Barriers, enablers and strategies for the treatment and control of hypertension in Nepal: a mixed-methods systematic review

Raja Ram Dhungana (raja.dhungana@gmail.com)
Government of Nepal Department of Livestock Services https://orcid.org/0000-0002-9610-6306

Zeljko Pedisic
Victoria University

Achyut Raj Pandey
ABT Associates Inc

Nipun Shrestha
Victoria University

Maximilian de Courten
Victoria University

Research

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Abstract

Background Untreated and uncontrolled hypertension are major challenges in the cascade of care of hypertension in Nepal. The challenge of achieving effective treatment and control of blood pressure is linked to various barriers to hypertension care at different layers of health system. Evidence is emerging globally on the effectiveness of multi-pronged hypertension control strategies. However, no systematic review of strategies of and factors associated with hypertension treatment and control in Nepal is available. Understanding country-specific factors influencing hypertension care is critical to address the gaps in the management of hypertension. This study aimed to systematically review published literature and synthesise the findings on barriers, enablers and strategies for hypertension treatment and control in Nepal.

Methods Six databases namely Embase, PubMed, Web of Science, CINAHL, ProQuest and WorldCat were systematically searched for studies investigating the factors or strategies in relation to hypertension treatment and control in Nepal. The methodological quality of selected articles was assessed using Mixed Methods Appraisal Tool. Themes on barriers and enablers were generated and framed under "health system" (HS), "provider" and "patient" domains, according to the framework synthesis approach. Findings on hypertension strategies were narratively synthesised.

Results We identified 14 published studies; one with mixed, three with qualitative and 10 with quantitative methods. Eight were related to barriers and enablers and six of hypertension strategies. The identified barriers associated with the HS were: lack of affordable services and lack of resources. The barriers at the provider's level were: communication gaps and long waiting hours for appointments. Poor help-seeking behaviour, non-adherence to medication, perceived side-effects of drugs, financial hardship and lack of family support were barriers identified at patient level. The following enablers were identified: positive illness perception, free essential healthcare services and family support. Strategies implemented across the HS, provider and patient were: establishing digital health records at health centres, health worker's capacity development, health education and yoga practice.

Conclusion There is a range of barriers for hypertension treatment and control in Nepal pertaining to the health system, providers, and patients. Comprehensive interventions are needed at all three levels to further improve management and control of hypertension in Nepal.

Registration: The protocol is registered in PROSPERO (registration number: CRD42020145823)

Background

A political declaration on the prevention and control of non-communicable diseases (NCDs), adopted by the General Assembly of the United Nations in 2011, emphasises the importance of developing national capacities to deal with NCDs, particularly in low- and middle-income countries (LMICs) [1]. Though more and more focus of the health care system is placed on tackling NCDs, the rate of hypertension treatment and control in LMICs is low [2]. Compared to high-income countries, LMICs reported substantially lower proportions of hypertension treatment (55.6% vs. 29.0%) and control (28.4% vs. 7.7%) in 2010 [3]. In three South Asian countries, namely, Bangladesh, India, and Pakistan, 31.9% of the hypertensive population was on anti-hypertensive medication and only 12.9% had controlled blood pressure [4].

Nepal, a South Asian low-income country landlocked between China and India hosts a population of 28 million, with 125 caste/ethnic groups living mostly in rural areas [38, 39]. Under the Ministry of Health and Population, the public HS in Nepal delivers health services to the community through the tertiary, district and primary care centres and health posts. Alongside public health services, private for-profit sectors and non-governmental organisations provide health services in Nepal [40].

With the growing burden of hypertension, untreated and uncontrolled hypertension is prevalent in Nepal [5-9]. The majority of hypertensive individuals were not aware of their condition (awareness = 40.0%); a half of those that were aware were not treated (treatment = 20.2%), and a half of those on treatment, the blood pressure was not controlled (control = 10.5%) in 2016 [10].

Gaps in hypertension treatment and control can be due to various factors associated with the health system (HS), health care providers, and individual patients. Systematic reviews and other studies from outside of Nepal suggest that lack of access, poor service delivery, unaffordability of health services and limited resources are common HS-related barriers [11-16]. Similarly, studies also found that the factors pertaining to patients and providers, such as lack of awareness and treatment, non-adherence to medicine, adverse drug effect, therapeutic inertia and communication gaps, were contributing to uncontrolled hypertension [13, 16-26].

The quantity and the scope of primary research on barriers, enablers and strategies for hypertension control are growing in Nepal. While new evidence from primary studies is evolving, a systematic review is needed for a comprehensive identification, understanding and synthesis of the factors affecting hypertension treatment and control in the Nepali context. Therefore, this review aimed to systematically synthesise evidence on barriers, enablers, and strategies for hypertension treatment and control in Nepal. By providing a comprehensive understanding of HS- and individual-level barriers and enablers and strategies of hypertension treatment and control, this review aims to inform the policymakers and related stakeholders about the gaps in hypertension care and contribute to the development of the contextual and problem-specific strategies to increase hypertension control in Nepal.

Theoretical Framework

A framework was used to organise patient, health care provider and HS (excluding patients and providers) barriers, enablers, and strategies for improving blood pressure control. The framework was derived from the theories used in previous studies on barriers [14, 16, 27], and adapted to also fit the enablers and strategies (Fig 1).
By “barriers”, we considered factors that prevent adequate blood pressure control [27]. These have previously been linked with hypertensive patients, providers, and HS [27]. Patient-reported barriers include, for example, knowledge, perception, beliefs, practices, self-efficacy, motivation, family support, affordability, and socio-cultural factors [16, 28]. These factors may affect health-seeking behaviours, treatment adherence, and follow-up care among patients, ultimately affecting blood pressure control [27]. The commonly cited barriers by the providers comprise a lack of skills, non-compliance with guidelines, lack of motivation and therapeutic inertia [16, 26, 27]. An interaction of patient and provider factors can sometimes lead to a communication gap and a poor relationship, affecting treatment adherence and follow-up care [29].

Furthermore, enablers are the personal or contextual factors that facilitate actions or a required behaviour for hypertension treatment and control. They include, for example, perceived severity of illness and family supports [30].

Strategies to improve blood pressure control include interventions, programs, or policies targeted at the providers, patients, or other components of the HS. Some of the proposed evidence-based strategies are: task shifting [31]; team-based care [32]; HS financing, including insurance coverage and co-payments of medical care [14]; HS arrangement, including improved service delivery [33]; using combination therapy [34] or polypill [35]; home-based blood pressure monitoring [36]; and educational interventions for patients and physicians [37].

**Methods**

We systematically searched the literature, selected eligible studies, assessed methodological quality, extracted relevant data and synthesised findings in accordance with the PRISMA-P guidelines [41]. The protocol has been registered in PROSPERO (registration number: CRD42020145823).

**Eligibility criteria**

We used the PICOS (P-Population; I- Intervention/Exposure; C-Control/Comparator; O-Outcome; S- Study design) model to frame our research questions and applied the same model to define the eligibility criteria for study selection.

**Population**

We included studies that provided information on any factors limiting or facilitating the performance of hypertension treatment and control in Nepal. These factors were associated with hypertensive patients, health care providers (physicians, nurses, healthcare workers, and others), or HS. We did not restrict studies according to participants’ age, gender, ethnicity, or comorbidity.

**Exposure/Intervention**

We defined the exposure as any factor that reportedly impedes or facilitates hypertension treatment and control in Nepal [14, 27], excluding the non-modifiable personal attributes such as age, gender and ethnicity of patients and providers. Strategies were any interventions, program or policies targeted to the providers or patients or other components of HS and delivered at the community or health facility level for improving hypertension treatment and control.

**Comparator/ control**

We included studies with or without comparators or control groups. Some of the comparators or controls were untreated, uncontrolled hypertensive individuals, general population, or patients who receive usual care.

**Outcomes**

The outcomes of interest were hypertension treatment, hypertension control and treatment adherence. By “hypertension treatment” we considered the use of at least one antihypertensive medication by hypertensive patients [14]. We defined “hypertension control” as maintaining systolic blood pressure of <140 mmHg and diastolic blood pressure of <90 mmHg in individuals under antihypertensive medication [14]. “Medication adherence” was defined as consistently taking antihypertensive medication as prescribed by the health care provider [14].

**Types of study**

We included observational and experimental studies that assessed and quantified barriers, enablers and strategies for hypertension treatment and control in Nepal. Qualitative studies that provided information on views and experiences of patients, providers and other related stakeholders regarding barriers and enablers of hypertension treatment and control were also included in the review. Studies were included regardless of language of publication or sample size. Reviews, case reports, case series and conference abstracts were not deemed eligible for inclusion.

**Search strategy**

The primary literature search was conducted in PubMed, Embase, Web of Science, CINAHL through EBSCOHost, ProQuest and WorldCat databases. We used a comprehensive search strategy to identify hypertension-related literature published between 2000 and 2019 in Nepal. Barriers, enablers and strategies for hypertension treatment and control are likely to change over time. Therefore, to provide the most relevant data for informing changes in HS, this review focused on the current situation, and we only searched for the recent literature (i.e. published in the last 19 years). We combined the search terms “hypertension” and “blood pressure” with the search term “Nepal”. The full search syntax is provided in Additional file 1.

The secondary search was done through Nepal Journals Online, Journal of Nepal Health Research Council, Journal of Kathmandu Medical College, Nepal Medical College Journal, Journal of Nepal Medical Association and Chitwan Medical College Journal to identify any relevant citations we may have missed in
the primary search. We also accessed websites of the Nepali Government, professional and regulatory organisations, and national and international agencies. Finally, we conducted forward citation searching for included studies using Google Scholar database and screened the reference lists of the included studies, as part of a backward citation search.

Data management

We used EndNote X8 (Clarivate Analytics, Philadelphia, PA, USA) and Rayyan QCRI [42] to manage the records and other data throughout the review. At first, we exported the citations to EndNote X8 to identify and remove the duplicates. The citations were then imported to Rayyan QCRI, to carry out screening and data extraction.

Selection process

Initially, two authors (RRD and ARP) independently screened the titles and abstracts of the articles and labelled them as ‘included’, ‘maybe’ and ‘excluded’. Full-texts of all records except the ones that were labelled ‘excluded’ by both reviewers were reviewed. The screening process was blinded, and all disagreements in study selection were settled by consensus between the two authors. A PRISMA flow chart illustrating the study selection process is presented in Figure 2.

Data extraction

Using a pre-designed form, two authors (RRD and ARP) independently extracted the following data from the included publications: study design; study types; study participants; study setting; statistical method; year of publication; information to assess the risk of bias; types and/or prevalence of barriers, enablers, and strategies; and the effects on hypertension awareness, treatment, and control. For strategies, we additionally extracted data related to the comparator group.

Quality assessment

We used the Mixed Methods Appraisal Tool (MMAT) to assess the methodological quality of the selected studies. MMAT includes a five-item checklist for different types of study design, with ‘Yes’ ‘No’ and ‘Can’t tell’ response options (Additional file 2). For each ‘Yes’ response, the given study was assigned a star (*). A study could, therefore, get an overall score between zero and five stars. For the purpose of this review, we used the following scoring system for overall methodological quality: 0-1 star = ‘low’; 2-3 stars = ‘fair’; 4-5 stars = ‘good’. Two authors (RRD and ARP) independently checked the quality of the studies. To resolve discrepancies between assessment scores provided by the two authors, a third author (NS) also assessed the quality of the studies using the same tools. All conflicts were resolved by consensus. We did not exclude studies from the review based on the outcome of quality assessment.

Data synthesis

Extracted qualitative data were coded and merged into themes. Themes relating to the institution, resource, finance and services were categorised as HS factors. Attitudes, perceptions, practices, behaviours and beliefs of patients and providers were listed under their respective domains. We calculated the odds ratios and their 95% confidence interval (CI) using the proportions of exposure and outcomes, where available. Due to the limited information on the exposures and outcome and large methodological heterogeneity between the selected studies, we summarised the quantitative study findings narratively.

Results

Study selection

The systematic search of six electronic databases resulted in a total of 2175 records. We removed 767 duplicates in Endnote and exported the remaining 1408 records to Rayyan QCRI for the title and abstract screening. We excluded 1302 ineligible titles and abstracts and retrieved the full-texts of the remaining 106. Out of 106 full-text articles that we assessed, 94 had either ineligible outcomes, were conducted among non-hypertensive or non-Nepali participants, or were reviews, case reports or conference abstracts. We did the forward and backward citation tracking for the remaining 12 studies and identified two additional records [43, 44]. No additional eligible studies were found in Nepalese journals and on websites of the Nepali Government, professional and regulatory organisation, and national and international agencies. Finally, we included 14 studies [43-56] for qualitative synthesis (see Fig 2). None of the studies were included in the quantitative synthesis of effect sizes because of the significant differences in their aims, methods, interventions/exposures and outcome measures.

Study characteristics

Most of the included studies were conducted in either province 3 (Kathmandu and its periphery, n=8) or province 4 (Kaski, n=4) or province 1 (Sunsari, n=2). The numbers of studies conducted in hospital (n=8) and community (n=6) settings were nearly equal. All the community-based studies included participants from the peri-urban (n=3) and urban (n=3) areas. Only a study conducted in Acham (hospital and community-based) included participants from a rural area [47]. Half of the studies were completed in 2015 or later, and more than half of the studies assessed the barriers and enablers of hypertension treatment and control [51-56]. They applied qualitative methods (n=3) [52-54], quantitative methods (n=4) [51, 56] and mixed methods (n=1) [55]. The majority of these studies investigated hypertension treatment as an outcome of interest (Table 1). Two studies also discussed adherence to antihypertensive medications [56]. Most studies collected data from hypertensive participants, except one study that interviewed community health workers to explore the barriers and enablers in utilising healthcare among the patients [54].
Six of the included studies that assessed the effectiveness of hypertension treatment and control strategies [45-50] were prospective comparative studies (n=2) [49, 50], randomised trials (n=2) [46, 48] and uncontrolled before and after studies (n=2) [45, 47]. Five out of these six studies reported systolic and diastolic blood pressure as outcome measures. The studies tested the effectiveness of health education [45-49] and combined, antihypertensive medication and yoga [48] interventions (Table 2).

The total number of participants in the included studies was 2854, with the individual sample sizes ranging from n=13 to n=1638. Only two studies applied multivariable analysis and reported adjusted effect sizes [46, 56]. Most of the qualitative and mixed methods studies (3 out of 4) applied a thematic analysis. Three studies [46, 47, 52] reported non-response rates, and they ranged between 9% and 27.8%.

**Study quality**

In quality assessment, 10 studies received three or more stars, suggesting they were of fair or good methodological quality (Additional file 2). Only two out of 11 quantitative studies [46, 56] adjusted for confounding.

**Barriers and enablers of hypertension treatment and control**

**Analysis of qualitative studies**

We identified several themes on barriers to hypertension treatment and control discussed in the qualitative studies. We categorised them into three major domains: HS excluding provider and patient (HS), providers and individual-level barriers (see Table 3). The unaffordability of health services [52], lack of human resources (primarily in rural areas) [54] and the unavailability of diagnostic tools [54] were the most often cited HS barriers.

All qualitative studies discussed the gap in communication between providers and patients that impeded medication adherence and routine follow up. Health care providers failed to deliver clear information to patients regarding medication dosage and duration, behaviour modification and need for routine monitoring. A hypertensive male participant from the study conducted by Shrestha et al. [52] complained that the doctor did not explain enough about his condition.

"... After check-up, I was told to take medicine. ...I was not told anything. So, I asked people who have heart disease to get the information regarding what food to eat, which food increases it, and which food controls pressure." [52]

A 55 years old participant with uncontrolled hypertension from a study in Kathmandu reported not visiting the doctor for one and half years since her last visit [55] as the doctor did not require for regular visits.

"I have not gone for follow-up. I am following the same regimen from last one and a half years. My doctor told me to visit him only if I had problems" [55]

In addition, other provider-related factors affecting hypertension treatment and control were: health worker's lack of interest in counselling for lifestyle modifications [52-55]; long waiting hours for the appointment [52, 55]; lack of national guidelines in hypertension treatment [55]; and provider's negative behaviours [52].

A wide variety of themes evolved from the qualitative analysis of the findings on patient's beliefs and practices that affect hypertension treatment and control (table 3). The factors impeding initiation of anti-hypertensive treatment were poor help-seeking behaviours [54] and reluctance to take medication due to perceived side effects and fear of long-term use [52, 55]. Other factors related mostly to treatment adherence and lack of routine monitoring of patients [54, 55].

Three studies discussed facilitating factors for improved hypertension treatment. These factors were: access to hypertension care [54], treatment adherence [52], patient's attitude [52] and family support [52, 53].

**Analysis of quantitative studies**

Five studies reported findings based on quantitative data. The studies identified the factors associated with hypertension treatment, adherence and control (Table 4). Affordability of the medication was associated with treatment adherence [56]. At the level of the provider, prescribing more than one type of pill was associated with greater non-adherence [56]. Similarly, long waiting hours (>20 minutes vs ≤ 20 minutes) and lack of proper counselling from health workers were associated with uncontrolled hypertension [55]. All quantitative studies reported various individual (patient) level factors that were significantly associated with untreated or uncontrolled hypertension or non-adherence to medication. These were: patients' lack of awareness on normal blood pressure target [51, 55]; lack of knowledge about possible complications of hypertension [51]; lost to follow up [55, 56]; non-adherence to medication [51, 55]; and lack of blood pressure monitoring [51].

Only one study discussed enablers separately. The study found that scores of dimensions of illness perception particularly timeline, treatment control and coherence were positively correlated with medication adherence [44]. The higher the participant perceived high blood pressure as a chronic condition (Spearman correlation coefficient (r)=0.23, p<0.05), the better was medication adherence. The more the participant believed that treatment can control the blood pressure (r=0.51, p<0.05), the higher the medication adherence score they had. The better the understanding of hypertension (coherence, r=0.22, p<0.05), the higher was the medication adherence [44].

**Strategies for hypertension treatment and control**
Out of six eligible studies in this category, all studies intervened at the patient level, two studies also investigated the effect of workforce strengthening (training and continuing education for integrated of NCD care) and one study evaluated the impact of new data recording systems at the healthcare centre (Table 5). The most frequently tested antihypertensive strategy was health education for hypertensive patients [45-47, 49], and three out of four studies found it effective in reducing blood pressure. Health education was applied as a component of comprehensive blood pressure management strategies [46, 47, 49] or as a single intervention [45]. Two studies [49, 50] also compared hydrochlorothiazide, enalapril and amlodipine as first-line antihypertensive drugs. The reduction in mean systolic and diastolic pressure was significantly higher with enalapril and amlodipine, compared with hydrochlorothiazide [49, 50]. The post-intervention changes in blood pressure did not vary significantly between enalapril and amlodipine groups after for weeks [49, 50]. The participant under amlodipine reported more adverse events such as peripheral oedema, shortness of breath and headache than the enalapril group.

One study studied yoga as an adjuvant therapy to medication and compared it with usual care [48]. The yoga group had a significant reduction in both systolic and diastolic blood pressure, compared with the 'medication-only' group after six weeks of follow-up.

One of the two studies that intervened at the level of healthcare providers found a significant improvement in blood pressure amongst their clients. The study trained and involved female community health volunteers for health education, blood pressure monitoring and referral of hypertensive cases. The intervention was effective in reducing blood pressure. The mixed-effect regression coefficient was −4.9 (95% CI: −7.8 to −2.0) for systolic blood pressure and −2.6 (95% CI: −4.6 to −0.7) for diastolic blood pressure [46]. One study implemented the intervention program at HS level, by establishing a new digital and electronic health record system along with capacity development for healthcare workers and providing health education to the patients. The intervention had no statistically significant effect on hypertension control [47].

**Discussion**

We systematically reviewed 14 studies to identify barriers, enablers and strategies of hypertension treatment and control in Nepal. Eight of them investigated various barriers and enablers within three domains: the HS, health care providers, and patients. The remaining six studies investigated the effectiveness of different hypertension treatment and control strategies, including health education and yoga across three domains.

The synthesis of the findings reported in qualitative studies generated two themes on HS-related barriers for hypertension control and treatment. They were the lack of affordable health care and HS resources including human and physical assets. Barriers relating to the availability and affordability of hypertension care services are common in low-income countries [14, 16]. The HS factors are complex and can have a strong influence on health service delivered by the provider and patient's access to care [14]. The unaffordability of the services and financial hardship at the individual level had linkages with poor help-seeking behaviours, non-adherence to medication and an irregular follow-up visit to health centres among hypertensive patients. The result from the quantitative analysis also showed a significant association between high cost of drugs and non-adherence to medication. The odds of not adhering to taking antihypertensive drugs were 5.2 times higher among the patients who reported the cost of medicine as expensive [56]. This study identified the provision of free essential medicines at government health facilities as a single facilitating factor relating to the Nepali HS. Providing free essential antihypertensive medicines at the primary health care centres was supposed to increase access to treatment of hypertension. Some of the calcium channel blockers and beta blockers are freely available at primary health care centres. However, regular, effective and efficient supply of these medicines is uncertain [57]. For example, most of the health facilities (HF) reported unavailability of atenolol (82% of HF) and amlodipine (90% of HF) in 2015 [58].

None of the strategies from the selected studies aimed either to minimise the financial burden associated with hypertension treatment or to increase access to hypertension care. Only one study was identified to report the effect of a strategy implemented at the level of the HS through establishing a new hospital-based health record system, but the strategy failed to show a significant impact on blood pressure control. To increase access to care and minimise excessive health expenditures, different HS financing approaches, including health insurance coverage and co-payments for medication, have been evaluated globally [14]. Nepal recently enacted the National Health Insurance Act 2017 and is moving towards universal health coverage for equitable access to high quality and affordable health care [59]. However, its overall impact on treatment and control of hypertension and its related complications is yet to be studied.

The communication gap between health workers and the patient was one of the most frequently studied barriers associated at provider level. The lack of clear messages on medication, dosage, lifestyle modification, and follow-up visits was reported as causes of non-adherence to medication and irregular follow-up visits, ultimately leading to suboptimal control of blood pressure. A recent study found that less medication (antihypertensive drugs) discussion between providers and patients was associated with six-fold increased odds of poor adherence to medication [60]. Jolles et al. also emphasised that effective communication between providers and patients could increase medication adherence and contributes to better control of high blood pressure [61].

Capacity development of the female health community volunteers (FCHV) and involving them for health education, blood pressure monitoring and linking the cases to health services was the only effective intervention implemented at the provider level. FCHV's involvement in screening the high-risk population and monitoring the hypertensive patients is cost-effective [62] and could minimise the burden of exhausted healthcare workers. After a very acclaimed performance on improving maternal and child health [63], new evidence suggests that community health workers could potentially work as the frontline cadres to fight against the burden of non-communicable diseases, including hypertension [64]. Task sharing, particularly with other non-physician health workers in understaffed and resource-poor settings, is an effective strategy for the management of hypertension [31]. Additionally, other approaches such as including pharmacists and nurses in a team and providing team-based hypertension care also showed positive effects on controlling high blood pressure [32].

Most of the barriers at the patient level were related to their knowledge, beliefs and practices. Being unaware of the normal blood pressure target, perceived side effects of drugs, fear of long-term use of medication, reluctance to take medicine, self-medication, non-adherence to medication, lack of follow-up visits, lack of blood pressure monitoring and not minimising risky health behaviours were the key barriers to hypertension treatment and control. Similar to our findings, Khatib et al. in their systematic review reported several patient-related barriers to hypertension treatment and control [16]. The authors categorised
them as capability barriers (e.g. lack of skills and knowledge), intention and motivation barriers (e.g. lack of motivation to adhere to a treatment, false belief, such as drug dependency) and medication barriers (e.g. self-perceived side effects, taste and dosage), among others [16].

Health education was the most commonly investigated strategy. One study used yoga as an adjuvant therapy to the medication and found that yoga was effective in blood pressure reduction. Globally, barriers relating to patient's knowledge, belief and practices have been tackled effectively using different approaches, such as simplifying the treatment using polypill [35] and increasing adherence to medication, routinely and conveniently monitoring hypertension through home-based blood pressure measurement [36] and providing education interventions to patients and physicians [37]. Future studies should evaluate the effectiveness of these strategies in the context of Nepali healthcare system.

Some limitations of this review stem from the characteristics of the included studies. Although the majority of studies scored three out of five in MMAT checklists, nearly all of them were subject to inadequate reporting of research methods. For example, out of six comparative or intervention studies, three studies did not present clear information on the estimation of the required sample size and participant selection. Furthermore, only two quantitative studies adjusted their analyses for confounding. These methodological limitations of the included studies may have affected their findings and consequently the findings of our review.

A meta-analysis could not be performed due to a large methodological heterogeneity and variation in outcome measures between the included studies. For example, the data on barriers and enablers were extracted from qualitative, quantitative and mixed methods studies. The data on antihypertensive strategies were from prospective comparative studies, randomised trials and uncontrolled before and after studies. The intervention modalities also varied across studies from health education to yoga.

**Conclusion And Implications**

This is the first study to systematically investigate barriers, enablers and strategies of hypertension treatment and control in Nepal. The study findings suggest that various factors associated with HS, providers and patients are responsible for poor hypertension care in Nepal. The findings of our review may assist policy makers and other public health stakeholders in designing future interventions for addressing barriers to hypertension treatment and control.

One example is promoting health education through female community health volunteers (FCHV) that has shown a positive effect in reducing blood pressure. However, more tailored and theory-driven research on the effectiveness of health education interventions is required to confirm the finding. Similarly, trial data available in Nepal are not sufficient to conclude that yoga is an effective antihypertensive therapy. Further large-scale studies are required to evaluate the effectiveness of yoga in hypertension treatment and control.

**Abbreviations**

<table>
<thead>
<tr>
<th>Abbreviation</th>
<th>Description</th>
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<tr>
<td>FCHV</td>
<td>Female health community volunteer</td>
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<tr>
<td>HF</td>
<td>Health facility</td>
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<tr>
<td>HS</td>
<td>Health system (without provider and patient)</td>
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<td>LMICs</td>
<td>Low- and middle-income countries</td>
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<td>MMAT</td>
<td>Mixed Methods Appraisal Tool</td>
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<td>NCDs</td>
<td>Non-communicable diseases</td>
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</table>

**Declarations**

**Availability of data and materials**

All data generated or analysed during this study are included in this published article and its additional files.

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**Authors’ Contributions**

RRD developed the study protocol, screened the records, extracted the data, assessed the quality of included studies, interpreted the findings and prepared the first draft of the manuscript. ARP screened the records, extracted the data, assessed the quality of included studies, and assisted in interpreting the findings. NS assisted in assessing the quality of the selected studies. ZP and MdC reviewed the study protocol. All authors contributed to writing the manuscript and its revisions. All authors read and approved the final manuscript.

**Ethics declarations**
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Tables

Table 1. Characteristics of the studies on barriers and enablers of hypertension treatment and control in Nepal
<table>
<thead>
<tr>
<th>Sn</th>
<th>Study ID</th>
<th>Study types</th>
<th>Sampling method</th>
<th>Data collection method</th>
<th>Participants</th>
<th>Sample size total (m/f)</th>
<th>Survey year</th>
<th>Survey site</th>
<th>Area</th>
<th>Outcome</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>Bhandari et al. (2015)</td>
<td>Cross-sectional study</td>
<td>Random sampling (154 from 975 hypertensive participants)</td>
<td>Semi-structured interview</td>
<td>Hypertensive persons of age 35 years and above</td>
<td>154 (NA/NA)</td>
<td>September to February 2010</td>
<td>Dharan sub-metropolitan city</td>
<td>Urban</td>
<td>Adherence to medication, hypertension control</td>
<td>Sex, education, occupation, family history of hypertension were adjusted. No information on gender distribution</td>
</tr>
<tr>
<td>2</td>
<td>Devkota et al. (2016)</td>
<td>Cross-sectional, mixed-methods (Quantitative and Qualitative) study</td>
<td>Stratified random sampling for the quantitative part</td>
<td>Structured interview</td>
<td>Hypertensive persons of age between 18 years and 70 years</td>
<td>118 (54/64)</td>
<td>January and July 2015</td>
<td>Kathmandu, Nepal</td>
<td>Peri-urban</td>
<td>Hypertension awareness, hypertension treatment,</td>
<td>No adjustment of confounding in analysis</td>
</tr>
<tr>
<td>3</td>
<td>Khan et al. (2013)</td>
<td>Cross-sectional quantitative study</td>
<td>Not clear</td>
<td>Structured interview</td>
<td>Treated hypertensive persons</td>
<td>79 (44/35)</td>
<td>2012</td>
<td>Pokhara</td>
<td>Urban</td>
<td>Hypertension control</td>
<td>No clear explanation on sampling procedures, no adjustment of confounding in analysis</td>
</tr>
<tr>
<td>4</td>
<td>Khanal et al. (2017)</td>
<td>Cross-sectional, qualitative study</td>
<td>Purposive sampling</td>
<td>In-depth interview</td>
<td>Health care providers (health assistants, pharmacy assistants, nurse, specialised nurse, practicing pharmacists, chief hospital pharmacist, doctors, and specialised doctor)</td>
<td>9 (4/5)</td>
<td>March 2015</td>
<td>Loknath Municipality, Kaski</td>
<td>N/A</td>
<td>Hypertension treatment</td>
<td>Small sample size, participants are not the hypertensive patients</td>
</tr>
<tr>
<td>5</td>
<td>Maharjan S (2016)</td>
<td>Cross-sectional, quantitative study</td>
<td>Purposive sampling</td>
<td>Structured interview</td>
<td>Hypertensive patients of age 18 years and above and who were on treatment</td>
<td>85 (53/32)</td>
<td>January and March 2016</td>
<td>Sahid Gangralal National Heart Center</td>
<td>Urban</td>
<td>Treatment adherence</td>
<td>No adjustment of confounding in analysis</td>
</tr>
<tr>
<td>6</td>
<td>Oli et al. (2014)</td>
<td>Cross-sectional, qualitative study</td>
<td>Non-probability sampling</td>
<td>In-depth interviews</td>
<td>Patients with confirmed hypertension, diabetes, heart disease for at least one year; Age 20 years and older</td>
<td>13 (6/7)</td>
<td>May and July 2013</td>
<td>Bhaktapur</td>
<td>Peri-urban</td>
<td>Hypertension treatment</td>
<td>Participants belonged to non-hypertensive patient as well</td>
</tr>
<tr>
<td>7</td>
<td>Shrestha et al. (2018)</td>
<td>Cross-sectional, qualitative study</td>
<td>Random sampling</td>
<td>Focus group discussion</td>
<td>Newly diagnosed hypertensive patients of age 18 years and above</td>
<td>35 (26/9)</td>
<td>Dec 2015 to Feb 2016</td>
<td>Dhulikhel Municipality</td>
<td>Peri-urban</td>
<td>Hypertension awareness, hypertension treatment, hypertension control</td>
<td>Non-response rate is 27.8</td>
</tr>
<tr>
<td>8</td>
<td>Simkhada R (2012)</td>
<td>Cross-sectional study</td>
<td>Random sampling</td>
<td>Structured interview, clinical measurements</td>
<td>Hypertensive participants diagnosed and under medication at least from last 6 months</td>
<td>147 (83/64)</td>
<td>1 Feb 2011-31 July 2011</td>
<td>Bir Hospital, Kathmandu, Nepal</td>
<td>NR</td>
<td>Hypertension control</td>
<td>Small sample size; no age and sex-adjusted analysis</td>
</tr>
</tbody>
</table>

Table 2: Characteristics of the studies on the effectiveness of strategies to improve hypertension treatment and control in Nepal
<table>
<thead>
<tr>
<th>Study ID</th>
<th>Study types</th>
<th>Study date and sites</th>
<th>Participants</th>
<th>Participant recruitment</th>
<th>Intervention</th>
<th>Sample size total (treatment/control)</th>
<th>Outcome</th>
<th>MMAT score</th>
</tr>
</thead>
<tbody>
<tr>
<td>Alam et al. (2007)</td>
<td>Prospective comparative study</td>
<td>Sep 2005 to July 2006; Kathmandu University Teaching Hospital; Dhiulikhel and Saheed Gangalal National Heart Center</td>
<td>Hypertensive patients diagnosed newly or not taking medication, of any age</td>
<td>Participants were selected and divided them into two groups: amlodipine and enalapril group</td>
<td>Amlodipine or enalapril for four weeks</td>
<td>72 (36/36)</td>
<td>Systolic blood pressure</td>
<td>***</td>
</tr>
<tr>
<td>Humagain et al. (2015)</td>
<td>Prospective comparative study</td>
<td>January 2015 to April 2015; Dhiulikhel Hospital, Kathmandu University Nepal</td>
<td>Hypertensive patients of 40-80 years of age attending cardiology OPD</td>
<td>Enrolled all the hypertensive patients who attended the study site during the study period and divided them into three groups based on the three antihypertensive medication they used: hydrochlorothiazide 25 mg or amlodipine 5 mg or enalapril 5 mg</td>
<td>Participants received an antihypertensive medication (any of hydrochlorothiazide 25 mg or Amlodipine 5 mg or Enalapril 5 mg) and health education.</td>
<td>172 (NA/NA)</td>
<td>Systolic blood pressure</td>
<td>***</td>
</tr>
<tr>
<td>Khadka et al. (2010)</td>
<td>Randomised controlled trial</td>
<td>2000-2002; Yoga and lifestyle clinics, Department of Physiology, BPKIHS</td>
<td>Essential hypertensive patients of 30-60 years of age, without diabetes</td>
<td>Purposively selected the participants and Systematically randomised them into yoga and control group</td>
<td>Treatment group participants underwent for yoga practice scheduled for half an hour at least for 6 days in a week for 6 weeks; yogic practices were strengthening exercise (5 min), yogic asanas (6-7 minutes), Shavasan (5 minutes), meditation (5 minutes), pranayam (4 minutes)</td>
<td>14 (7/7)</td>
<td>Systolic blood pressure</td>
<td>***</td>
</tr>
<tr>
<td>Kumar et al. (2019)</td>
<td>Uncontrolled Before and After study</td>
<td>1 December 2010–31 May 2018; Achham, Nepal</td>
<td>Hypertensive patients of age 18 years and above</td>
<td>Among the 488 hypertensive patients enrolled in the NCD program, the study selected 86% (340 out of 408) patients (all with at least two follow up visits and having at least 12 week interval between baseline and endline</td>
<td>Intervention includes NCD management capacity development for healthcare workers, establishment of digital and electronic health records in the centre and individual (patient) level counselling for preventive and treatment of NCDs.</td>
<td>340 (340/-)</td>
<td>Hypertension control</td>
<td>***</td>
</tr>
<tr>
<td>Neupane et al. (2018)</td>
<td>Clustered randomised trial</td>
<td>2015-2016; Lekhanath municipality, Kaski</td>
<td>General population (normotensive, pre-hypertensive, hypertensive) of 25-65 years of age</td>
<td>Participants were selected from 14 clusters. The municipality was divided into 15 clusters out of which 14 clustered were randomised into intervention and control group, 1:1 ratio.</td>
<td>The treatment group participants received health education on major NCD risk factors, had their blood pressure measurement and were referred (only for hypertensive participants) to health centre by the trained Female Community Health Volunteer in every four months/year.</td>
<td>1638 (939/699)</td>
<td>Systolic blood pressure</td>
<td>****</td>
</tr>
<tr>
<td>Sharma et al. (2014)</td>
<td>Uncontrolled Before and After study</td>
<td>August 2012 to April 2013; Sankalpa Pharmacy, Kaski, Nepal</td>
<td>Hypertensive patients of 31 years and above attending the pharmacist run hypertension clinic</td>
<td>Enrolled all the eligible hypertensive patients who attended the study site</td>
<td>Healthcare workers (pharmacist) led one-to-one counselling - contained three counselling session at 0 date, two months and four months. The contents of counselling were lifestyle modification, treatment, and control of hypertension</td>
<td>50 (50/-)</td>
<td>Systolic blood pressure</td>
<td>***</td>
</tr>
</tbody>
</table>

Table 3. Barriers and enablers of hypertension treatment and control in Nepal
Table 4. Barriers reported in quantitative studies

<table>
<thead>
<tr>
<th>Study ID</th>
<th>Domain</th>
<th>Findings</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bhandari et al. (2015)</td>
<td>Health system related</td>
<td>Exposure: high cost of medicine (expensive vs. affordable)</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: non-adherence</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 5.2; 95% CI: 1.1-23.9</td>
</tr>
<tr>
<td></td>
<td>Provider related</td>
<td>Exposure: multi-pills (more than one vs. one antihypertensive drug prescription)</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: non-adherence</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 5.3; 95% CI: 1.2-23.7</td>
</tr>
<tr>
<td></td>
<td>Patient related</td>
<td>Exposure: lost to follow up (irregular vs regular)</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: non-adherence</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 6.4; 95% CI: 1.2-33.4</td>
</tr>
<tr>
<td>Devkota et al. (2016)</td>
<td>Health system related</td>
<td>Exposure: long waiting hours (&gt;20 minutes vs. ≤ 20 minutes)</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 2.7; 95% CI: 1.1-6.8</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Exposure: no health education from health worker</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 2.6, 95% CI 1.2-6.4</td>
</tr>
<tr>
<td></td>
<td>Patient related</td>
<td>Exposure: lack of awareness of blood pressure target</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: untreated hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 3.2, 95% CI: 1.3-8.2</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Exposure: nonadherence to medication</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 8.8, 95% CI: 3.3-23.4</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Exposure: lost to follow-up</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 3.3, 95% CI: 1.2-9.1</td>
</tr>
<tr>
<td>Simkhad (2012)</td>
<td>Health system related</td>
<td>Exposure: lack of awareness on blood pressure target</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 2.8, 95% CI: 1.3-6.1</td>
</tr>
<tr>
<td></td>
<td>Provider related</td>
<td>Exposure: lack of awareness on BP complication</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 3.7, 95% CI: 1.5-9.1</td>
</tr>
<tr>
<td></td>
<td>Patient related</td>
<td>Exposure: nonadherence to medication</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 3.3, 95% CI: 1.5-7.1</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Exposure: lack of blood pressure monitoring</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Outcome: uncontrolled hypertension</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Odds ratio = 2.9, 95% CI: 1.1-8.1</td>
</tr>
</tbody>
</table>

Table 5. Findings of the studies on the effectiveness of strategies to improve hypertension treatment and control in Nepal
<table>
<thead>
<tr>
<th>Study ID</th>
<th>Comparison</th>
<th>Sample size total (treatment/control)</th>
<th>Participants in analysis</th>
<th>Outcome 1: Systolic blood pressure</th>
<th>Findings</th>
<th>Overall MMAT score</th>
</tr>
</thead>
<tbody>
<tr>
<td>Alam et al. (2007)</td>
<td>Enalapril vs amlodipine</td>
<td>72 (36/36)</td>
<td>72</td>
<td>mean (sd) reduced from 162.1 (22.0) mmHg to 131.3 (11.4) mmHg in enalapril group and from 158.2 (16.2) mmHg to 132.6 (11.7) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td>Humagain et al. (2015)</td>
<td>Comparison 1: hydrochlorothiazide 25 mg vs amlodipine 5 mg</td>
<td>172 (NA/NA)</td>
<td>172</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Comparison 2: hydrochlorothiazide 25 mg vs enalapril 5 mg</td>
<td>172 (NA/NA)</td>
<td>172</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td></td>
<td>Comparison 3: amlodipine 5 mg vs enalapril 5 mg</td>
<td>172 (NA/NA)</td>
<td>172</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td>Khadka et al. (2010)</td>
<td>yoga vs routine care</td>
<td>14 (7/7)</td>
<td>14</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td>Kumar et al. (2019)</td>
<td>multilevel intervention, no comparison group</td>
<td>340 (340/-)</td>
<td>340</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td>Neupane et al. (2018)</td>
<td>Health education, blood pressure monitoring, and referral vs usual care</td>
<td>1638 (939/699)</td>
<td>1468</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
<tr>
<td>Sharma et al. (2014)</td>
<td>Health education, no comparison group</td>
<td>50 (50/-)</td>
<td>50</td>
<td>mean (sd) reduced from 103.1 (11.5) mmHg to 85.2 (8.1) mmHg in enalapril group and from 100.3 (7.3) mmHg to 87.3 (8.4) mmHg in amlodipine group</td>
<td>***</td>
<td></td>
</tr>
</tbody>
</table>

Additional Files

Additional file 1:
Search strategies. (DOCX 88 KB)

Additional file 2:
Quality assessment of the selected studies. (xlsx 13 KB)

Figures
2175 records were identified through database searching
CINAHL (n = 331)
EMBASE (n = 1052)
ProQuest (n=42)
PubMed (n=467)
Web of Sciences (n=275)
WorldCat (n=8)

767 duplicates were removed

Records screened (n = 1408)

Records excluded through title and abstract screening (n=1302)

94 Full-text articles excluded
- Non-relevant outcome (n = 71)
- Non-specific study population (n=9)
- Review and case report (n=4)
- Conference abstracts (n=10)

Full-text articles assessed for eligibility (n = 106)

Studies included in primary search (n = 12)

Studies included in qualitative synthesis (n = 14)

2 records were identified through backward and forward citation search

Figure 1
Study selection flow diagram
Figure 2

Conceptual framework of barriers, enablers, and strategies for hypertension control

Supplementary Files

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- PRISMA2009checklist.doc
- Additionalfile1.Searchstrategy.docx
- Additionalfile2.Qualityassessment.xlsx