assessment in the year before study enrollment | 73.9 (68.7-78.5) | 34.9 (31.0-39.0) | 79.8 (76.6-82.7) | 40.2 (37.8-42.6) |
| At least a health check-up in the year before study enrollment | NA | 57.2 (53.0-61.2) | NA | 68.5 (66.2-70.8) |
| BP assessment ${ }^{3}$ | NA | 48.2 (42.7-53.7) | NA | 49.6 (46.7-52.6) |

$\mathrm{N}=$ weighted study population. $\mathrm{IQR}=$ interquartile range. $\mathrm{PAB}=$ Panamanian balboa. $\mathrm{BP}=$ blood pressure $. \mathrm{NA}=$ not applicable. $\mathrm{BMI}=$ body mass index.
MFI = monthly family income. SBP = systolic blood pressure. $\mathrm{DBP}=$ diastolic blood pressure. $\%=$ percentages. CIs = confidence intervals.
${ }^{1}$ According to World Health Organization (WHO).
${ }^{2}$ SBP/DBP $\geq 140 / 90 \mathrm{mmHg}$ and no self-reported medical history of hypertension.
${ }^{3}$ BP assessment among those with at least a health check-up in the year before study enrollment. Missing data on the PREFREC 2010 study: BP measurements ( $\mathrm{N}=83$ $177 ; 11 \%)$, MFI ( $\mathrm{N}=8675 ; 7.8 \%$ ). Missing data on the ENSPA 2019 study: BP measurements ( $\mathrm{N}=167088 ; 12 \%$ ), MFI ( $\mathrm{N}=65961 ; 5 \%$ ), physical inactivity ( $\mathrm{N}=259$ 434; 19\%), BMI ( $\mathrm{N}=164047$; 12\%).
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participants reported having secondary education (an increase of 13.9 percentage points for men and 8.3 percentage points for women). Fewer men reported having higher education $(28.7 \%$ vs $21.4 \%)$, while a slight increase was present in women ( $21.3 \%$ vs $22.1 \%$ ) (Table 1). Regarding MFI, fewer women reported MFI of $<250$ PAB in the ENSPA (2019), compared to the PREFREC (2010) ( $31.7 \%$ vs $23.9 \%$ ), whereas MFI of $250-999$ PAB and MFI of $\geq 1,000$ PAB increased by $8.3 \%$ and $2.4 \%$, respectively. Among men, MFI of $<250 \mathrm{PAB}$ remained unchanged (20.0\%), MFI of $250-999$ PAB decreased by $5.7 \%$, and MFI of $\geq 1,000 \mathrm{PAB}$ increased by $6.0 \%$ (Table 1 ).

## Established risk factors for hypertension

The prevalence of current and ex-smokers in PREFREC (2010) was $11.7 \%$ and $47.0 \%$ in men, respectively, and among women $3.9 \%$ and $15.7 \%$, respectively. In the ENSPA (2019) study, the prevalence of current smokers was $7.6 \%$ for men and $2.4 \%$ for women, whereas the prevalence of ex-smokers was $4.2 \%$ in men and $1.0 \%$ in women. Regarding BMI, we found that the prevalence of overweight in men was $32.7 \%$ ( $95 \%$ CI: 27.7-38.1) in PREFREC (2010) and 41.1\% ( $95 \%$ CI: $36.8-45.6$ ) in ENSPA (2019). In contrast, among women, obesity was present in 35.3\% (95\% CI: 31.8-39.0) of participants in PREFREC (2010) and in 49.1\% (95\% CI: 46.551.7) in ENSPA (2019). Physical inactivity in PREFREC (2010) was present in $20.4 \%$ and $12.3 \%$ of men and women, respectively, and in ENSPA (2019), it accounted for $52.5 \%$ of men and $66.9 \%$ of women (Table 1).

## Hypertension

In the PREFREC (2010), the prevalence of hypertension and hypertension unawareness in men were $51.6 \%$ ( $95 \%$ CI: $45.7-57.5$ ) and $32.3 \%$ ( $95 \%$ CI: $25.4-40.1$ ), respectively, and in women $46.0 \%$ ( $95 \%$ CI: 42.1-49.9) and $16.1 \%$ ( $95 \%$ CI: 12.6-20.4), respectively (Table 1 and Fig 2). In the ENSPA (2019), the prevalence of hypertension and hypertension unawareness in men were $46.5 \%$ ( $95 \%$ CI: 42.1-51.0) and $52.3 \%$ ( $95 \%$ CI: 45.9-58.6), respectively, and in women $42.1 \% ~(95 \% ~ C I: ~ 39.6-44.7) ~ a n d ~ 33.3 \% ~(95 \% ~ C I: ~ 29.8-37.0), ~ r e s p e c t i v e l y ~(T a b l e ~ 1 ~ a n d ~$ Fig 2). As shown in the S2 and S3 Tables, after merging all data from both studies and adjusting for all other variables, there was a $21 \%$ borderline decrease in the odds of having hypertension in the ENSPA (2019) study compared to the PREFREC study (ENSPA OR: 0.79; 95\% CI: $0.62-1.02$ ), but a $70 \%$ increase in the odds of being unaware (ENSPA OR: $1.70 ; 95 \%$ CI: $1.15-$ 2.51), respectively.

## Healthcare-related factors

We found that fewer men reported having a BP assessment in the year before study enrollment in the ENSPA (2019) study compared to the PREFREC (2010) study (73.9\% [95\% CI: 68.778.5] in PREFREC (2010) vs $34.9 \%$ [ $95 \%$ CI: 31.0-39.0] in ENSPA (2019)). Similarly, women displayed a decrease from $79.8 \%$ ( $95 \%$ CI: 76.6-82.7) in PREFREC (2010) to $40.2 \% ~(95 \% \mathrm{CI}$ : 37.8-42.6) in ENSPA (2019) (Fig 3). Additionally, in ENSPA (2019), 57.2\% of men and 68.5\% of women had a health check-up in the year before study enrollment. Amongst them, $48.2 \%$ of men and $49.6 \%$ of women reported having their BP assessed the year before study enrollment.

## Risk factors associated with unawareness of hypertension

Table 2 shows the crude and adjusted analysis for the associations between each exposure variable and hypertension unawareness, stratified by the study year. Regarding demographic factors, men were more likely to be unaware, compared with women, in the PREFREC (2010)


Fig 2. Prevalence and unawareness of hypertension stratified by sex and the study year. $\mathrm{BP}=$ blood pressure. The PREFREC (2010) study: men $(\mathrm{n}=834$ [ $\mathrm{N}=255481])$, women $(\mathrm{n}=1899$ [ $\mathrm{N}=531326])$. The ENSPA (2019) study: men ( $\mathrm{n}=1317$ [ $\mathrm{N}=666751]$ ), women ( $\mathrm{n}=3334$ [ $\mathrm{N}=677584]$ ).
https://doi.org/10.1371/journal.pone.0276222.g002
(OR: 2.31; 95\% CI: 1.29-4.12) and in the ENSPA (2019) (OR: 2.09; 95\% CI: 1.37-3.17) (model B in Table 2). Whereas individuals aged $<50$ years were more likely to be unaware, compared with those aged $\geq 50$ years, in PREFREC (2010) (OR: $1.84 ; 95 \%$ CI: $1.06-3.17$ ) and in ENSPA (2019) (OR: 2.43; 95\% CI: 1.57-3.76) (model B in Table 2). No association with ethnicity was found (S4 and S5 Tables).

When considering hypertension risk factors, obesity was associated with lower odds of having hypertension unawareness, compared to a normal weight, in PREFREC (2010) (OR: 0.48; $95 \%$ CI: $0.25-0.91$ ), but not in ENSPA (2019) (OR: $0.76 ; 95 \%$ CI: $0.44-1.29$ ) (model B in Table 2). Similarly, physical inactivity was associated with decreased odds of having hypertension unawareness in PREFREC (2010) (OR: $0.43 ; 95 \%$ CI: $0.22-0.84$ ), but not in ENSPA (2019) (OR: $0.88 ; 95 \%$ CI: $0.57-1.36$ ) (model B in Table 2). Family history of hypertension had a borderline inverse association for hypertension unawareness in PREFREC (2010) (OR: 0.62; $95 \%$ CI: $0.36-1.06$ ) and an inverse association in ENSPA (2019) (OR: $0.18 ; 95 \%$ CI: $0.11-0.27$ ) (model B in Table 2). No association was found with tobacco consumption regardless of the study year, and self-reported medical history of diabetes had a borderline inverse association with hypertension unawareness in ENSPA (2019) (OR: 0.55 ; 95\% CI: $0.31-1.00$ ) (model B in Table 2).


Fig 3. $B P$ assessment in the year before study enrollment stratified by sex and the study year. $\mathrm{BP}=\mathrm{blood}$ pressure. The PREFREC (2010) study: men ( $\mathrm{n}=834$ [ $\mathrm{N}=255481]$ ), women ( $\mathrm{n}=1899$ [ $\mathrm{N}=531$ 326]). The ENSPA (2019) study: men ( $n=1317$ [ $\mathrm{N}=666751]$ ), women ( $\mathrm{n}=3334$ [ $\mathrm{N}=677584]$ ).
https://doi.org/10.1371/journal.pone.0276222.g003
Furthermore, having a BP assessment in the year before study enrollment was associated with decreased odds of being unaware in PREFREC (2010) (OR: $0.46 ; 95 \% \mathrm{CI}: 0.25-0.87$ ) and in ENSPA (2019) (OR: 0.35 ; 95\% CI: $0.23-0.53$ ) (model B in Table 2).

When examining the association between socioeconomic factors and unawareness, in PREFREC (2010), higher odds of having hypertension unawareness was found with having no/primary education (OR: 2.27; 95\% CI: 1.06-4.86), compared to higher education, and in ENSPA (2019), with living in a non-urban region (OR: 1.63; 95\% CI: 1.03-2.60), compared to living in an urban region (model B in Table 2).

## Discussion

Our findings indicate a borderline decrease in the odds of having hypertension between the PREFREC (2010) and ENSPA (2019) studies, but an increase in the odds of being unaware of hypertension. Further, there was an increase in the prevalence of overweight and obesity in the provinces of Panama and Colon.

Panama has experienced exceptional economic growth over the past three decades, evidenced by the increase in GDP per capita, which has doubled in the last ten years [12]. This growth, in turn, has produced a rapidly expanding urbanization associated with increased migration from rural to urban areas [30,31], which has augmented the number of people exposed to lifestyle factors related to hypertension, such as westernized dietary habits, excess sodium in the diet, harmful alcohol consumption, stress, and physical inactivity [32-34]. It is likely that these factors have contributed to the increase in the prevalence of obesity in Panama over the past three decades (from $3.8 \%$ in men and $7.6 \%$ in women, in 1982 , to $16.9 \%$ in men

Table 2. Crude and adjusted logistic regression analysis of hypertension unawareness (SBP/DBP $\geq \mathbf{1 4 0} / \mathbf{9 0} \mathbf{~ m m H g}$ without a self-reported medical history of hypertension) in hypertensive individuals aged between 30 and 75 years by the study year (PREFREC 2010 and ENSPA 2019) in the provinces of Panama and Colon. Odds ratio (OR) and 95\% confidence intervals (CIs).

|  | PREFREC 2010 |  |  | ENSPA 2019 |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Crude analysis OR (95\% CI) | Adjusted analysis |  | Crude analysis | Adjusted analysis |  |
|  |  | OR (95\% CI) |  | OR (95\% CI) | OR (95\% CI) |  |
|  |  | Model A | Model B |  | Model A | Model B |
| Demographic Factors |  |  |  |  |  |  |
| Sex |  |  |  |  |  |  |
| Women (reference) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Men | 2.49 | 2.30 | 2.31 | 2.19 | 2.13 | 2.09 |
|  | (1.60-3.89) | (1.33-3.96) | (1.29-4.12) | (1.62-2.97) | (1.42-3.20) | (1.37-3.17) |
| Age group |  |  |  |  |  |  |
| $\geq 50$ years (reference) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| $<50$ years | 1.69 | 1.68 | 1.84 | 2.22 | 2.29 | 2.43 |
|  | (1.07-2.65) | (1.01-2.80) | (1.06-3.17) | (1.60-3.09) | (1.54-3.40) | (1.57-3.76) |
| Established risk factors for hypertension |  |  |  |  |  |  |
| Tabaco consumption |  |  |  |  |  |  |
| Non-smoker | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Ex-smoker | 1.28 | 0.90 | 0.81 | 0.89 | 1.34 | 1.24 |
|  | (0.78-2.10) | (0.50-1.62) | (0.45-1.47) | (0.37-2.12) | (0.38-4.72) | (0.33-4.68) |
| Current smoker | 1.64 | 0.56 | 0.56 | 1.53 | 0.94 | 0.73 |
|  | (0.66-4.10) | (0.16-1.95) | (0.16-1.97) | (0.68-3.42) | (0.34-2.58) | (0.27-1.98) |
|  |  |  |  |  |  |  |
| Underweight | 3.38 | 2.62 | 2.36 | 0.69 | 0.89 | 0.88 |
|  | (0.88-12.98) | (0.52-13.11) | (0.46-12.01) | (0.11-4.49) | (0.24-3.35) | (0.24-3.32) |
| Normal weight | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Overweight | 0.70 | 0.64 | 0.63 | 0.96 | 1.09 | 1.16 |
|  | (0.41-1.18) | (0.37-1.09) | (0.36-1.10) | (0.60-1.54) | (0.65-1.85) | (0.68-1.98) |
| Obesity | 0.45 | 0.45 | 0.48 | 0.60 | 0.79 | 0.76 |
|  | (0.25-0.80) | (0.24-0.84) | (0.25-0.91) | (0.39-0.94) | (0.47-1.34) | (0.44-1.29) |
| Physical inactivity |  |  |  |  |  |  |
| No (reference) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Yes | 0.54 | 0.47 | 0.43 | 0.95 | 0.92 | 0.88 |
|  | (0.29-0.99) | (0.25-0.88) | (0.22-0.84) | (0.68-1.33) | (0.61-1.39) | (0.57-1.36) |
|  |  |  |  |  |  |  |
| No (reference) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Yes | 0.51 | 0.60 | 0.62 | 0.17 | 0.18 | 0.18 |
|  | (0.31-0.83) | (0.35-1.00) | (0.36-1.06) | (0.11-0.24) | (0.12-0.27) | (0.11-0.27) |
| Self-reported medical history of diabetes |  |  |  |  |  |  |
| No (reference) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Yes | 0.85 | 1.00 | 1.09 | 0.31 | 0.57 | 0.55 |
|  | (0.45-1.60) | (0.51-2.00) | (0.54-2.19) | (0.18-0.54) | (0.32-1.02) | (0.31-1.00) |
| BP assessment in the year before study enrollment |  |  |  |  |  |  |
| No (reference) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) | 1 (ref) |
| Yes | 0.31 | 0.48 | 0.46 | 0.25 | 0.35 | 0.35 |
|  | (0.18-0.54) | (0.27-0.86) | (0.25-0.87) | (0.18-0.36) | (0.23-0.52) | (0.23-0.53) |
| Socioeconomic Factors |  |  |  |  |  |  |
| Region |  |  |  |  |  |  |
| Urban (reference) | 1 (ref) |  | 1 (ref) | 1 (ref) |  | 1 (ref) |

Table 2. (Continued)


SBP = systolic blood pressure. $\mathrm{DBP}=$ diastolic blood pressure. $\mathrm{OR}=$ odds ratio. CIs $=$ confidence intervals. $\mathrm{BP}=$ blood pressure. $\mathrm{BMI}=$ body mass index.
PAB = Panamanian Balboa.
Model A = adjusted by sex, age group, ethnicity, BMI categories, physical inactivity, family history of hypertension, self-reported medical history of diabetes, tobacco consumption, and BP assessment in the year before study enrollment.

Model B = further adjusted by region, education, and monthly family income.
${ }^{1}$ According to World Health Organization (WHO).
https://doi.org/10.1371/journal.pone.0276222.t002
and $23.8 \%$ in women, in 2008 [35]) and continues to rise as suggested by our study. We also found that more than half of individuals in the ENSPA (2019) study were physically inactive, a higher estimate than that reported in other countries in Latin America and the Caribbean [36].

The aforementioned findings could explain our high estimates of prevalence of hypertension compared with those reported by other countries in the Americas region [7, 9-11, 37-39]. Nevertheless, we found a decrease in the prevalence of hypertension between the PREFREC (2010) and ENSPA (2019) study, as opposed to other countries in the region, such as Peru, that reported an increasing prevalence of hypertension attributed to recent economic growth [10], whereas the prevalence in Brazil [37], Canada [7], and the United States [9] has remained unchanged. In agreement with our study, recent studies have reported a decrease in the prevalence of hypertension despite an increase in obesity [4, 11]. Our results suggest that other risk factors associated with hypertension such as salt intake, smoking, and exposure to dietary fatty acids have improved [4, 11]. For example, in the case of smoking, in 2005, Panama adopted the WHO Framework Convention on Tobacco Control. Since then, two laws were introduced: a law regulating tobacco control in 2008, and a tobacco tax increase in 2009 [40, 41]. Associated with these laws, there has been a decrease in tobacco consumption and a reduction in the incidence of acute myocardial infarction [41]. Likewise, we found an improvement in education levels: an important and well-known determinant of health [15, 32, 39, 42]. Our study suggests that as health determinants, such as education, improve and risk factors for hypertension, such as tobacco consumption, are diminished, the prevalence of hypertension as well as its potential complications are reduced.

Our results showed an increased prevalence of hypertension unawareness across both studies in the provinces of Panama and Colon. Similarly, a significant increase in the prevalence of hypertension unawareness was found in Canada [7], the United States [9], and in Peru [10]. In contrast, recent findings from other studies performed in Chile [11] and Brazil [36], showed either a plateau or a decrease in hypertension unawareness. When global trends in hypertension were examined between 1990 to 2019 in Latin America and the Caribbean, a slight decrease in hypertension unawareness was found until the mid-2000s before flattening [4]. The increase in hypertension unawareness in some countries has been attributed to reduced funding directed to hypertension programs, a fragmented healthcare system, lack of educational programs, and lack of BP screening in the younger population (18-44 years) [7, 9, 10].

Our study identified a decrease in the prevalence of BP assessment in the year before study enrollment between the PREFREC (2010) and ENSPA (2019) study. As expected, BP assessment in the year before study enrollment was a factor strongly associated with decreased odds of having hypertension unawareness in both studies, which is consistent with previous research suggesting that the frequency of BP assessment is related to awareness [4, 9, 43-45]. Furthermore, we found that approximately half of the individuals in the ENSPA (2019) study who participated in health check-ups did not have a BP assessment, pointing to a significant flaw of these health check-ups and a lack of compliance with the national healthcare program standards [21]. Of note, the current national healthcare program states that BP levels should be measured in the right arm at every health check-up in patients 18 years or older.

Several factors were associated with decreased odds of having hypertension unawareness, including female sex, older age ( $\geq 50$ years), obesity, physical inactivity, family history of hypertension, and BP assessment in the year before study enrollment. Women were less likely to be unaware, compared to men, a finding consistent with a large body of evidence on sex and health-seeking behaviors [39, 43, 45-51]. Older individuals ( $\geq 50$ years) were less likely to have hypertension unawareness compared to younger individuals ( $<50$ years). This is likely due to a higher healthcare utilization by older individuals, as other studies have suggested [32, 39, 45-49, 51, 52]. Another factor associated with a lower risk of hypertension unawareness was having a family history of hypertension, which is consistent with previous findings [43, 46, 49, 52].

Paradoxically, in the PREFREC (2010) study, obesity and physical inactivity were found to be associated with decreased odds of having hypertension unawareness. One possible explanation may be that these groups have been targeted by screening programs and have more contact with health professional [32, 39, 44, 46-52]. On the other hand, the inverse association with hypertension unawareness was not observed in the ENSPA (2019) study, suggesting a decrease in regular health check-ups or reduced screening in this group of individuals.

Socioeconomic factors related to hypertension unawareness differed across both studies. We found in the PREFREC (2010) study that having no education/primary education and in the ENSPA (2019) study that living in a non-urban region were associated with increased odds of having hypertension unawareness. The lack of association of education and unawareness in the ENSPA (2019) study could be explained by the finding of improved educational levels, when compared to the PREFREC (2010) study. Both of these socioeconomic factors have been previously recognized in many countries [10, 32, 39, 44, 46-48, 51]. However, among Chinese men and women, having a low educational level was associated with a lower risk of being unaware [52]. Additionally, compared to other studies [47, 50], we did not find an association between household income and hypertension unawareness.

Hypertension has emerged as an important risk factor in younger populations (aged 25-49 years) [53]. A previous Panamanian study reported that hypertension was the most common risk factor associated with stroke in young adults [54]. Likewise, a prospective study from

China found that having early-onset hypertension ( $<45$ years) increases CVD mortality compared with late-onset hypertension ( $\geq 65$ years) [55]. Therefore, greater emphasis should be placed on screening the younger population, as stated in our national healthcare program [21].

Our study has several limitations. First, the PREFREC (2010) and ENSPA (2019) are crosssectional study designs and thus, causality should not be inferred. Second, medical history of hypertension was based on self-reporting and there may have been reporting biases. Moreover, although both studies collected three BP measurements for each participant, these were taken during a single visit (diagnosis requires two or more visits) and in different arms. However, this is a common approach in large-scale epidemiological studies, and because of design and time constrains, measurements in both arms could not be assessed [10, 26, 46, 52, 56]. Third, in the two studies, physical inactivity was assessed with different methodologies, so we were not able to analyze variations across studies. Fourth, we did not perform a temporal analysis of trend over time. Finally, although our study was limited to two Panamanian provinces that contain around $60 \%$ of the country's population, the results cannot be extrapolated to the whole country. Moreover, the provinces of Panama and Colon are predominantly urban, with the province of Colon having the highest concentration of people of Afro-Panamanians, who have a higher risk of hypertension and obesity compared to other ethnic groups in the country $[25,35]$. The strength of this study was the large study population as well as the sampling methodology that allowed us to provide robust information.

## Conclusions

Although the study found a reduction in the prevalence of hypertension, our results also showed a worrisome increase in hypertension unawareness. These findings may be due, in part, to a weakening of CVD prevention healthcare program, as evidenced by a decrease in BP assessment in the year before study enrollment when one compares the PREFREC (2010) and the ENSPA (2019) studies. In addition, almost half of study participants who reported having had a health check-up in the year prior to the ENSPA (2019) study, denied having had a BP assessment.

Our study points to the need of strengthening the CVD healthcare program implemented in 2014 [21]. Future research aiming to develop interventions to improve early diagnosis and reduce risk factors associated with hypertension is warranted. It is likely that if the finding related to hypertension unawareness is not promptly addressed, we could expect a rebound in the decrease in the prevalence of hypertension and the observed decline in CVD mortality [15, 57].

## Supporting information

S1 Table. Definition of exposure variables based on the study year.
(PDF)
S2 Table. Crude and adjusted logistic regression analysis of hypertension prevalence (SBP/ DBP $\geq 140 / 90 \mathbf{m m H g}$ detected during the study and/or self-reported medical history of hypertension) in participants aged between 30 and 75 years in the provinces of Panama and Colon using merged data from the PREFREC (2010) and ENSPA (2019) studies. Odds ratio (OR) and $95 \%$ confidence intervals (CIs).
(PDF)
S3 Table. Crude and adjusted logistic regression analysis of hypertension unawareness (SBP/DBP $\geq 140 / 90 \mathrm{mmHg}$ without a self-reported medical history of hypertension) in hypertensive participants aged between 30 and 75 years in the provinces of Panama and

Colon using merged data from the PREFREC (2010) and ENSPA (2019) studies. Odds ratio (OR) and 95\% confidence intervals (CIs).
(PDF)
S4 Table. Crude and adjusted logistic regression analysis of hypertension unawareness (SBP/DBP $\geq 140 / 90 \mathrm{mmHg}$ without a self-reported medical history of hypertension) in hypertensive participants aged between 30 and 75 years in the PREFREC study in the provinces of Panama and Colon. Odds ratio (OR) and 95\% confidence intervals (CIs). (PDF)

S5 Table. Crude and adjusted logistic regression analysis of hypertension unawareness (SBP/DBP $\geq 140 / 90 \mathrm{mmHg}$ without a self-reported medical history of hypertension) in hypertensive participants aged between 30 and 75 years in the ENSPA study in the provinces of Panama and Colon. Odds ratio (OR) and $95 \%$ confidence intervals (CIs). (PDF)

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## References

1. World Health Organization. Global Status Report on noncommunicable diseases. 2014 [cited 10 June 2021]. Available from: http://apps.who.int/iris/bitstream/handle/10665/148114/9789241564854_eng. pdf?sequence=1
2. Pan American Health Organization. Annual Report of the Director of the Pan American Sanitary Bureau 2020. Saving Lives and Improving Health and Well-Being. Washington, DC: PAHO. 2020.
3. Mills KT, Stefanescu A, He J. The global epidemiology of hypertension. Nat Rev Nephrol 2020; 16: 223-37. https://doi.org/10.1038/s41581-019-0244-2 PMID: 32024986
4. Zhou B, Carrillo-Larco RM, Danaei G, Riley LM, Paciorek CJ, Stevens GA, et al. Worldwide trends in hypertension prevalence and progress in treatment and control from 1990 to 2019: a pooled analysis of 1201 population-representative studies with 104 million participants. Lancet 2021. https://doi.org/10. 1016/S0140-6736(21)01330-1 PMID: 34450083
5. Krist AH, Davidson KW, Mangione CM, Cabana M, Caughey AB, Davis EM, et al. US Preventive Services Task Force. Screening for Hypertension in Adults: US Preventive Services Task Force reaffirmation recommendation statement. JAMA 2021; 325: 1650-56.
6. Mills KT, Bundy JD, Kelly TN, Reed JE, Kearney PM, Reynolds K, et al. Global Disparities of Hypertension Prevalence and Control: A Systematic Analysis of Population-Based Studies From 90 Countries. Circulation 2016; 134: 441-50. https://doi.org/10.1161/CIRCULATIONAHA.115.018912 PMID: 27502908
7. Leung AA, Williams JVA, McAlister FA, Campbell NRC, Padwal RS. Worsening Hypertension Awareness, Treatment, and Control Rates in Canadian Women Between 2007 and 2017. Can J Cardiol 2020; 36: 732-739. https://doi.org/10.1016/j.cjca.2020.02.092 PMID: 32299635
8. Zhou B, Danaei G, Stevens GA, Bixby H, Taddei C, Carrillo-Larco RM, et al. Long-term and recent trends in hypertension awareness, treatment, and control in 12 high-income countries: an analysis of 123 nationally representative surveys. Lancet 2019; 394: 639-51. https://doi.org/10.1016/S0140-6736 (19)31145-6 PMID: 31327564
9. Muntner P, Hardy ST, Fine LJ, Jaeger BC, Wozniak G, Levitan EB, et al. Trends in Blood Pressure Control Among US Adults with Hypertension, 1999-2000 to 2017-2018. JAMA 2020; 324: 1190-1200. https://doi.org/10.1001/jama.2020.14545 PMID: 32902588
10. Villarreal-Zegarra D, Carrillo-Larco RM, Bernabe-Ortiz A. Short-term trends in the prevalence, awareness, treatment, and control of arterial hypertension in Peru. J Hum Hypertens 2021; 35: 462-71. https://doi.org/10.1038/s41371-020-0361-1 PMID: 32518303
11. Passi-Solar Á, Margozzini P, Mindell JS, Ruiz M, Valencia-Hernandez CA, Scholes S. Hypertension care cascade in Chile: a serial cross-sectional study of national health surveys 2003-2010-2017. BMC Public Health 2020; 20:1397. https://doi.org/10.1186/s12889-020-09483-x PMID: 32928176
12. The World Bank website. Panama Data. 2019 [cited 26 July 2021]. Available from: https://data. worldbank.org/country/PA
13. International Monetary Fund website. Report for Selected Countries and Subjects: October 2019 [cited 13 July 2021]. Available from: https://www.imf.org/en/Publications/WEO/weo-database/2019/October/ weo-report?c=311,213,314,313,316,339,218,223,228,233,238,321,243,248,253,328,258,336,263, $268,343,273,278,283,288,293,361,362,364,366,369,298,299, \& s=$ NGDPRPPPPC,NGDPDPC,\&sy= 2010\&ey=2019\&ssm=0\&scsm=1\&scc=0\&ssd=1\&ssc=0\&sic=0\&sort=country\&ds=.\&br=1
14. The World Bank website. Gini Index (World Bank estimate)-Panama. 2019 [cited 18 July 2021]. Available from: https://data.worldbank.org/indicator/SI.POV.GINI?locations=PA
15. Quiel L, Moreno Velásquez I, Gómez B, Motta J, Herrera-Ballesteros V. Social determinants and cardiovascular disease mortality in Panama, 2012-2016. BMC Public Health 2019; 19:199. https://doi.org/ 10.1186/s12889-019-6508-8 PMID: 30770742
16. Castro F, Zúñiga J, Higuera G, Carrión Donderis M, Gómez B, Motta J. Indigenous Ethnicity and Low Maternal Education Are Associated with Delayed Diagnosis and Mortality in Infants with Congenital Heart Defects in Panama. PLOS ONE 2016; 11: e0163168. https://doi.org/10.1371/journal.pone. 0163168 PMID: 27648568
17. Castro F, Shahal D, Tarajia M, et al. Baseline characteristics, survival and direct costs associated to treatment of gastric cancer patients at the National Oncology Institute of Panama from 2012 to 2015: a hospital-based observational study. BMJ Open 2017; 7:e017266. https://doi.org/10.1136/bmjopen-2017-017266 PMID: 28947456
18. Pan American Health Organization. Health in the Americas+, 2017 Edition. Summary: Regional Outlook and Country Profiles. Washington, DC: PAHO. 2017.
19. The Gorgas Memorial Institute of Health Studies website. Summary sheet of non-communicable diseases. 2019 [cited 20 July 2021]. Available from: http://www.gorgas.gob.pa/SIGENSPA/documentos/5. \%20Factsheet/HOJA\%20RESUMEN_ENT.pdf
20. National Institute of Statistics and Census website. Deaths and mortality rate of the main causes of death in the Republic of Panama. 2019 [cited 27 May 2021]. Available from: https://inec.gob.pa/ archivos/P00140176420210302150121Cuadro\%2013.pdf
21. Panamanian Ministry of Health. Technical-Administrative Standards: Adult and Elderly Health Programs. 2018 [cited 5 May 2021]. Available from: https://www.educacioninterprofesional.org/sites/ default/files/fulltext/2018/2018_pan_norma_adulto_adulto_mayor.pdf
22. Moreno Velásquez I, Castro F, Gómez B, Cuero C, Motta J. Chronic Kidney Disease in Panama: Results from the PREFREC Study and National Mortality Trends. Kidney Int Rep 2017; 2: 1032-41. https://doi.org/10.1016/j.ekir.2017.05.016 PMID: 29270512
23. The Gorgas Memorial Institute of Health Studies website. Interactive Geographic Information System of the National Health Survey of Panama. 2019 [cited 26 July 2021]. Available from: http://gorgas.gob.pa/ SIGENSPA/Inicio.htm
24. National Institute of Statistics and Census. Territorial Distribution and Internal Migration in Panama: Census 2010 [cited 10 July 2021]. Available from: https://www.inec.gob.pa/archivos/

## P0705547520200925152431Distribución\%20Territorial\%20y\%20Migración\%20Interna\%20en\%

 20Panamá-Censo2010_F.pdf25. Mc Donald Posso AJ, Motta Borrel JA, Fontes F, Cruz Gonzalez CE, Pachón Burgos AA, Cumbrera Ortega A. High Blood Pressure in Panama: Prevalence, Sociodemographic and Biologic Profile, Treatment, and Control (STROBE). Medicine (Baltimore) 2014; 93: e101.
26. World Health Organization. The WHO STEPwise approach to noncommunicable disease risk factor surveillance. WHO STEPS Surveillance Manual; 2017. pp. 184 [cited 9 July 2021]. Available from: https://www.who.int/ncds/surveillance/steps/STEPS_Manual.pdf
27. Unger T, Borghi C, Charchar F, Khan NA, Poulter NR, Prabhakaran D, et al. 2020 International Society of Hypertension Global Hypertension Practice Guidelines. Hypertension 2020; 75: 1334-57. PMID: 32370572
28. Harvard T.H. Chan website. Obesity Prevention Source: Why Use BMI? [cited 27 July 2021]. Available from: https://www.hsph.harvard.edu/obesity-prevention-source/obesity-definition/obesity-definition-fullstory/
29. World Health Organization. Global Physical Activity Surveillance: Analysis Guide [cited 9 July 2021]. Available from: https://www.who.int/ncds/surveillance/steps/resources/GPAQ_Analysis_Guide.pdf
30. United Nations Development Programme website. Panama in brief. 2021 [cited 12 August 2021]. Available from: https://www.pa.undp.org/content/panama/es/home/countryinfo/\#Econom\�\�a
31. Pittí A, Gaudin Y, Hess S. Characterization of rural areas in Panama based on national statistics: social, economic and demographic approach. Mexico City: Economic Commission for Latin America and the Caribbean; 2021. 48 p.
32. Li J, Shi L, Li S, Xu L, Qin W, Wang H. Urban-rural disparities in hypertension prevalence, detection, and medication use among Chinese Adults from 1993 to 2011. Int J Equity Health 2017; 16: 50. https:// doi.org/10.1186/s12939-017-0545-7 PMID: 28288635
33. Ibrahim MM, Damasceno A. Hypertension in developing countries. Lancet 2012; 380: 611-9. https:// doi.org/10.1016/S0140-6736(12)60861-7 PMID: 22883510
34. Ibrahim MM. Hypertension in Developing Countries: A Major Challenge for the Future. Curr Hypertens Rep 2018; 20: 38. https://doi.org/10.1007/s11906-018-0839-1 PMID: 29717393
35. Sasson M, Lee M, Jan C, Fontes F, Motta J. Prevalence and Associated Factors of Obesity among Panamanian Adults. 1982-2010. PLoS ONE 2014; 9: e91689. https://doi.org/10.1371/journal.pone. 0091689 PMID: 24621825
36. Guthold R, Stevens GA, Riley LM, Bull FC. Worldwide trends in insufficient physical activity from 2001 to 2016: a pooled analysis of 358 population-based surveys with 1.9 million participants. Lancet Glob Health 2018; 6: e1077-86. https://doi.org/10.1016/S2214-109X(18)30357-7 PMID: 30193830
37. Souza NP de, Cesse EÂP, Souza WV de, Fontbonne A, Carvalho Barreto MNS de, Le Goff M, et al. Temporal variation in prevalence, awareness and control of hypertension in urban and rural areas in Northeast Brazil between 2006 and 2016. Cad Saúde Pública 2020; 36: e00027819. https://doi.org/10. 1590/0102-311X00027819 PMID: 32321071
38. Campos-Nonato I, Hernández-Barrera L, Pedroza-Tobías A, Medina C, Barquera S. Hypertension in Mexican adults: prevalence, diagnosis and type of treatment. Ensanut MC 2016. Salud Pública México 2018; 60: 233-43.
39. Geldsetzer P, Manne-Goehler J, Marcus M-E, Ebert C, Zhumadilov Z, Wesseh CS, et al. The state of hypertension care in 44 low-income and middle-income countries: a cross-sectional study of nationally representative individual-level data from 1.1 million adults. Lancet 2019; 394:652-62. PMID: 31327566
40. Official Digital Gazette. Law No. 13-Which Adoption Measures to Control Tobacco and its Harmful Effects on Health. 2008 [cited 11 August 2021]. Available from: https://www.tobaccocontrollaws.org/ files/live/Panama/Panama\%20-\%20Law\%2013\%20of\%202008.pdf
41. Jan C, Lee M, Roa R, Herrera V, Politis M, Motta J. The Association of Tobacco Control Policies and the Risk of Acute Myocardial Infarction Using Hospital Admissions Data. PLoS ONE 2014; 9: e88784. https://doi.org/10.1371/journal.pone.0088784 PMID: 24520421
42. Fernandez R.M. (2019) Gross Domestic Product and Health. In: Leal Filho W., Wall T., Azul A., Brandli L., Özuyar P. (eds) Good Health and Well-Being. Encyclopedia of the UN Sustainable Development Goals. Springer, Cham.
43. Chau K, Girerd N, Zannad F, Rossignol P, Boivin J-M. Health-related determinants of undiagnosed arterial hypertension: a population-based study. Fam Pract 2019; 36: 276-83. https://doi.org/10.1093/ fampra/cmy075 PMID: 30165649
44. Zhou J, Fang S. Association between Undiagnosed Hypertension and Health Factors among MiddleAged and Elderly Chinese Population. Int J Environ Res Public Health 2019; 16: 1214. https://doi.org/ 10.3390/ijerph16071214 PMID: 30987361
45. Chow CK, Teo KK, Rangarajan S, Islam S, Gupta R, Avezum A, et al. Prevalence, Awareness, Treatment, and Control of Hypertension in Rural and Urban Communities in High-, Middle-, and Low-Income Countries. JAMA 2013; 310: 959-68. https://doi.org/10.1001/jama.2013.184182 PMID: 24002282
46. Li T, Song X, Wu J, Li Z, Li L, Yu Y, et al. Awareness of hypertension and related factors in northeastern China: a cross-sectional study. J Hum Hypertens 2020; 34: 43-50. https://doi.org/10.1038/s41371-019-0263-2 PMID: 31548618
47. Ahmed S, Tariquijaman Md, Rahman MdA, Hasan MdZ, Hasan MdM. Inequalities in the prevalence of undiagnosed hypertension among Bangladeshi adults: evidence from a nationwide survey. Int J Equity Health 2019; 18: 33. https://doi.org/10.1186/s12939-019-0930-5 PMID: 30770739
48. Fenech G, Vallée A, Cherfan M, Kab S, Goldberg M, Zins M, et al. Poor Awareness of Hypertension in France: The CONSTANCES Population-Based Study. Am J Hypertens 2020; 33: 543-51. https://doi. org/10.1093/ajh/hpaa018 PMID: 32202627
49. Lee H-Y. Socioeconomic Disparities in the Prevalence, Diagnosis, and Control of Hypertension in the Context of a Universal Health Insurance System. J Korean Med Sci 2017; 32: 561-7. https://doi.org/10. 3346/jkms.2017.32.4.561 PMID: 28244279
50. Hasan MdM, Tasnim F, Tariqujjaman Md, Ahmed S, Cleary A, Mamun A. Examining the prevalence, correlates and inequalities of undiagnosed hypertension in Nepal: a population-based cross-sectional study. BMJ Open 2020; 10: e037592. https://doi.org/10.1136/bmjopen-2020-037592 PMID: 33004393
51. Park JE, Park J-H, Chang SJ, Lee JH, Kim SY. The Determinants of and Barriers to Awareness and Treatment of Hypertension in the Korean Population. Asia Pac J Public Health 2019; 31: 121-35. https://doi.org/10.1177/1010539518825006 PMID: 30678483
52. Santosa A, Zhang Y, Weinehall L, Zhao G, Wang N, Zhao Q, et al. Gender differences and determinants of prevalence, awareness, treatment and control of hypertension among adults in China and Sweden. BMC Public Health 2020; 20: 1763. https://doi.org/10.1186/s12889-020-09862-4 PMID: 33228600
53. Vos T, Lim SS, Abbafati C, Abbas KM, Abbasi M, Abbasifard M, et al. Global burden of 369 diseases and injuries in 204 countries and territories, 1990-2019: a systematic analysis for the Global Burden of Disease Study 2019. Lancet 2020; 396: 1204-22. https://doi.org/10.1016/S0140-6736(20)30925-9 PMID: 33069326
54. Romero L, Cigarruista Y, Mackay P, Sánchez R, Serrano A, Vega I, et al. Factores asociados a enfermedades cerebrovasculares en adultos jóvenes. Complejo Hospitalario Metropolitano Dr. Arnulfo Arias Madrid. 2008-2012. Panamá. Revista medico científica 2013; 26; 39-48.
55. Wang C, Yuan Y, Zheng M, Pan A, Wang M, Zhao M, et al. Association of Age of Onset of Hypertension with Cardiovascular Diseases and Mortality. J Am Coll Cardiol 2020; 75: 2921-30. https://doi.org/10. 1016/j.jacc.2020.04.038 PMID: 32527401
56. Bryan S, Saint-Pierre Larose M, Campbell N, Clarke J, Tremblay MS. Resting blood pressure and heart rate measurement in the Canadian Health Measures Survey, cycle 1. Health Rep 2010; 21:71-8. PMID: 20426229
57. Carrión Donderis M, Moreno Velásquez I, Castro F, Zúñiga J, Gómez B, Motta J. Analysis of mortality trends due to cardiovascular diseases in Panama, 2001-2014. Open Heart 2016; 3: e000510. https:// doi.org/10.1136/openhrt-2016-000510 PMID: 28123756
