

## Original Research Article

# Prescription pattern and rationality of antihypertensive drugs in patients of type 2 diabetes with hypertension: a pilot study

A. S. Bhore, Kalyani Khandare\*, K. A. Bansod

Department of Pharmacology, Dr. Panjabrao Deshmukh Memorial Medical College, Amravati, Maharashtra, India

**Received:** 17 December 2018

**Accepted:** 29 January 2019

**\*Correspondence:**

Dr. Kalyani Khandare,

E-mail: [kalyanikhandare18@gmail.com](mailto:kalyanikhandare18@gmail.com)

**Copyright:** © the author(s), publisher and licensee Medip Academy. This is an open-access article distributed under the terms of the Creative Commons Attribution Non-Commercial License, which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

### ABSTRACT

**Background:** As presence of hypertension (HTN) in type 2 diabetes significantly increases risk of morbidity and mortality, its control with rational use of antihypertensives is essential. Authors performed this study to understand the current prescribing pattern of antihypertensives in patients of type 2 diabetes and their assess the rationality to recommendations of JNC-8 guidelines.

**Methods:** Authors performed a cross sectional survey of prescription of diagnosed patients of diabetes with HTN at a tertiary care hospital. Prescription of patients attending medicine OPD were scanned. Data was collected over 2 month and analyzed with descriptive statistics.

**Results:** In 76 patients analysed, mean age was  $54.9 \pm 9.3$  years and 51.3% were females. Insulin and oral antidiabetic agents were prescribed in 16 (21.1%) and 73 (96.1%) patients respectively. One, two, three and four antihypertensive drugs were prescribed in 44 (57.9%), 28 (36.8%), 3 (3.9%) and 1 (1.3%) patients respectively and 24 (31.6%) of them were fixed dose combinations. Among antihypertensives, angiotensin converting enzyme inhibitors (ACEIs), angiotensin receptor blockers (ARBs), thiazide diuretics, beta blockers (BBs) and calcium channel blockers (CCBs) were present in 29 (38.2%), 26 (34.2%), 22 (28.9%), 21 (27.6%) and 14 (18.4%) respectively. In individual drugs, ramipril (28.9%) was most common followed by telmisartan (26.3%), hydrochlorothiazide (26.3%), atenolol (16, 21.1%) and amlodipine (17.1%) were frequently prescribed. In adjuvant drugs, antiplatelet (52.6%) and statins (56.6%) were common. This pattern of prescription was as per the recommendations of recent JNC-8 guidelines for treatment of hypertension.

**Conclusions:** ACEIs/ARB are the most preferred for HTN treatment in type 2 diabetes mellitus. Prescription of antihypertensives in our set up was rational as per JNC-8 recommendations.

**Keywords:** Guidelines, Hypertension, Prescription, Type 2 diabetes

### INTRODUCTION

Type 2 diabetes mellitus (T2DM) is rapidly growing epidemic in India and currently affects 7.4 million people with prevalence of 8.3%.<sup>1</sup> Hypertension (HTN) is the most common comorbid condition associated with T2DM.<sup>2</sup> Prevalence of HTN in T2DM is substantial and can be seen in more than 80% of patients.<sup>3</sup> Presence of

HTN in T2DM increases the risk of micro- and macro-vascular complications.<sup>2,4</sup>

Thus, a robust control of HTN is essential in patients with T2DM. Joint national committee 8 (JNC-8) guidelines provide recommendations for management of HTN.<sup>5</sup> JNC-8 recommendations are commonly followed by Indian physicians.<sup>6</sup> A target blood pressure (BP) of

<140/90 is advised by the most guidelines in patients T2DM.<sup>6</sup>

Achievement of better control of HTN in patients with T2DM is possible with use of multiple anti-hypertensive medications. Among these, ACEIs/ARBs, CCBs and thiazide diuretics are recommended as first line agents in T2DM.<sup>5</sup>

Despite availability of multiple effective antihypertensive drugs, control of HTN remains poor.<sup>7</sup> Therefore, assessment of drug prescribing pattern is necessary to ascertain that the current trends are according to guideline recommendations. This can identify any deviations from recommended therapy and help to take corrective measures.

Different studies have demonstrated that prescribing patterns of antihypertensive in T2DM are as per the international recommendations.<sup>8,9</sup> Understanding the prescribing practices help identify gaps in adequate therapy for control of HTN which is important especially in patients with T2DM. However, there have been no studies conducted at our institute to understand the prescribing pattern of antihypertensives in T2DM patients. Hence, authors planned this cross-sectional study.

**METHODS**

The study was conducted in patients of T2DM with HTN attending the medicine outpatient department (OPD) of a tertiary care hospital.

Ethical approval was taken from institutional ethics committee approved the protocol for the study. A cross-sectional, single centre, observational study at the medicine outpatient setting.

**Inclusion criteria**

- Patients of age ≥18 years,
- Both genders,
- Diagnosed as type 2 diabetes mellitus with hypertension by the physician,
- Has been prescribed treatment for HTN.

**Exclusion criteria**

- Patients requiring admission to the hospital,
- Pregnant and lactating women,
- Patient not willing to reveal their prescription data.

After taking the informed consent from the patients, the data from their prescription was recorded in case record form (CRF). The data collected includes demographic data like age, gender, and details of the drugs prescribed. Prescription of patients with T2DM and HTN in one-month duration were captured.

**Parameters assessed**

Total number of drugs per prescription was calculated. Among medications for T2DM and HTN, total number, drug class, individual drugs, dose and frequency of administration were recorded. Also, total number of drugs per prescription was calculated by counting all the drugs in prescription.

**Assessment of rationality to guidelines**

The prescribed treatment for HTN was then compared with recommendations of JNC-8 guidelines<sup>5</sup> to ascertain whether treatment was as per the recommendations or not.

**Statistical analysis**

The captured data was entered in to Microsoft excel sheet version 2010. Data was analyzed with descriptive statistics in excel sheet only. Categorical variables were presented as frequency and percentages. Continuous variables were presented as mean and standard deviation.

**RESULTS**

**Baseline characteristics**

In one-month prescription data captured in CRF, total 76 patients were included. Mean age of the patients was 54.9±9.3 years and most of them were in age group of 51-60years. Proportion of females was 51.3% (Table 1). Provides demographic characteristics of the patients.

**Table 1: Demographic characteristics.**

Parameter	Observation (%)
<b>Age (years)</b>	
Mean±SD	54.9±9.3
<b>Age groups</b>	
<40	6 (7.9)
41-50	20 (26.3)
51-60	27 (35.5)
>60	23 (30.3)
<b>Sex</b>	
Male	37 (48.7)
Female	39 (51.3)

**Total number of drugs per prescription**

Figure 1 shows that most prescriptions contained 5 to 7 drugs (51.3%) in total followed by <5 drugs (23.7%) and 8 to 10 drugs (19.7%).

**Oral antidiabetic agents prescribing pattern**

Metformin (80.3%), sulfonylureas (68.4%) and alpha glucosidase inhibitors (26.3%) were common antidiabetic drugs prescribed in study patients (Table 2).

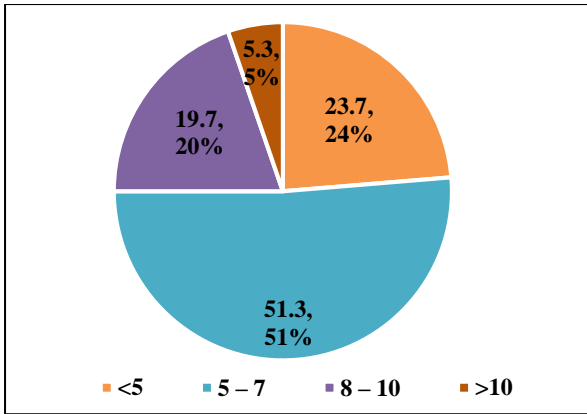


Figure 1: Total number of drugs per prescription.

Table 2: Details of antidiabetic drugs.

Anti-diabetic drugs	Observation (%)
Metformin	61 (80.3)
Sulfonylureas	52 (68.4)
Alpha glucosidase inhibitors	20 (26.3)
Thiazolidinediones	5 (6.6)
Insulin	16 (21.1)

**Antihypertensive prescribing pattern**

Among antihypertensive drugs, most patients were prescribed with either one (57.9%) or two (36.8%) antihypertensives.

Table 3: Details of the antihypertensive drugs.

Anti-hypertensive drugs	Observation (%)
<b>Total prescribed</b>	
One	44 (57.9)
Two	28 (36.8)
Three	3 (3.9)
Four	1 (1.3)
Prescribed as fixed dose combination	24 (31.6)
<b>Antihypertensive class</b>	
RAAS blocker	55 (72.4)
ACEIs	29 (38.2)
ARBs	26 (34.2)
Thiazide diuretic	22 (28.9)
Beta-blockers	21 (27.6)
Calcium channel blockers	14 (18.4)

Among them, 31.6% were prescribed in the form of fixed dose combinations (FDCs). Combined ACEIs and ARBs- the renin angiotensin aldosterone system (RAAS) blockers were most commonly prescribed antihypertensive class (72.4%) followed by thiazide diuretics (28.9%) and beta-blockers (27.6%) as shown in (Table 3).

As outlined in Table 4, ramipril (28.9%), telmisartan (17.1%), hydrochlorothiazide (26.3%), atenolol (21.1%) and amlodipine (17.1%) were the common drugs prescribed form their antihypertensive class.

Table 4: Prescribing frequency of individual anti-hypertensive drugs.

Antihypertensive drugs	Observation (%)
<b>ACEIs</b>	
Ramipril	22 (28.9)
Enalapril	5 (6.6)
Lisinopril	2 (2.6)
<b>ARBs</b>	
Telmisartan	13 (17.1)
Losartan	12 (15.8)
Olmesartan	1 (1.3)
<b>Thiazide diuretics</b>	
Hydrochlorothiazide	20 (26.3)
Chlorthalidone	2 (2.6)
<b>Beta-blockers</b>	
Atenolol	16 (21.1)
Metoprolol	4 (5.3)
Carvedilol	1 (1.3)
<b>CCB</b>	
Amlodipine	13 (17.1)
Nifedipine	1 (1.3)

**Concomitant drugs**

Among concomitant medications, antiplatelet and hypolipidemic drugs were prescribed in 52.6% and 56.6% patient respectively. Aspirin was prescribed in all cases either as single drug (39.5%) or in combination with clopidogrel (13.2%). Rosuvastatin and atorvastatin were prescribed with near equal frequency as shown in (Table 5).

Table 5: Details of other drugs prescribed.

Drugs	Observation (%)
Antiplatelets	40 (52.6)
Aspirin	30 (39.5)
Aspirin + Clopidogrel	10 (13.2)
Hypolipidemic	43 (56.6)
Rosuvastatin	22 (28.9)
Atorvastatin	21 (27.6)

**Rationality of antihypertensive drugs**

As per JNC-8 guidelines<sup>5</sup>, ACEIs/ARBs, Thiazide diuretics or CCBs are among the preferred choices for management of HTN in patients with T2DM. Authors observed similar pattern with most common use of ACEIs/ARBs, thiazide diuretics and CCBs in total covering treatment of over 80% patient of T2DM with HTN.

## DISCUSSION

In this study, mean age of the patients was 54.9±9.3 years suggesting patients who are above 50 years of age have higher propensity of development of cardiometabolic disorders like HTN and T2DM. Most patients received metformin, sulfonylureas and insulin which are cornerstone of T2DM treatment since past many decades. In India, these are most frequently prescribed anti-diabetic agents. A study from Indore city in India identified similar pattern of antidiabetic drug prescription where metformin and glimepiride were the most frequently prescribed drugs.<sup>10</sup> Another study from Navi Mumbai also reported that metformin dominated the antidiabetic therapy.<sup>11</sup> This is according to the American diabetes association (ADA) guidelines recommendations where metformin is advised as a first choice of drug for treatment of T2DM.<sup>12</sup>

In present study, prescription of single or two antihypertensive agents was more common. A study from Shekar AM from India reported that prescription of antihypertensive monotherapy was more common than polytherapy (60.14% vs 39.86% respectively) in patients with diabetes.<sup>13</sup> Study from Ethiopia by Abegaz et al, reported monotherapy in 50% of the patients.<sup>14</sup> Dhanraj et al, also reported that 41% of T2DM patients in their study received monotherapy treatment.<sup>9</sup> The selection of monotherapy versus polytherapy varies according to the BP levels and that makes prescribing pattern slightly different in different studies. Requirement of more than one antihypertensive agent suggest more severe form of HTN which needs adequate control.

We observed that antagonists of renin-angiotensin-aldosterone system were the most common drugs prescribed for control of HTN in patients. Following these, thiazide diuretics were second most frequently used agents.

Multiple studies have reported similar prescribing practices. Dhanraj et al, reported that ACEIs (59%) and ARBs (52%) were most frequently prescribed antihypertensives in their study from Punjab, India.<sup>9</sup> A study from Jammu by Kaur S et al, reported ACEIs (33.8%) as most frequently prescribed drugs for control of HTN.<sup>15</sup> Another study from Sharma JK et al, reported that ACEIs (85.36%) were the most common drugs for HTN treatment in T2DM.<sup>16</sup> These studies suggest that ACEIs or ARBs remain the first-choice agent in HTN with T2DM. The JNC-8 guidelines also recommend these drugs as first choice agents in patients of HTN with T2DM.<sup>5</sup> They have shown to provide end-organ protection and reduce the incidence of adverse cardiovascular outcomes and mortality in T2DM patients with HTN.<sup>17</sup>

Among other choices of drugs, thiazide diuretics and beta-blockers were prescribed with nearly similar frequency and CCBs were used in lesser number of

patients. However, a study from Dhanraj et al reported CCBs as second choice agents (29%) compared to diuretics (27%) or beta-blockers (14%).<sup>9</sup> Contrasting to this, Sharma et al. reported beta-blockers (33.44%) as second most frequently prescribed drugs followed by calcium channel blockers (29.95%).<sup>16</sup> This suggest the choice of second drug might vary in different regions of India depending on the local practices and preferences of physicians and patients.

However, a lower use of CCBs in present study calls for action to improve their prescribing in HTN with T2DM as JNC-8 guidelines have recommended CCBs among the first-line agents than beta blockers.<sup>5</sup> As far as individual therapies are concerned, there are no specific recommendations on individual molecules and can be preferred as per physician choices. Authors observed ramipril, telmisartan, hydrochlorothiazide, amlodipine and atenolol among the preferred drugs from their respective class.

As HTN and T2DM increase the risk of cardiovascular disease, preventive strategies are used in many of the patients. In supporting to this, authors observed prescription of antiplatelets and hypolipidemic drugs in 52.6% and 56.6% patients respectively.

This suggest that these individuals either had past CV event or might be at a higher risk of CV events qualifying for statin therapy as outlined in 2013 guidelines from American college of Cardiology and American heart association.<sup>18</sup> Considering a long list of medication prescribed in patients T2DM with HTN, it is not unusual to observed 5 to 6 drugs per prescription authors observed that over half of the study patients had 5 to 7 drugs in one prescription.

Presence of comorbidities like CV disease, stroke, infections increase the number of drugs per prescription in such patients. Having more than 10 drugs in prescription of these patient points to the fact that their disease is not controlled well requiring multiple therapy. Therefore, this suggests that efforts are necessary to improve the control of hyperglycemia and hypertension in these patients.

## CONCLUSION

In patients of HTN with T2DM, ACEIs/ARBs are most common drugs prescribed with most of them being monotherapy or dual drug therapy. Prescription of CCBs is on the lower side suggesting need to improve physician and patients' awareness on possible benefits with these drugs compared to those of beta blockers. Concomitant use of statins and antiplatelet suggest preventive strategies are also undertaken in these patients who are at risk of CV disease. In all, the prescribing pattern of antihypertensives in patients of HTN with T2DM is rational and is as per guideline recommendations.

## ACKNOWLEDGEMENTS

Authors would like to thank Dr. Sumanth MM, Assistant Professor, Department of Community Medicine, M.M.C and R.I., Mysore for assisting with the statistical work.

*Funding: No funding sources*

*Conflict of interest: None declared*

*Ethical approval: The study was approved by the Institutional Ethics Committee*

## REFERENCES

1. IDF. Diabetes atlas. 8th ed. Available at: <http://www.diabetesatlas.org/>.
2. Long AN, Dagogo-Jack S. Comorbidities of diabetes and hypertension: mechanisms and approach to target organ protection. *J Clin Hypertens*. 2011;13(4):244-51.
3. Jelinek HF, Osman WM, Khandoker AH, Khalaf K, Lee S, Almahmeed W, et al. Clinical profiles, comorbidities and complications of type 2 diabetes mellitus in patients from United Arab Emirates. *BMJ Open Diab Res Care*. 2017;5(1):e000427.
4. Hurst C, Thinkhamrop B. The association between hypertension comorbidity and microvascular complications in type 2 diabetes patients: a nationwide cross-sectional study in Thailand. *Diab Metabol J*. 2015;39(5):395-404.
5. James PA, Oparil S, Carter BL, Cushman WC, Dennison-Himmelfarb C, Handler J, et al. 2014 evidence-based guideline for the management of high blood pressure in adults: report from the panel members appointed to the eighth joint national committee (JNC 8). *JAMA*. 2014;311(5):507-20.
6. Hiremath JS, Katekhaye VM, Chamle VS, Jain RM, Bhargava AI. Current practice of hypertension in India: Focus on blood pressure goals. *J Clin Hypertens*. 2016;10(12):OC25.
7. Anchala R, Kannuri NK, Pant H, Khan H, Franco OH, Di Angelantonio E, et al. Hypertension in India: a systematic review and meta-analysis of prevalence, awareness, and control of hypertension. *J Hypert*. 2014;32(6):1170.
8. Alavudeen SS, Alakhali KM, Ansari SMA, Khan NA. Prescribing pattern of antihypertensive drugs in diabetic patients of southern province, Kingdom of Saudi Arabia. *ARS Pharm*. 2015;56(2):109-14.
9. Dhanaraj E, Raval A, Yadav R, Bhansali A, Tiwari P. Prescription pattern of antihypertensive agents in T2DM patients visiting tertiary care centre in north India. *Int J Hypertens*. 2012;520915.
10. Vengurlekar S, Shukla P, Patidar P, Bafna R, Jain S. Prescribing pattern of antidiabetic drugs in indore city hospital. *Indian J Pharm Sci*. 2008;70(5):637-40.
11. Agarwal A, Jadhav P, Deshmukh Y. Prescribing pattern and efficacy of anti-diabetic drugs in maintaining optimal glycemic levels in diabetic patients. *J Basic Clin Pharm*. 2014;5(3):79-83.
12. American Diabetes Association. Standards of medical care in diabetes-2017 abridged for primary care providers. *Clinical diabetes: a publication of the American Diabetes Association*. 2017Jan;35(1):5. Available at: <http://clinical.diabetesjournals.org/content/35/1/5>.
13. Pavani V, Cidda M, Krishna TR, Parmar MY, Nalini M. Study of prescribing patterns of antihypertensive drugs. *Int J Pharm Bio Sci*. 2012;2(2):317-27.
14. Abegaz TM, Tefera YG, Abebe TB. Antihypertensive drug prescription patterns and their impact on outcome of blood pressure in Ethiopia: a hospital-based cross-sectional study. *Int Pharm Res Practice*. 2017;6:29.
15. Kaur S, Gupta S, Kumar D, Lal M, Gilani Z. Prescribing pattern of antihypertensive drugs in a tertiary care hospital in Jammu-A Descriptive study. *JK-Pract*. 2012;17(4):38-41.
16. Sharma JK, Parmar SP, Trivedi HR. A study of prescribing pattern of antihypertensive drugs in hypertensive patients with co morbid diabetes in a tertiary care teaching hospital. *Int J Basic Clin Pharm*. 2018;(3):375-80.
17. Hao G, Wang Z, Guo R, Chen Z, Wang X, Zhang L, et al. Effects of ACEI/ARB in hypertensive patients with type 2 diabetes mellitus: a meta-analysis of randomized controlled studies. *BMC Cardiovasc Disord*. 2014;14(1):148.
18. Stone NJ, Robinson JG, Lichtenstein AH, Merz CN, Blum CB, Eckel RH, et al. 2013 ACC/AHA guideline on the treatment of blood cholesterol to reduce atherosclerotic cardiovascular risk in adults: A report of the American college of cardiology/American heart association task force on practice guidelines. *J Am Coll Cardiol*. 2014;63(2):2889-934.

**Cite this article as:** Bhore AS, Khandare K, Bansod KA. Prescription pattern and rationality of antihypertensive drugs in patients of type 2 diabetes with hypertension: a pilot study. *Int J Res Med Sci* 2019;7:982-6.