## Proceedings

# Prescribing Pattern of Anti-Hypertensive Drugs and Adherence to JNC VII Guideline ${ }^{+}$ 

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Published: 14 November 2018


#### Abstract

Introduction: Hypertension is an important public health concern because of its associated morbidity, mortality and economic impact on society. It is a significant risk factor for cardiovascular, cerebrovascular and renal complications. A number of national and international guidelines for the management of hypertension have been published. The Joint National Committee (JNC) in 2003 published a series of guidelines to recommend the appropriate antihypertensive therapy based on the best available evidence. Objectives: This drug utilization study was intended to find out the preferred drug group prescribed either alone or in combinations and their adherence to the JNC 7 guidelines. Materials and Methods: This was a prospective cross-sectional study. Drug utilization data on 100 hypertensive patients were collected from various hospitals in Nepal. The patients who received antihypertensive drugs during their treatment period in SPSS V. 16.0. The prescribed drugs were compared with JNC 7 guidelines. Results: It was found that $40 \%$ of patients were male and $60 \%$ were female. The largest subset of female hypertensive patients ( $45.0 \%$ ) were in the age group of $>60$ years and a plurality of male hypertensive patients $(45.0 \%)$ were in the age group of $40-60$ years. It was found that $45 \%$ of the patients had Stage 1 hypertension, $32 \%$ of the patients were in prehypertension, $17 \%$ of the patients had Stage 2 hypertension. The most frequently prescribed antihypertensive drug regimens were angiotensin receptor blockers (ARBs) ( $32.44 \%$ ), ARB + thiazide (15.94\%), diuretics (11.59\%), calcium channel blockers (CCBs) + beta blockers ( $9.42 \%$ ) and CCBs $(8.7 \%)$. Thirty-nine percent received monotherapy while the remaining $61 \%$ received combination therapy. Seventy-four percent of the total prescriptions followed JNC 7 guidelines. Conclusion: There is a need to follow official guidelines in managing hypertension as a chronic disease, since these guidelines are based on various clinical trials, and the successful attainment of a target BP in patients will be made much easier by implementing them. National health policymakers should consider the evaluation and treatment of hypertension as a right in the public health system for better outcomes in terms of morbidity and mortality from hypertension.


Keywords: HTN; JNC 7 guidelines; anti-hypertensive; drug utilization; ARBs

## 1. Introduction

Hypertension (HTN) is the most common cardiovascular disorder and is regarded as major public health problem [1]. HTN is not a cause of death by itself, but it is an important risk factor for cardiovascular mortality and morbidity. HTN remains one of the most important preventable contributors [2]. HTN is the single most important cause of preventable mortality. It is a key contributing factor in the development of cardiovascular and cerebrovascular disease, and a major
cause of stroke, myocardial infarction, heart failure and kidney disease, among others. It leads to death if not detected early and treated appropriately.

Antihypertensive drug therapy has improved remarkably in the last 56 years. The first antihypertensive agent which was developed in 1958 was chlorothiazide, the first thiazide diuretic used in treatment of HTN, which was developed from the antibiotic sulfanilamide. After the grand success of chlorothiazide in the treatment of HTN, soon more agents were developed, such as beta blockers, calcium channel blockers, angiotensin-converting enzymes, angiotensin receptor blockers and renin inhibitors $[3,4]$. Due to the increase in the number of drugs for the treatment of HTN, different complications and problems have been arising.

To counteract the problem, 39 major professional, public and voluntary organizations and seven federal agencies formed a committee known as the Joint National Committee (JNC) [5].

## The Key Messages of the JNC 7 Report

In persons older than 50 years, systolic blood pressure (BP) greater than 140 mmHg is a greater risk factor for HTN. Cardiovascular disease (CVD) risk doubles for each increment of 20/10 mmHg. Those who are normotensive at 55 years of age have a $90 \%$ lifetime risk of developing hypertension. Prehypertension, defined as a systolic BP of $120-139 \mathrm{mmHg}$ or a diastolic BP of $80-89 \mathrm{mmHg}$, requires lifestyle modifications to prevent the progressive rise in blood pressure and CVD. Hypertension will be controlled only if patients are motivated to stay on their treatment plan. According to JNC 7 guidelines, the measured blood pressure is classified by the below blood pressure classification [5].

In JNC7 guideline blood pressure is classified and simplified to four categories, which are given below (Table 1).

Table 1. Contemporary blood pressure classification guidelines. DBP: diastolic blood pressure; SBP: systolic blood pressure.

| Subset | JNC 7 Category | SBP/DBP |
| :---: | :---: | :---: |
| 1. | Normal | $<120 / 80$ |
| 2. | Prehypertension | $120-139 / 80-89$ |
| 3. | Hypertension Stage 1 | $140-159 / 90-99$ |
| 4. | Hypertension Stage 2 | $\geq 160 / \geq 100$ |

## 2. Methodology

### 2.1. Study Site

The study was carried out in Lalitpur Heart Clinic, Jawalakhel Nepal Heart Care Centre, Jhamsikhel and Kritipur Community Health Centre (Janapoly clinic), Kritipur and Grande International Hospital, Dhapasi.

### 2.2. Study Period

Data collection was carried out for 2 months over June-July 2014.

### 2.3. Study Design

This study was designed as a prospective cross-sectional analysis.

### 2.4. Study Population

The study included all willing patients who visited the cardiac care center or the hospital of Kathmandu Valley for the treatment of hypertension and who were positively diagnosed.

### 2.5. Inclusion Criteria

- The patients who visited in the center and diagnosed with hypertension and taking antihypertensive medicines.
- Hypertensive patients who agreed to share their information and responded to the questionnaire.


### 2.6. Exclusion Criteria

- The patients who visited the center and who were not diagnosed with hypertension.


### 2.7. Sample Size

The study included a total of 100 patients from four treatment centers: 26 from the Lalitpur Heart Clinic, 23 from the Nepal Heart Care Centre, 37 from the Janapoly clinic and 14 from the Grande International Hospital.

### 2.8. Data Collection Tools and Techniques

Data collection forms were filled from all the patients who gave consent for research. The required information was obtained from the medical records of the patients, such as their names, ages, genders, addresses, blood pressures, weights, prescriber names, prescriber offices, the antihypertensive drugs used, any associated diseases and medications for those associated diseases.

### 2.9. Statistical Tools

All the data collected from different centers were coded as per variables and data entry was done in SPSS V 16.0 data sheet and analyzed by the help of statistical software SPSS V 16.0. The analyzed data was expressed in percentage.

## 3. Results

Our data shows that 40 patients (40\%) involved were males and 60 ( $60 \%$ ) were females, indicating a $20 \%$ higher prevalence of hypertension in the female population.

From Figure 1 it can be seen that the plurality of female hypertension patients ( $45.0 \%$ ) were in the age group of $>60$ years and the plurality of male hypertension patients ( $45.0 \%$ ) were in the age group of 40-60 years, indicating an earlier onset of hypertension in male population in our study group. The proportions of female and male hypertension patients in the age group of $<40$ years were $11.7 \%$ and $17.5 \%$ respectively, which was the smallest age group.


Figure 1. Age distribution of patients.
Table 2 shows that the prevalence of Stage 1 hypertension over other stages of hypertension was higher. Among 100 patients $45 \%$ of the patients had Stage 1 hypertension, $32 \%$ of the patients had
prehypertension, $17 \%$ of the patients had Stage 2 hypertension and the proportion in the normal stage was found to be $6 \%$.

Table 2. Stages of hypertension.

| Subset | Stage of Hypertension | Proportion (\%) |
| :---: | :---: | :---: |
| 1. | Prehypertension | 32 |
| 2. | Stage 1 hypertension | 45 |
| 3. | Stage 2 hypertension | 17 |
| 4. | Normal | 6 |

In Figure 2 it can be seen that $38 \%$ and $7 \%$ of male and female patients, respectively, were found to have diabetes mellitus. A high risk of coronary heart disease was found in $13 \%$ of female patients and in $18 \%$ of male patients. Similarly, asthma was found in $5 \%$ of male patients and in $9 \%$ of female patients. Diabetes mellitus comorbid with a high risk of coronary heart disease was found at an incidence of $4 \%$ in both male and female patients and diabetes mellitus together with chronic kidney disease was found in $2 \%$ of female patients.


Figure 2. Comorbidities of the patients.
Table 3 shows that angiotensin receptor blockers alone was the most commonly prescribed regimen of antihypertensive drugs in $30.44 \%$ patients, followed by angiotensin receptor blockers (ARBs) + thiazide in $15.95 \%$, calcium channel blockers (CCBs) alone in $12.32 \%$, CCBs + beta blockers in $9.42 \%$, beta blockers in $8.7 \%$, thiazide diuretics and angiotensin-converting enzyme (ACE) inhibitors in $6.52 \%$ respectively, ARB + CCB in $3.62 \%$, diuretic combinations in $1.45 \%$, and alpha blockers and beta blockers + thiazide in $0.72 \%$ patients, respectively.

Table 4 shows that ARBs were the most commonly prescribed group of antihypertensive drugs in $42.84 \%$ of non-hypertensive patients by physicians' orders, followed by CCBs, beta blockers, diuretic combinations and ARB + thiazide group, each used in $14.29 \%$ of patients with normal blood pressure.

Table 5 shows that angiotensin receptor blockers were the most commonly prescribed group of antihypertensive drugs in $28.21 \%$ of prehypertensive patients, followed by calcium channel blockers in $20.51 \%$, thiazide diuretics and ACE inhibitors each in $15.39 \%$, ARBs + thiazide in $7.69 \%$, beta blockers in $5.13 \%$ and others in $2.56 \%$.

Table 3. Percentage of patients administered each antihypertensive drug regimen.

| Subset | Antihypertensive Medicine | Proportion of Patients (\%) |
| :---: | :---: | :---: |
| 1. | Thiazide diuretics | 6.52 |
| 2. | Angiotensin-converting enzyme inhibitors (ACEs) | 6.52 |
| 3. | Angiotensin receptor blockers (ARB) | 30.44 |
| 4. | Calcium channel blockers (CCBs) | 12.32 |
| 5. | Beta blockers | 8.7 |
| 6. | ARB + thiazide | 15.95 |
| 7. | ARB + CCB | 3.62 |
| 8. | CCB + beta blockers | 9.42 |
| 9. | Alpha blockers | 0.72 |
| 10. | High-ceiling diuretics | 3.62 |
| 11. | Beta blockers + thiazide | 0.72 |
| 12. | $H i g h ~ C e i l i n g ~ d i u r e t i c s+~ P o t a s s i u m ~ S p a r i n g ~ d i u r e t i c s ~$ | 1.45 |

Table 4. Percentage of normotensive patients prescribed each drug regimen.

| Subset | Antihypertensive Medicine | Proportion of Patients (\%) |
| :---: | :---: | :---: |
| 1. | Angiotensin receptor blockers | 42.84 |
| 2. | Calcium channel blockers | 14.29 |
| 3. | Beta Blockers | 14.29 |
| 4. | ARB + thiazide | 14.29 |
| 6. | High-ceiling + Potassium-sparing diuretics | 14.29 |

Table 5. Percentage of prehypertensive patients prescribed each drug regimen.

| Subset | Antihypertensive Medicine | Proportion of Patients (\%) |
| :---: | :---: | :---: |
| 1. | Thiazide diuretics | 15.39 |
| 2. | Angiotensin-converting enzyme inhibitors | 15.39 |
| 3. | Angiotensin receptors blockers | 28.21 |
| 4. | Calcium channel blockers | 20.51 |
| 5. | Beta blockers | 5.13 |
| 6. | ARB + thiazide | 7.69 |
| 7. | ARB + CCB | 2.56 |
| 8. | CCB + beta blockers | 2.56 |
| 9. | High-ceiling + Potassium-sparing diuretics | 2.56 |

Table 6 shows that angiotensin receptor blockers were the most commonly prescribed group of antihypertensive drugs in $28.12 \%$ of Stage 1 patients, followed by ARB + thiazide and CCB + beta blocker each in $18.75 \%$, beta blockers alone in $10.94 \%$ and calcium channel blockers alone in $7.81 \%$. It was found that only $3.13 \%$ patients were prescribed thiazide only

Table 6. Percentage of Stage 1 patients prescribed each drug regimen.

| Subset | Antihypertensive Medicine | Proportion of Patients (\%) |
| :---: | :---: | :---: |
| 1. | Thiazide diuretics | 3.13 |
| 2. | Angiotensin-converting enzyme inhibitors | 4.69 |
| 3. | Angiotensin receptor blockers | 28.12 |
| 4. | Calcium channel blockers | 7.81 |
| 5. | Beta blockers | 10.94 |
| 6. | ARB + thiazide | 18.75 |
| 7. | ARB + CCB | 13.13 |
| 8. | CCB + beta blockers | 18.75 |
| 9. | Alpha blockers | 1.56 |
| 10. | High-ceiling diuretics | 1.56 |
| 11. | Beta blockers + thiazide | 1.56 |

The largest proportions of Stage 1 patients prescribed a single drug were those taking ARBs at $28.12 \%$ and those taking beta blockers at $10.94 \%$, which are smaller than the corresponding proportions reported in studies conducted in Bangalore for monotherapy with ARBs (62\%) and beta blockers (15\%).

Table 7 shows that angiotensin receptor blockers were the most commonly prescribed group of antihypertensive drugs, used in $40.0 \%$ of Stage 2 patients, followed by ARB + thiazide at $24.0 \%$, calcium channel blockers at $12.0 \%$, beta blockers and ARB + CCB each at $8.0 \%$ and thiazide diuretics and high-ceiling diuretics each in $4 \%$ patients respectively in Stage 2 hypertension.

Table 7. Percentage of Stage 2 patients prescribed each drug regimen.

| Subset | Antihypertensive Medicine | Proportion of Patients (\%) |
| :---: | :---: | :---: |
| 1. | Thiazide diuretics | 4 |
| 3. | Angiotensin receptor blockers | 40 |
| 4. | Calcium channel blockers | 12 |
| 5. | Beta blockers | 8 |
| 6. | ARB + thiazide | 24 |
| 7. | ARB + CCB | 8 |
| 9. | High-ceiling diuretics | 4 |

Figure 3 shows that hypertensive patients with diabetes mellitus as an associated disease were the largest group with a comorbidity, so hypoglycemic drugs were prescribed for 51 patients, similarly hypolipidemic drugs were used in 28 patients to treat high coronary risk disease, antiasthmatic drugs were used in 15 patients for the treatment of asthma, and antiplatelets and antidepressant drugs were prescribed to 5 and 1 patients respectively.


Figure 3. Associated drugs in used in hypertension patients.
The proportions of patients prescribed one, two, three or more drugs are shown in Table 8. It can be seen that $39.0 \%$ patients received monotherapy and $61.0 \%$ received combination therapy. Among all patients, $48.0 \%$ took two drugs, $12.0 \%$ took three drugs and $1.0 \%$ took four or more drugs.

Table 8. Proportion of patients prescribed each number of drugs.

| Subset | Number of Drugs Prescribed | Proportion of Patients (\%) |
| :---: | :---: | :---: |
| 1. | One drugs | 39 |
| 2. | Two drugs | 48 |
| 3. | Three drugs | 12 |
| 4. | Four or more drugs | 1.0 |

All the prescription information collected from different Heart care Centers of the Kathmandu Valley were analyzed according to JNC 7 guidelines. Figure 4 shows that Lalitpur Heart Clinic, Nepal heart Care Center, Grande International Hospital and Janapoly clinic's prescriptions followed JNC 7 guidelines by $92.3 \%, 86.95 \%, 71.42 \%$ and $54.05 \%$, respectively, whereas a greater percentage of prescriptions which did not comply with JNC 7 guidelines were found in the Janapoly clinic (i.e., $45.95 \%$ ), with the numbers for Grande International Hospital, Nepal Heart Care Center and Lalitpur Heart Clinic being 28.58\%, 13.05\% and $7.7 \%$ respectively.


Figure 4. Compliance with JNC 7 Guidelines.
Cost analysis indicated that most patients were charged below 1500 rupees per month for the treatment of hypertension with associated drugs, and only the four-case numbers $60,63,40$, and 1 paid more than this. It was found that $25 \%$ of patients paid less than 500 rupees per month for their treatment, $50 \%$ patients were charged below 1000 rupees per month, and the remaining $25 \%$ of the patients were charged below 1500 rupees per month. The patients without comorbidities were charged below rupees 600 per month in only one case (i.e., case number 27 ).

The patients were divided into different categories according to their blood pressure, and the cost of the total drugs which they used during their treatment was analyzed on a monthly basis. Our data in Figure 5 show that the $4 \%$ of normal stage patients paid less than 500 rupees, and $2 \%$ were charged 501-1000 rupees per month for their treatment. In the prehypertensive stage, $22 \%$ patients were charged below Rs 500 and $10 \%$ were charged Rs 501-1000 per month for their treatment. In both Stage 1 and Stage 2 hypertension, most patients paid less than 500 rupees, and there was a similar proportion of patients who were charged between 1001 and 1500 rupees and more than 1501 rupees.


Figure 5. Stage-wise total cost analysis of the drugs. HTN: hypertension.

## 4. Discussion

The main aim of antihypertensive therapy is to prevent hypertension-related morbidity, mortality and complications [6]. As antihypertensive prescription is required for the entire life, the side effects, quality of life and cost of drugs are also important aspects. Keeping all factors in mind, various prescribing guidelines have been formulated. Treatment guidelines can be monitored by several methods. Drug utilization studies are one such method [6].

Our study group showed a higher prevalence of hypertension in elderly patients (i.e., $>40$ years). However, most females showed an earlier onset of hypertension than males (i.e., $20 \%$ greater than in males).

Data also revealed that there was a greater proportion of female patients than male patients who had hypertension along with diabetes mellitus and HTN with asthma, but the percentage of male patients with high-risk coronary disease was higher than in female patients. In hypertension along with its comorbidities, the most frequently prescribed group of drugs are ARBs. The associated drugs which were mostly prescribed in comorbidities were hypoglycemic drugs because the number of patients with DM was greater. Similarly, hypolipidemic drugs, anti-asthmatic drugs, antiplatelet and anti-depressant drugs were also prescribed to patients [7].

Hypertension is placed into different categories as given by the JNC 7 guidelines, and our data analysis results showed that patients with Stage 1 hypertension were observed in greater number ( $45 \%$ ), followed by prehypertension ( $32 \%$ ), Stage 2 hypertension ( $17 \%$ ), and normotension ( $6 \%$ ) [8].

This study also found that JNC 7 guidelines were followed in $74 \%$ of the study group. In all stages of hypertension, the most frequently prescribed group of drugs were ARBs, in prehypertension ARBs were followed by CCBs, thiazide, ACE inhibitors and beta blockers. In Stage 1 hypertension, ARBs were followed by ARB + thiazide and CCB + beta blocker. ARBs were also highly prescribed in Stage 2, which was followed ARB + thiazide, CCB, and beta blocker. More than half (i.e., $61 \%$ ) of the total patients in this study received two or more antihypertensive drugs [9-11].

After analyzing all the prescriptions collected from different Heart care Centers of the Kathmandu Valley, we found most of the institutes followed the seventh report of the Joint National Committee. In general, our data showed that $74 \%$ of the prescriptions followed the JNC guidelines [12,13].

Cost analysis showed that $57 \%$ of patients were charged below 500 rupees per month for their treatment. About $33 \%$ of total patients were charged between 501 and 1000 rupees per month, and $5 \%$ of patients were charged from 1001 to 1500 and more than 1500 rupees per month [14].

## 5. Conclusions

Based on our analysis of 100 prescriptions, the results of the current study were partly in accordance with the JNC 7 guidelines. Hence, larger extended studies are required to draw noticeable inferences.

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