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# High blood pressure: prevalence and adherence to guidelines in a population-based cohort 

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## Summary

QUESTIONS UNDER STUDY: High blood pressure, the single leading health risk factor worldwide, contributes greatly to morbidity and mortality. This study aimed to add to the understanding of diagnosed and undiagnosed high blood pressure in Switzerland and to evaluate adherence to hypertension guidelines.
METHODS: Included were 3962 participants from the first (2001-2003) and second (2010-2011) follow-ups of the population-based Swiss Cohort Study on Air Pollution and Lung and Heart Disease in Adults. High blood pressure was defined as blood pressure $\geq 140 / 90 \mathrm{~mm} \mathrm{Hg}$ and the prevalence of doctor-diagnosed hypertension was based on questionnaire information.
RESULTS: High blood pressure was found in $34.9 \%$ of subjects, $49.1 \%$ of whom were unaware of this condition; $30.0 \%$ had doctor-diagnosed hypertension and, although $82.1 \%$ of these received drug treatments, in only $40.8 \%$ was blood pressure controlled ( $<140 / 90 \mathrm{~mm} \mathrm{Hg}$ ). Substantial first-line beta-blocker use and nonadherence to comorbidity-specific prescription guidelines were observed and remained mostly unexplained. Age-adjusted

[^0]rates of unawareness and uncontrolled hypertension were more than $20 \%$ higher than in the USA.
CONCLUSIONS: There is room for improvement in managing hypertension in Switzerland. Population-based observational studies are essential for identifying and evaluating unmet needs in healthcare; however, to pinpoint the underlying causes it is imperative to facilitate linkage of cohort data to medical records.

Key words: hypertension; blood pressure; prevalence; awareness; treatment; control; guideline adherence; Switzerland

## Introduction

High blood pressure (HBP) accounts for $17.8 \%$ of deaths and $7.0 \%$ of disability-adjusted life years (DALYs) globally, making it the number one single risk factor worldwide [1]. In Switzerland, a country with universal healthcare coverage and which ranks fourth in per capita total health expenditure [2], an estimated $27 \%$ of adults have hypertension [3] and $18.7 \%$ of deaths and $7.9 \%$ of DALYs [4] are attributed to this risk factor. The burden of disease related to hypertension is mainly mediated by ischaemic heart disease and stroke [5].
In Switzerland, concordant national data on hypertension prevalence from surveys, primary care or hospital discharge statistics are available [6]. However, when hypertension prevalence is based on diagnosed hypertension, the true prevalence of hypertension may be underestimated owing to the often asymptomatic nature of HBP. Regional estimates of undiagnosed hypertension in Switzerland vary from 6 to $13 \%$ of the adult population, this range may in part be explained by differences in age distributions and the age-standardisation of the reported rates in the respective studies $[7,8]$. However, current nationwide populationbased data covering varied geographical and cultural re-
gions is lacking. Once diagnosed, one may expect uniformity in managing hypertension in a small and economically well-to-do country like Switzerland with its own guidelines for the management of hypertension [9]; however, national information on guideline adherence is scarce.
Managing hypertension according to guidelines remains a practical challenge, especially in a poly-morbid, polymedicated population where more than one conflicting guideline may apply and drugs with antihypertensive properties are used for the treatment of other diseases. Moreover guidelines worldwide differ [9-12], and more than one document may influence clinical practice in a given country.
This study aimed to describe the prevalence, awareness, treatment and control of hypertension in Switzerland, and to evaluate adherence to the Swiss Society of Hypertension ( SSH ) guidelines in regards to drug choice.

## Methods

## Study population

This analysis was based on the Swiss Cohort Study on Air Pollution and Lung and Heart Disease in Adults (SAPALDIA), with participants originally recruited from the general adult population in eight communities covering the range of geographical (urban, rural, alpine) and cultural (German-, French-, and Italian-speaking) conditions in Switzerland [13]. Beginning with a baseline assessment in 1991, the first and second follow-ups were in 2001-2003 (SAPALDIA 2) and 2010-2011 (SAPALDIA 3), respectively. Blood pressure (BP) measurement was added from SAPALDIA 2 onwards. Ethical approval was obtained from the Swiss Academy of Medical Sciences’ Central Ethical Committee and the appropriate cantonal ethics committees. Participants gave written informed consent. This analysis focused on data from SAPALDIA 3 using elements from SAPALDIA 2 for added depth. Included were SAPALDIA 2 and 3 participants for whom all relevant data were available: that is, questionnaire information on morbidities, medications, lifestyle and clinical information on blood pressure, weight and height $(\mathrm{n}=3962)$ (see appendix, supplementary fig. S1).

## Questionnaire information

Morbidity data were collected during an in-person interview with the use of questionnaires. Questions included: "Has a doctor ever told you that you have one of the following diseases?" followed by a list of diseases including hypertension, arrhythmia, asthma, etc. Only participants with asthma who had also had at least one attack in the last year or were taking asthma medication were considered as asthmatics in this analysis. Participants with atrial fibrillation were defined as those with arrhythmia taking either vitamin K antagonists or platelet aggregation inhibitors (excluding heparin). All medications taken were listed in a questionnaire. Drugs considered asantihypertensive were angiotensin converting-enzyme inhibitors (ACEIs), angiotensin II receptor blockers (ARBs), beta-blockers (BBs), calcium antagonists, diuretics, renin inhibitors and other hypertensives with anatomical therapeutic chemical
(ATC) code C02. Questionnaire information on smoking (never, former, current), alcohol consumption (number of standard glasses/day), fruit and vegetable intake (number of servings/day) and physical activity (minutes of moderate and/or vigorous activity/week) was also collected. Neighbourhood-level socioeconomic index, a 0 - to 100-point scale based on education, head of household occupation, rent and crowding derived from 2000 census data [14], was allocated on the basis of SAPALDIA 2 home address. Degree of urbanisation (high, medium, low) was allocated on the basis of home address according to the European-wide degree of urbanisation (DEGURBA) classification for local administrative units - usually communes.

## Clinical measurements

Blood pressure was measured twice by trained field workers using an automatic device (705CP, OMRON, Tokyo, Japan) with a cuff of appropriate size (after having measured arm circumference) and using the Riva-Rocci method, in the sitting position after minimum 10 minutes rest, on the left arm, with 3 minutes between measurements. Blood pressure values are expressed as mean systolic and mean diastolic pressures in mm Hg . Height and weight were measured when the participants were without shoes and wearing light clothing.

## High blood pressure and doctor-diagnosed

 hypertensionHBP was defined as systolic blood pressure $\geq 140$ and/ or diastolic blood pressure $\geq 90 \mathrm{~mm} \mathrm{Hg}$, simply described as HBP or "blood pressure $\geq 140 / 90$ " from now onwards. Doctor-diagnosed hypertension (ddHT) was based on the following question "Has a doctor ever told you that you have hypertension?" HBP unawareness refers to those with HBP but not ddHT and is expressed as a percentage of those with HBP. Uncontrolled ddHT refers to those with ddHT and HBP and is expressed as a percentage of those with ddHT.

## Guideline adherence

Guideline adherence was evaluated in those with ddHT and was based on the Swiss Society of Hypertension 2009 guidelines [15] in effect at SAPALDIA 3 data collection. Specific recommendations are described in table 1. Current Swiss Society of Hypertension guidelines are available on their website [9].
Recommended lifestyle goals were combined in a lifestyle score, one point attributed for each recommendation successfully met; not smoking, low risk alcohol consumption ( $\leq 1$ glass/day by women and $\leq 2$ glasses/day by men) [16], fruit/vegetable consumption $\geq 5 /$ day, body mass index $<25$ $\mathrm{kg} / \mathrm{m}^{2}$ and regular physical activity ( 150 minutes of moderate or 60 minutes of vigorous activity per week alone or in an equivalent combination with moderate activity).

## Statistical analysis

Unless noted otherwise, data presented are from SAPALDIA 3. Descriptive results are presented as number and percent for categorical variables and as median and interquartile range for quantitative variables. Chi-squared
tests, t-tests and Wilcoxon rank-sum tests were used for bivariate analysis as appropriate. Several multivariable logistic regression models were constructed using the backwards stepwise method to assess determinants of the following outcomes; ddHT, HBP, HBP unawareness, uncontrolled ddHT and new onset of cardiovascular events (angina pectoris, myocardial infarction, heart failure and stroke). Covariates systematically considered in all models were: sex, age, language, degree of urbanisation, education, neighbourhood-level socioeconomic index and lifestyle score. Covariates additionally considered in some models were: systolic blood pressure, HBP, HBP unawareness, uncontrolled ddHT and a binomial variable for concomitant cardiovascular or metabolic disease (dyslipidaemia, diabetes, angina pectoris, myocardial infarction, heart failure or stroke). As neighbourhood-level socioeconomic index was available only for SAPALDIA 2 addresses, in sensitivity analyses the participants who had moved during the SAPALDIA 2-3 interval were excluded to determine if observed associations persisted. Since association patterns did not differ, results are not presented. All p-values $<0.05$ were considered statistically significant. Statistical analysis was performed using STATA 13 (StataCorp. 2013. Stata: Release 13. Statistical Software. College Station, TX: StataCorp LP.)

## Results

Available characteristics of included and excluded participants at baseline and the first and second follow-ups are presented in supplementary table S1. In comparison with the excluded participants, our study sample was better educated, included fewer smokers and more physically active individuals having less HBP and/or ddHT, but having similar prevalences of other comorbidities. The analytical sample consisted of 3962 participants, $50 \%$ men and $50 \%$ women, with a median age of 60.4 years, ranging from 37 to 81 years old.

High blood pressure and hypertension; prevalence, awareness, and control
Of the 3986 participants, $34.9 \%$ had HBP (blood pressure $\geq 140 / 90$ ) and $30.0 \%$ had ddHT. These two groups overlapped to some degree resulting in $47.1 \%$ having HBP and/ or ddHT. Of those with HBP, 49.1\% did not report a diagnosis of hypertension, and were therefore considered unaware, representing $17.1 \%$ of the analytical sample as a whole. Of those with ddHT, $59.2 \%$ had HBP and were therefore considered to have uncontrolled ddHT. Prevalence of HBP, ddHT, and HBP and/or ddHT was statistically significantly higher in men than in women ( $\mathrm{p}<0.001$ ). Prevalence data including sex differences are described in figure 1. Even after controlling for age, lifestyle score, presence of concomitant cardiovascular or metabolic disease, language, education level, degree of urbanisation and neighbourhood level socioeconomic level, men were more likely to have HBP (odds ratio [OR] 1.59, $95 \%$ confidence interval [CI] 1.38-1.85, p $<0.001$ ) and ddHT (OR 1.38, $95 \%$ CI 1.18-1.62, $\mathrm{p}<0.001$ ).
Systolic blood pressure $\geq 160$ and/or diastolic blood pressure $\geq 100 \mathrm{~mm} \mathrm{Hg}$ was found in $19.5 \%, 33.0 \%, 27.0 \%$ and $20.8 \%$ of those with ddHT, uncontrolled ddHT, HBP and HBP unawareness, respectively. These blood pressure values correspond to grade 2 or 3 hypertension. Detailed dis-


Figure 1
Prevalence of high blood pressure and doctor-diagnosed hypertension, stratified by gender (\% [95\%CI])
p -values for sex differences determined with chi-squared tests, * p $<0.001,{ }^{a} p=0.262,{ }^{b} p=0.104$
$\mathrm{CI}=$ confidence interval; ddHT = doctor diagnosed hypertension; HBP $=$ high blood pressure

Table 1: Selected 2009 Swiss Society of Hypertension guidelines [15].

## Lifestyle improvements for all

Smoking cessation, limited alcohol consumption, a diet rich in fruits and vegetables and poor in salt, physical activity, weight control

## Drug treatment

First-line drugs: ACEI, ARB, CA, diuretics
Alternative drug: beta-blockers (not recommend as a first-line drug as some studies show them to be ineffective in reducing stroke and coronary heart disease and to have negative effects on lipid and glucose metabolism. However, beta-blockers are permitted to initiate therapy when certain comorbidities are present) [26]

## Comorbidity specific treatement, recommended drugs

| Coronary heart disease | ACEI, ARB, BB, CA |
| :--- | :--- |
| Heart failure | ACEI, ARB, BB, diuretic |
| Diabetes | ACEI, ARB, BB*, CA |
| Dyslipidaemia | ACEI, ARB, CA |
| Migraines | BB |
| Asthma | avoid BB* |

Target blood pressure <140/90 in general
$<130 / 80$ in those with diabetes or kidney disease ${ }^{\dagger}$
$\mathrm{ACEI}=$ angiotensin converting enzyme inhibitor; $\mathrm{ARB}=$ angiotensin II receptor blocker; $\mathrm{BB}=$ beta-blocker; CA = calcium antagonist

* Beta-blockers are not recommended in people with abnormal glucose tolerance, but permitted in overt diabetes, blood pressure reduction being the main goal. The careful use of highly cardioselective beta-blockers is permitted in individuals with stable asthma if a good indication to beta-blocker use is present and with adequate monitoring [26].
${ }^{\dagger}$ Note, the 2015 Swiss guidelines differ, placing target blood pressure at <140/85 in those with diabetes or kidney disease [9].
tribution of blood pressure severity in these groups is described in table 2.


## Determinants of high blood pressure unawareness

A multivariable logistic regression with HBP unawareness as outcome was adjusted for age, sex, SAPALDIA 2 systolic blood pressure, presence of concomitant cardiovascular or metabolic disease, language, education level, degree of urbanisation, neighbourhood socioeconomic status and lifestyle score. Those more likely to be unaware of HBP were men (OR 1.42, 95\% CI 1.09-1.86, p = 0.010), younger participants (OR 0.88 per 10-year increase, $95 \% \mathrm{CI}$ $0.78-1.00, \mathrm{p}=0.047$ ), those with lower systolic blood pressure (OR 0.90 per $2-\mathrm{mm} \mathrm{Hg}$ increase, $95 \%$ CI $0.88-0.92$, $\mathrm{p}<0.001$ ), those without concomitant cardiovascular disease (OR 2.35, 95\% CI $1.82-3.02, \mathrm{p}<0.001$ ), those with lower neighbourhood-level socioeconomic status (OR 0.86 per 10 -point increase, $95 \%$ CI $0.75-0.99, \mathrm{p}=0.031$ ) and those with higher lifestyle score (OR 1.36 per point increase, $95 \%$ CI 1.19-1.55, p $<0.001$ ).

## Target blood pressure and determinants of uncontrolled hypertension

Based on our blood pressure measurements, the general target blood pressure of $<140 / 90 \mathrm{~mm} \mathrm{Hg}$ was reached by $40.8 \%$ of the 1188 participants with ddHT. Of the 232 participants with ddHT and diabetes or kidney disease, only $23.7 \%$ (18.6-29.7\%) reached the treatment goal of $<130 / 80$ mm Hg.
A multivariable logistic regression with uncontrolled ddHT as outcome was adjusted for age, sex, SAPALDIA 2 systolic blood pressure, presence of concomitant cardiovascular or metabolic disease, lifestyle score, language, education level, degree of urbanisation and neighbourhood-level socioeconomic status. More likely to have uncontrolled ddHT were those with higher SAPALDIA 2 systolic blood pressure (OR 1.08 per 2-mm Hg increase, $95 \%$ CI 1.06-1.10, p $<0.001$ ), and those with lower lifestyle score, though only with borderline statistical significance (OR 0.88 per point increase, $95 \%$ CI $0.77-1.00, \mathrm{p}=0.057$ ).

## Outcomes of high blood pressure, unawareness and uncontrolled hypertension

To determine the outcomes of HBP, HBP unawareness and uncontrolled ddHT and to evaluate possible overestimation of HBP due to the white coat effect (blood pressure measurements only being available from the study centre visits),
we assessed the 10-year morbidity in the respective groups from SAPALDIA 2. Controlling for age and sex, the OR for developing angina pectoris during the SAPALDIA 2-3 interval in those with HBP compared with those with normal blood pressure in SAPALDIA 2 was 2.2 (1.2-4.1, p = 0.013). Similarly, ORs for developing myocardial infarction, heart failure, stroke and any of these four cardiovascular diseases were $3.3(1.8-6.0, \mathrm{p}<0.001), 2.5(1.3-4.7, \mathrm{p}=$ $0.004), 1.7(0.9-3.2, \mathrm{p}=0.088)$ and $2.2(1.6-3.1, \mathrm{p}<0.001)$, respectively. Adjusted for age and sex, the ORs of developing cardiovascular outcomes during the SAPALDIA 2-3 interval in those unaware compared with those aware of their HBP at SAPALDIA 2 were for some outcomes higher than one and for some lower than one but never reached statistical significance. In contrast, after adjusting for age and sex, the ORs of developing angina pectoris, myocardial infarction, heart failure or stroke during the SAPALDIA 2-3 interval were higher than one in those with uncontrolled compared to those with controlled ddHT at SAPALDIA 2, although not reaching statistical significance.

## Hypertension pharmacotherapy

Overall, $82.1 \%$ of the 1188 participants with ddHT took blood pressure-lowering agents, $42.3 \%$ of these took one drug, $36.3 \%$ took two and $21.4 \%$ took three or more. Of those taking at least two, $65.0 \%$ used a combined fixed dose preparation. When compared with subjects with drugtreated ddHT, those with non-drug-treated ddHT had higher lifestyle scores (Wilcoxon rank-sum $\mathrm{p}=0.030$ ), and higher mean systolic blood pressure ( 147.9 mm Hg [145.5-150.3] compared with $142.6 \mathrm{~mm} \quad \mathrm{Hg}$ [141.5-143.8]) and diastolic blood pressure $(87.7 \mathrm{~mm} \mathrm{Hg}$ [86.4-89.0] compared with 81.3 mm Hg [80.6-82.0], p $\leq 0.001$ ).

## Beta-blockers as a first-line drug

Though the Swiss Society of Hypertension did not recommend beta-blockers as a first-line drug, except in the presence of some comorbidities, $100(24.2 \%, 20.4-28.7 \%)$ of the 412 hypertensive participants on monotherapy took beta-blockers. Providing possible explanations, $8.0 \%$ had coronary heart disease, $1.0 \%$ had heart failure, $9.0 \%$ had atrial fibrillation, $7.0 \%$ had diabetes, $16.0 \%$ had migraines and $34.0 \%$ had at least one of the above listed comorbidities. First-line beta-blocker prescription did not vary by age ( $\mathrm{p}=0.385$ ).

| Blood pressure ( mm Hg ) | In those with ddHT $(n=1188)$ | In those with uncontrolled ddHT $(\mathrm{n}=703)$ | In those with HBP ( $\mathrm{n}=1381$ ) | In those unaware of their HBP ( $\mathrm{n}=678$ ) |
| :---: | :---: | :---: | :---: | :---: |
| Normal < 140/90 | $\begin{aligned} & 40.8 \\ & (38.1-43.7) \end{aligned}$ | - | - | - |
| $\begin{aligned} & \text { Grade 1* } \\ & \text { 140-159 / 90-99 } \end{aligned}$ | $\begin{aligned} & 39.6 \\ & (36.9-42.5) \end{aligned}$ | $\begin{aligned} & 67.0 \\ & (63.4-70.4) \end{aligned}$ | $\begin{aligned} & 73.0 \\ & (70.6-75.3) \end{aligned}$ | $\begin{aligned} & 79.2 \\ & (76.0-82.1) \end{aligned}$ |
| $\begin{aligned} & \text { Grade 2* } \\ & 160-179 / 100-109 \end{aligned}$ | $\begin{aligned} & 15.5 \\ & (13.5-17.7) \end{aligned}$ | $\begin{aligned} & 26.2 \\ & (23.0-29.6) \end{aligned}$ | $\begin{aligned} & 21.8 \\ & (19.7-24.1) \end{aligned}$ | $\begin{aligned} & 17.3 \\ & (14.6-20.3) \end{aligned}$ |
| $\begin{aligned} & \text { Grade } 3^{*} \\ & \geq 180 / 110 \end{aligned}$ | $\begin{gathered} 4.0 \\ (3.1-5.3) \end{gathered}$ | $\begin{array}{\|l\|} \hline 6.8 \\ (5.2-9.0) \end{array}$ | $\begin{aligned} & \hline 5.2 \\ & (4.2-6.5) \end{aligned}$ | $\begin{gathered} \hline 3.5 \\ (2.4-5.2) \end{gathered}$ |

[^1]*Grades 1-3 represent systolic blood pressure and/or diastolic blood pressure within the respective ranges. Participants were allocated to a group based on highest level of blood pressure, be it systolic or diastolic.

## Comorbidity specific treatment

Antihypertensive drug classes recommended on the basis of comorbidity as well as the distribution of guideline adherence are described in table 3. Of those with drug treated ddHT and dyslipidaemia ( $\mathrm{n}=406$ ), $18.2 \%$ were taking neither ACEIs, ARBs nor calcium antagonists, despite recommendations. Only $33.8 \%$ of this seeming guideline nonadherence could be explained by the presence of other comorbidities where beta-blockers or diuretics were permitted or are commonly prescribed (coronary heart disease, heart failure, atrial fibrillation, diabetes, migraines or kidney disease). Beta-blockers were prescribed in 8 (19.5\%) individuals with both asthma and hypertension. All eight had asthma medication and in six the prescribed betablockers were cardioselective. Of the possible explanations for beta-blocker prescription (coronary heart disease, heart failure, migraines or atrial fibrillation), only one of these asthmatic participants had migraines and one had atrial fibrillation.

## Discussion

High blood pressure and hypertension: prevalence, awareness and control
In this analysis from the population-based SAPALDIA cohort $50 \%$ had either ddHT or blood pressure $\geq 140 / 90$. Thirty percent had ddHT, among whom $60 \%$ had uncontrolled ddHT. Some uncontrolled ddHT may reflect a white coat effect or may have been intentional, accounting for orthostatic hypotension. Still, rates of uncontrolled HBP were high and the distribution of blood pressure severity was equally surprising, as $21 \%$ of those unaware of their HBP had measurements $\geq 160 / 100 \mathrm{~mm} \mathrm{Hg}$. Furthermore, recent research indicates that the 140 mm Hg target is not low enough; a 120 mm Hg target results in significantly lower cardiovascular mortality and morbidity in nondiabetic patients without previous stroke but at high risk of cardiovascular events [17]. The association of both HBP and uncontrolled ddHT with long-term cardiovascular outcomes reinforces the validity of our measurements and underlines the importance of early diagnosis and proper treatment. Similar odds of cardiovascular outcomes between those aware and unaware of their HBP were expected as HBP is harmful regardless of having a diagnosis or not.

After definitions were harmonised, our results were comparable to other Swiss data on hypertension prevalence, awareness and control [3, 7, 18] (see supplementary table S2). When comparing age-specific harmonised definitions from our study and 2009-2010 US data [19], we found that the prevalence of HBP and/or antihypertensive drug intake were similar, whereas awareness and control were respectively $20.0-37.4 \%$ and $23.8-34.4 \%$ lower in our data (see supplementary table S3). This is consistent with the literature which shows lower awareness and control in Europe compared to the USA [20-22].
In the current study, we assessed the prevalence, determinants and outcomes of HBP (defined as blood pressure $\geq 140 / 90 \mathrm{~mm} \mathrm{Hg}$ ) and hypertension as diagnosed by a doctor. The white coat effect, which should be excluded by performing 24-hour or home blood pressure measurement, and masked hypertension may have led to respectively over- or underestimation of HBP. Because of their longterm participation in SAPALDIA, our sample may be healthier and more compliant than the general population. Our results might thus underestimate the true population prevalence of diagnosed or undiagnosed hypertension. On the other hand, our blood pressure values are the mean of two measurements from a single visit and may overestimate an individual's usual blood pressure [23]. Still, 140/90 remains an arbitrary cut-off across a spectrum of abovenormal blood pressure within which cardiovascular risk increases [24]. Approximately 50\% of DALYs attributed to above-normal blood pressure occur below this 140/90 cutoff [5].

## Groups at higher risk

Identifying groups at higher risk for HBP unawareness and uncontrolled ddHT is challenging; accordingly, the results from the literature differ. In our study, younger age, male sex, healthier lifestyle, lower neighbourhood-level socioeconomic status and lower SAPALDIA 2 systolic blood pressure were associated with HBP unawareness. Uncontrolled hypertension was associated with higher SAPALDIA 2 systolic blood pressure. These findings are, in part, echoed in studies from Switzerland and elsewhere [7, 8, 19]. Differences in healthcare utilisation, drug prescription or adherence trends may explain these results, although we were unable to asses this in the current study.

| Comorbidity (number of participants) and recommended drug classes | Only recommended drugs prescribed n (\%) | Mix of recommended and nonrecommended drugs prescribed n (\%) | Only nonrecommended drugs prescribed n (\%) |
| :---: | :---: | :---: | :---: |
| Coronary heart disease ( $\mathrm{n}=86$ ) ACEI, ARB, BB, CA | 49 (57.0\%) | 36 (41.9\%) | 1 (1.2\%) |
| Heart failure $(\mathrm{n}=35)$ ACEI, ARB, BB, diuretic | 25 (71.4\%) | 8 (22.9\%) | 2 (5.7\%) |
| Diabetes ( $\mathrm{n}=156$ ) ACEI, ARB, BB, CA | 68 (43.6\%) | 82 (52.6\%) | 6 (3.8\%) |
| Dyslipidaemia ( $\mathrm{n}=406$ ) ACEI, ARB, CA | 118 (29.1\%) | 214 (52.7\%) | 74 (18.2\%) |
| Migraines ( $\mathrm{n}=73$ ) BB | 16 (21.9\%) | 17 (23.3\%) | 40 (54.8\%) |
| Asthma ( $\mathrm{n}=41$ ) avoid BB | 33 (80.5\%) | 5 (12.2\%) | 3 (7.3\%) |
| ACEI = angiotensin converting enzyme inhibitor; ARB = angiotensin II receptor blocker; BB = betablocker; CA = calcium antagonist; ddHT = doctor-diagnosed hypertension |  |  |  |

## Pharmacotherapy

High rates of uncontrolled hypertension despite high treatment rates might be explained by the complexity of clinical situations, or by physicians' inertia in changing drugs or increasing dose even when blood pressure targets remain unmet as previously described in Switzerland [25]. Also, drug compliance is challenging because hypertension is chronic and often asymptomatic. In some cases fixed-combination antihypertensive preparations may be preferred to remedy this issue [10].
Several practical challenges arise in aligning pharmacotherapy to guidelines. Guidelines differ worldwide, and several documents potentially influence practitioners in a given country. Even within one document, conflicting recommendations may apply to the same individual. Also, drug choice is influenced by efficacy, tolerance, security and price. Lastly, antihypertensive drugs are used in the treatment of other conditions independently of hypertension. These elements may explain some of the seeming guideline nonadherence.
In our analysis, nonadherence was particularly high regarding beta-blockers. The Swiss Society of Hypertension did not recommend beta-blockers as first-line drugs since studies report them to be ineffective in reducing stroke and coronary heart disease and to have negative effects on lipid and glucose metabolism [26]. Similarly, US and UK guidelines [11, 12] did not generally encourage beta-blockers as first-line drugs. However, the European Society of Hypertension guidelines considered beta-blockers as appropriate for initial treatment [10, 27]. Since $24.2 \%$ of firstline drugs in our data were beta-blockers and their prescription remained largely unexplained by comorbidities, Swiss doctors were possibly unaware of the Swiss Society of Hypertension recommendation or were following European guidelines.
In the recommendation to avoid beta-blockers when asthma is present, the Swiss Society of Hypertension does nuance the statement [26]. However, a recent review recommends prudence, even when using cardioselective beta-blockers in hypertensive asthmatics [28]. Twenty percent of asthmatics with hypertension were receiving betablockers (mainly cardioselective), but clear indications for beta-blocker treatment were not evident in our data.

## Needed action in hypertension management

While there is room for improvement in overall guideline adherence, one main concern is the high prevalence of uncontrolled hypertension. A European-wide survey, including Switzerland, showed that physicians underestimate their role in unmet blood pressure targets [29]. What should be done about guideline nonadherence, particularly in regard to unmet blood pressure goals? Redon et al. propose some practical solutions: increasing both awareness of the harmful nature of hypertension and patient accountability and simplifying treatment with combined preparations [30].

## Strengths and limitations

Participants in the SAPALDIA cohort are characterised in great detail and reflect the geographical and cultural diversity of Switzerland. Yet the Swiss health care system is
highly fragmented and its organisation does not allow for efficient access to objective information on medical diagnoses and treatment. In this study, diagnoses and medication intake were self-reported. The study was not designed to address particularly the issue of uncontrolled hypertension, so information on regularity of blood pressure measurements, drug compliance, or reasons for nonadherence to medical recommendations is lacking. Therefore we cannot fully explain the observed unawareness and noncontrol of hypertension; however, their high prevalence is unlikely to be due to bias or participant behaviour alone.

## Conclusions

Prevalence of HBP, ddHT, HBP unawareness and uncontrolled ddHT were high. Nonadherence to comorbidityspecific drug prescription guidelines remained high and largely unexplained, particularly concerning the use of beta-blockers as first-line drugs. For HBP, a major cardiovascular risk factor, there is still room for substantial improvement from prevention to diagnosis, treatment and control, even in Switzerland, a high-income country. Our results underline the value of population-based cohorts for the identification and characterisation of potential unmet needs in healthcare practice. To further improve the value of these cohorts, it would be essential to facilitate efficient linkage of cohort data to medical records.

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## Appendix: Supplementary figure and tables

## Figure S1

Selection of study population.
BP = blood pressure; SAPALDIA $=$ Swiss Cohort Study on Air Pollution and Lung and Heart Disease in Adults.


|  | SAPALDIA 1 included | SAPALDIA 1 excluded | p -value | SAPALDIA 2 included | SAPALDIA 2 excluded | p-value | SAPALDIA 3 included | SAPALDIA 3 excluded | p-value |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Participants | 3962 | 5689 |  | 3962 | 4085 |  | 3962 | 2126 |  |
| Men | 1984 (50.1\%) | 2761 (48.5\%) | 0.136 | 1984 (50.1\%) | 1883 (46.1\%) | <0.001 | 1984 (50.1\%) | 913 (42.9\%) | <0.001 |
| Age years (IQR) | $\begin{aligned} & 41.2(31.6, \\ & 48.9) \end{aligned}$ | $\begin{aligned} & 42.2 \text { (32.1, } \\ & 51.6) \end{aligned}$ | 0.003 | $52(42.5,59.8)$ | $\begin{aligned} & 53.5(43.7, \\ & 62.8) \end{aligned}$ | <0.001 | $\begin{aligned} & 60.4(50.9, \\ & 68.0) \end{aligned}$ | $\begin{aligned} & 60.1 \text { (50.5, } \\ & 69.5) \end{aligned}$ | 0.447 |
| German-speaking French-speaking Italian-speaking | $\begin{aligned} & 2148 \text { (54.2\%) } \\ & 1267 \text { (32.0\%) } \\ & 546 \text { (13.8\%) } \end{aligned}$ | $\begin{array}{r} 2758 \text { (48.5\%) } \\ 2004(35.2 \%) \\ 925(16.3 \%) \end{array}$ | <0.001 | $\begin{array}{r} 2148 \text { (54.2\%) } \\ 1267(32.0 \%) \\ 546 \text { (13.8\%) } \end{array}$ | $\begin{array}{r} 2106 \text { (51.6\%) } \\ 1317 \text { (32.2\%) } \\ 662 \text { (16.2\%) } \end{array}$ | 0.005 | $\begin{array}{r} \hline 2150 \text { (54.3\%) } \\ 1278 \text { (32.3\%) } \\ 534 \text { (13.5\%) } \end{array}$ | $\begin{aligned} & 663 \text { (55.9\%) } \\ & 351 \text { (29.6\%) } \\ & 171 \text { (14.4\%) } \end{aligned}$ | 0.214 |
| Highest education Primary Secondary ${ }^{\#}$ Tertiary | $\begin{array}{\|r} 424 \text { (10.7\%) } \\ 2753 \text { (69.6\%) } \\ 781 \text { (19.7\%) } \end{array}$ | $\begin{array}{r} 1199 \text { (21.2\%) } \\ 3644 \text { (64.3\%) } \\ 826 \text { (14.6\%) } \end{array}$ | <0.001 | $\begin{gathered} 168 \text { (4.2\%) } \\ 2559 \text { (64.6\%) } \\ 1235 \text { (31.2\%) } \end{gathered}$ | $\begin{array}{r} 534 \text { (13.1\%) } \\ 2698 \text { (66.1\%) } \\ 847 \text { (20.8\%) } \end{array}$ | <0.001 | $\begin{array}{\|c} 168 \text { (4.2\%) } \\ 2559 \text { (64.6\%) } \\ 1235 \text { (31.2\%) } \end{array}$ | $\begin{array}{r} 246 \text { (11.6\%) } \\ 1421 \text { (66.9\%) } \\ 457 \text { (21.5\%) } \end{array}$ | <0.001 |
| Neighbourhood-level socioeconomic index (0-to 100-point scale) |  |  |  | $\begin{aligned} & 64.2(57.5, \\ & 71.0) \end{aligned}$ | $\begin{aligned} & 63.4 \text { (55.6, } \\ & 70.3) \end{aligned}$ | 0.001 | $\begin{aligned} & 64.2(57.5, \\ & 71.0) \end{aligned}$ | $64(56.4,71.1)$ | 0.557 |
| Degree of urbanisation <br> High <br> Medium <br> Low |  |  |  | $\begin{array}{r} 1187 \text { (30.0\%) } \\ 2244(56.8 \%) \\ 523(13.2 \%) \end{array}$ | $\begin{array}{\|c} 1485(37.1 \%) \\ 2146 \text { (53.6\%) } \\ 370 \text { (9.2\%) } \end{array}$ | <0.001 | $\begin{array}{r} 1121 \text { (28.3\%) } \\ 2277 \text { (57.5\%) } \\ 564 \text { (14.2\%) } \end{array}$ | $\begin{array}{r} 605 \text { (30.0\%) } \\ 1165 \text { (57.8\%) } \\ 246 \text { (12.2\%) } \end{array}$ | 0.065 |
| Blood pressure |  |  |  |  |  |  |  |  |  |
| Systolic BP, mm Hg (IQR) |  |  |  | $\begin{aligned} & 124.5(112.5, \\ & 138.0) \end{aligned}$ | $\begin{aligned} & 127 \text { (114.0, } \\ & 139.5) \end{aligned}$ | <0.001 | $\begin{aligned} & \hline 131.5(120, \\ & 144.5) \end{aligned}$ | $\begin{aligned} & 131 \text { (120.0, } \\ & 143.5) \end{aligned}$ | 0.748 |
| Diastolic BP, mm Hg (IQR) |  |  |  | $\begin{aligned} & 79.0 \text { (72.0, } \\ & 86.5) \end{aligned}$ | $\begin{aligned} & 79.0 \text { (72.0, } \\ & 87.0) \end{aligned}$ | 0.481 | $\begin{aligned} & 78.5(72.0, \\ & 85.0) \end{aligned}$ | $\begin{aligned} & 79.0 \text { (72.0, } \\ & 85.5) \end{aligned}$ | 0.303 |
| HBP and/or ddHT |  |  |  | 1265 (31.9\%) | 1090 (41.2\%) | <0.001 | 1866 (47.1\%) | 724 (58.3\%) | $<0.001$ |
| HBP (BP $\geq 140 / 90$ ) <br> Unaware |  |  |  | $\begin{array}{r} 1032 \text { (26.0\%) } \\ 677 \text { (65.6\%) } \end{array}$ | $\begin{aligned} & 728 \text { (29.2\%) } \\ & 445 \text { (61.1\%) } \end{aligned}$ | $\begin{aligned} & 0.006 \\ & 0.055 \end{aligned}$ | $\begin{array}{\|r\|} \hline 1381 \text { (34.9\%) } \\ 678 \text { (49.1\%) } \end{array}$ | $\begin{aligned} & 331 \text { (34.8\%) } \\ & 173 \text { (55.4\%) } \end{aligned}$ | $\begin{aligned} & 0.977 \\ & 0.043 \end{aligned}$ |
| ddHT Uncontrolled |  |  |  | $\begin{aligned} & 588 \text { (14.8\%) } \\ & 355 \text { (60.4\%) } \end{aligned}$ | $\begin{aligned} & 645 \text { (18.7\%) } \\ & 283 \text { (58.4\%) } \end{aligned}$ | $\begin{array}{\|l\|} \hline<0.001 \\ 0.502 \end{array}$ | $\begin{array}{\|r\|} \hline 1188 \text { (30.0\%) } \\ 703 \text { (59.2\%) } \end{array}$ | $\begin{aligned} & 532 \text { (28.4\%) } \\ & 139 \text { (58.6\%) } \end{aligned}$ | $\begin{aligned} & 0.225 \\ & 0.881 \end{aligned}$ |
| Lifestyle characteristics |  |  |  |  |  |  |  |  |  |
| Never smokers Former smokers Current smokers | $\begin{array}{r} \hline 1960(49.5 \%) \\ 945(23.9 \%) \\ 1052(26.6 \%) \end{array}$ | $\begin{aligned} & \hline 2269 \text { (39.9\%) } \\ & 1236 \text { (21.7\%) } \\ & 2180(38.3 \%) \end{aligned}$ | <0.001 | $\begin{array}{r} \hline 1825 \text { (46.1\%) } \\ 1269(32.0 \%) \\ 868(21.9 \%) \end{array}$ | $\begin{aligned} & 1569 \text { (38.5\%) } \\ & 1255 \text { (30.8\%) } \\ & 1248(30.6 \%) \\ & \hline \end{aligned}$ | <0.001 | $\begin{array}{r} \hline 1787 \text { (45.1\%) } \\ 1542 \text { (38.9\%) } \\ 633 \text { (16.0\%) } \end{array}$ | $\begin{aligned} & 859 \text { (40.8\%) } \\ & 764 \text { (36.3\%) } \\ & 480 \text { (22.8\%) } \end{aligned}$ | <0.001 |
| Low risk alcohol consumption <br> $\leq 1$ glass/day in women $\leq 2$ glasses/day in men |  |  |  | $\begin{aligned} & 1916 \text { (96.9\%) } \\ & 1701 \text { (85.7\%) } \end{aligned}$ | $\begin{aligned} & 690 \text { (94.0\%) } \\ & 460 \text { (87.5\%) } \end{aligned}$ | 0.118 | $\begin{array}{\|l\|l\|} \hline 1910 \text { (96.6\%) } \\ 1759 \text { (88.2\%) } \end{array}$ | $\begin{aligned} & 560 \text { (96.2\%) } \\ & 456 \text { (89.8\%) } \end{aligned}$ | 0.340 |
| Fruit and vegetable intake $\geq 5 /$ day |  |  |  | 867 (21.9\%) | 530 (20.3\%) | 0.132 | 916 (23.1\%) | 243 (22.1\%) | 0.464 |
| Sufficient physical activity |  |  |  | 2913 (73.5\%) | 1745 (67.5\%) | <0.001 | 2967 (74.9\%) | 707 (67.6\%) | <0.001 |
| $\begin{aligned} & \mathrm{BMI}<25 \mathrm{~kg} / \mathrm{m}^{2} \\ & \geq 25 \text { to }<30 \mathrm{~kg} / \mathrm{m}^{2} \\ & \geq 30 \mathrm{~kg} / \mathrm{m}^{2} \end{aligned}$ | $\begin{array}{c\|} \hline 2773 \text { (70.1\%) } \\ 1002 \text { (25.3\%) } \\ 178 \text { (4.5\%) } \\ \hline \end{array}$ | $\begin{array}{\|l\|} \hline 3528 \text { (63\%) } \\ 1581 \text { (28.2\%) } \\ 490 \text { (8.8\%) } \\ \hline \end{array}$ | <0.001 | $\begin{array}{r} \hline 1934 \text { (48.8\%) } \\ 1472(37.2 \%) \\ 556(14.0 \%) \\ \hline \end{array}$ | $\begin{array}{r} 1132 \text { (42.9\%) } \\ 1001 \text { (38.0\%) } \\ 503 \text { (19.1\%) } \\ \hline \end{array}$ | <0.001 | $\begin{array}{\|r} \hline 1669 \text { (42.1\%) } \\ 1568 \text { (39.6\%) } \\ 725 \text { (18.3\%) } \\ \hline \end{array}$ | $\begin{array}{\|l\|} \hline 395 \text { (40.1\%) } \\ 395 \text { (40.1\%) } \\ 194 \text { (19.7\%) } \\ \hline \end{array}$ | 0.437 |
| Comorbidities (self-reported doctor-diagnosed) |  |  |  |  |  |  |  |  |  |
| Angina pectoris |  |  |  | 66 (1.7\%) | 62 (1.8\%) | 0.721 | 82 (2.1\%) | 39 (2.8\%) | 0.103 |
| Myocardial infarct |  |  |  | 47 (1.2\%) | 54 (1.5\%) | 0.181 | 92 (2.3\%) | 75 (4\%) | <0.001 |
| Heart failure |  |  |  | 47 (1.2\%) | 40 (1.5\%) | 0.233 | 61 (1.5\%) | 64 (3.4\%) | <0.001 |
| Atrial fibrillation |  |  |  |  |  |  | 119 (3.0\%) | 27 (2\%) | 0.040 |
| Stroke |  |  |  | 41 (1.0\%) | 48 (1.4\%) | 0.164 | 68 (1.7\%) | 24 (1.7\%) | 0.953 |
| Dyslipidaemia |  |  |  | 339 (38.7\%) | 94 (46.3\%) | 0.045 | 994 (25.1\%) | 375 (20.3\%) | <0.001 |
| Diabetes |  |  |  | 90 (2.3\%) | 151 (4.4\%) | <0.001 | 282 (7.1\%) | 162 (8.6\%) | 0.044 |
| Asthma |  |  |  | 137 (3.5\%) | 155 (3.8\%) | 0.384 | 172 (4.3\%) | 76 (3.7\%) | 0.208 |
| Migraines |  |  |  | 289 (7.3\%) | 276 (8.0\%) | 0.267 | 289 (7.3\%) | 138 (8.3\%) | 0.182 |
| $\mathrm{BMI}=$ body mass index; $\mathrm{BP}=$ blood pressure; ddHT = doctor-diagnosed hypertension; $\mathrm{HBP}=$ high blood pressure; IQR = interquartile range (25th, 75th percentile) \# including apprenticeships |  |  |  |  |  |  |  |  |  |


| Table S2: Comparison of hypertension prevalence between SAPALDIA 3 and other Swiss data. |  |  |
| :--- | :--- | :--- |
|  | SAPALDIA 3 2010-2011 <br> Percent (95\% CI) | Other Swiss data <br> Percent (95\% CI) |
| Prevalence ddHT | $30.0(28.6-31.4)$ | $27^{\#}$ |
| Prevalence of HT: HBP and/or taking antihypertensive drugs | $48.5(46.9-50.0)$ | $36(35-38)^{*}$ |
| Awareness: those with ddHT among those with HBP and/or taking antihypertensive drugs | $58.7(56.4-60.8)$ | $63(61-65)^{*}$ |
| Control: those with BP<140/90 among those with HBP and/or taking antihypertensive drugs | $28.0(26.1-30.1)$ | $23(21-25)^{*}$ |
| Here definitions of "hypertension", "awareness" and "control" were adopted from the CoLaus study. <br> \# Swiss Health Survey 2012, aged 15-75+ years [1] <br> * CoLaus 2003-2006, aged 35-75 years [2] SAPALDIA 3 participants were aged 37-81 years <br> BP = blood pressure; CI = confidence interval; ddHT = doctor-diagnosed hypertension; HT = hypertension; HBP = high blood pressure (blood pressure $\geq 140 / 90 ~ m m ~ H g) ~$ |  |  |


| Table S3: Prevalence comparison between SAPALDIA 3 and United States data. |
| :--- |

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Figures (large format)



[^0]:    ACEI angiotensin converting-enzyme inhibitor
    ARB angiotensin II receptor blocker
    BB beta-blocker
    BP blood pressure
    CI confidence interval
    DALY disability adjusted life year
    ddHT doctor-diagnosed hypertension
    HBP high blood pressure
    OR odds ratio
    SAPALDIA Swiss Cohort Study on Air Pollution and Lung and Heart Disease in Adults

    SSH Swiss Society of Hypertension

[^1]:    ddHT = doctor diagnosed hypertension; HBP = high blood pressure

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