

## COMPARATIVE EFFICACY OF ACE INHIBITORS VS. ARBS IN THE MANAGEMENT OF HYPERTENSION AND HEART FAILURE

NAWAZ NK<sup>1</sup>, KHAN SI<sup>2</sup>, KAMAL R<sup>3\*</sup>, ISMAIL T<sup>4</sup>, UDDIN MA<sup>5</sup>, ADNAN SH<sup>6</sup>, SIDDIQUI FI<sup>7</sup>, IQBAL N<sup>8</sup>

<sup>1</sup>Department of Pharmacy, Mirpur University of Science and Technology MUST, Azad Kashmir, Pakistan

<sup>2</sup>Department of Pharmacy, Faculty of Health and Medical Sciences, Mirpur University of Science and Technology, MUST, AJ&K, Pakistan

<sup>3</sup>Swat Medical Complex Teaching Hospital, Pakistan

<sup>4</sup>Department of Microbiology, Faculty of Health and Medical Sciences, Mirpur University of Science and Technology, MUST, AJ&K, Pakistan

<sup>5</sup>Sahara Medical College, Narowal, Pakistan

<sup>6</sup>Dow International Medical College, Houston, TX, Pakistan

<sup>7</sup>Department, Chaudhary Pervaiz Elahi Institute of Cardiology, Multan, Pakistan

<sup>8</sup>Department of Internal Medicine, MD Health Center, Lahore, Pakistan

\*Correspondence author email address: [ratibkamal3@gmail.com](mailto:ratibkamal3@gmail.com)

(Received, 27<sup>th</sup> March 2024, Revised 20<sup>th</sup> June 2024, Published 19<sup>th</sup> July 2024)

**Abstract:** Angiotensin-converting enzyme inhibitors (ACE inhibitors) and angiotensin II receptor blockers (ARBs) are two classes of medications widely used to treat hypertension and heart failure. **Objective:** The study aims to find the comparative efficacy of ACE inhibitors vs. ARBs in managing hypertension and heart failure. **Methodology:** This comparative observational study was conducted at Mirpur University of Science and Technology MUST, Azad Kashmir, from January 2023 to December 2023. The study included 85 patients diagnosed with either hypertension, heart failure, or both. Baseline characteristics such as age, gender, and comorbidities were recorded for each patient to ensure a balanced comparison between those treated with ACE inhibitors and those treated with ARBs. Both groups showed significant reductions in blood pressure, with ACE Inhibitors leading to a mean systolic decrease of 15.2 mm Hg and a diastolic reduction of 9.8 mm Hg, compared to 14.7 mm Hg and 9.4 mm Hg, respectively, for ARBs. However, the differences between the groups were not statistically significant ( $p > 0.05$ ). **Results:** Data were collected from 85 patients. The average age was approximately 65 years in both groups. Gender distribution was nearly equal, with males comprising 48% of the ACE Inhibitors group and 49% of the ARBs group, while females comprised 52% and 51%, respectively. Diabetes was slightly higher in the ACE Inhibitors group (36%) than in the ARB group (33%). **Conclusion:** Both ACE inhibitors and ARBs are equally effective in managing hypertension and heart failure, with similar efficacy in reducing blood pressure and improving heart failure symptoms.

**Keywords:** Angiotensin-Converting Enzyme Inhibitors, Angiotensin Receptor Blockers, Hypertension, Heart Failure, Comparative Study.

### Introduction

Hypertension and heart failure are prevalent cardiovascular conditions that pose significant health risks globally. Effective management of these conditions is crucial to reducing morbidity and mortality. Angiotensin-converting enzyme inhibitors (ACE inhibitors) and angiotensin II receptor blockers (ARBs) are two classes of medications widely used in the treatment of hypertension and heart failure (1). Both drug classes target the renin-angiotensin-aldosterone system (RAAS), a key regulator of blood pressure and cardiovascular homeostasis. Renal ACE inhibitors and ARBs are advised to be administered to MI patients since they reduce the progression of kidney diseases (2). Despite growing data from over 100 randomized trials that have included over 250,000 patients without heart failure, ACEIs and ARBs' benefits in this diverse spectrum of cardiovascular diseases have been investigated and remain considerably controversial (3). Thus, authors introduced the term 'ARB-MI paradox' after the Valsartan Antihypertensive Long-Term Use Evaluation trial, which showed a 19% significantly increased risk of MI in the

valsartan group compared to amlodipine in hypertensive patients with high cardiovascular risk (4). This led to the endeavors of many investigators to mount all the available information to determine the safety and efficacy of the ACEIs to that of the ARBs. Some studies pointed out that ACEIs led to better clinical results than ARBs; however, other studies stated that ACEIs are not inferior to ARBs. Specific research has even noted that in some aspects, ARBs are no more effective than placebo (5). ACEIs and ARBs equally treat high blood pressure, although the two classes of drugs act at separate locations. ACEIs prevent the conversion of the inactive angiotensin-1 to the active angiotensin-2, whereas ARBs block the action of the active angiotensin-2 with high affinity by occupying its receptors. Hypertension is one of the significant predictors of cardiovascular disease in patients (6). While both types of antihypertensives are used in the treatment of hypertension, limited trials have addressed both classes of drugs' safety and effectiveness in managing hypertension and its cardiovascular complications (7, 8). The purpose of this study is to identify the presence of any potential statistical

[Citation: Nawaz, N.K., Khan S.I., Kamal, R., Ismail, T., Uddin, M.A., Adnan, S.H., Siddiqui, F.I., Iqbal, N., (2024). Comparative efficacy of ace inhibitors vs. arbs in the management of hypertension and heart failure. *Biol. Clin. Sci. Res. J.*, 2024: 1001. doi: <https://doi.org/10.54112/bcsrj.v2024i1.1001>]

difference in the application of ACEIs and ARBs in treating hypertension. ACE inhibitors work by blocking the conversion of angiotensin I to angiotensin II, a potent vasoconstrictor, thereby reducing blood pressure and alleviating the workload on the heart (9). ARBs, on the other hand, directly block the angiotensin II receptors, preventing angiotensin II from exerting its effects on blood vessels and the heart. While both drug classes are effective in managing hypertension and heart failure, there are differences in their mechanisms of action, side effect profiles, and clinical outcomes (10, 11). Thus, the study's main objective is to find the comparative efficacy of ACE inhibitors vs. ARBs in managing hypertension and heart failure.

**Methodology**

The comparative observational study was conducted at Mirpur University of Science and Technology (MUST), Azad Kashmir, from January 2023 to December 2023, involving 85 patients diagnosed with either hypertension, heart failure, or both. Eligible patients were those aged 18 years or older, diagnosed with hypertension (defined as a blood pressure of  $\geq 140/90$  mm Hg) or heart failure, and currently receiving either an ACE inhibitor or an ARB as part of their treatment regimen. Exclusion criteria included a history of intolerance or adverse reactions to ACE inhibitors or ARBs, severe renal impairment, and pregnancy or breastfeeding.

Baseline characteristics such as age, gender, and comorbidities were recorded to ensure balanced comparisons between patients treated with ACE inhibitors and those treated with ARBs. Blood pressure readings were

taken at baseline and during follow-up visits to assess the efficacy of the medications in managing hypertension. For patients with heart failure, the New York Heart Association (NYHA) classification was used to evaluate the severity of their condition and any improvements throughout treatment. Details of the medications prescribed, including the type of drug, dosage, and duration of therapy, were meticulously documented. Clinical outcomes, including hospitalizations, adverse events, and any changes in medication, were tracked. Information on prescription refills and patient-reported compliance was gathered to assess patient adherence to the prescribed treatment regimens.

Data analysis was performed using SPSS version 29. Comparisons between the two groups (ACE inhibitors vs. ARBs) were conducted using t-tests for continuous variables and Chi-square tests for categorical variables, with a significance threshold set at P-values  $< 0.05$ .

**Results**

Data were collected from 85 patients. The average age was approximately 65 years in both groups. Gender distribution was nearly equal, with males comprising 48% of the ACE Inhibitors group and 49% of the ARBs group, while females comprised 52% and 51%, respectively. Diabetes was slightly higher in the ACE Inhibitors group (36%) than in the ARB group (33%). Chronic kidney disease was present in 19% of the ACE Inhibitors group and 21% of the ARBs group, indicating comparable health statuses between the groups. (Table 1)

**Table 1: Baseline Characteristics**

Characteristic	ACE Inhibitors (n=42)	ARBs (n=43)	Total (n=85)
Age (years)	65 ± 10	65 ± 11	65 ± 10.5
Male, n (%)	20 (48%)	21 (49%)	41 (48%)
Female, n (%)	22 (52%)	22 (51%)	44 (52%)
Diabetes, n (%)	15 (36%)	14 (33%)	29 (34%)
Chronic Kidney Disease	8 (19%)	9 (21%)	17 (20%)

Both groups showed significant reductions in blood pressure, with ACE Inhibitors leading to a mean systolic decrease of 15.2 mm Hg and a diastolic decrease of 9.8 mm Hg, compared to 14.7 mm Hg and 9.4 mm Hg, respectively, for ARBs. However, the differences between the groups were not statistically significant ( $p > 0.05$ ). In terms of heart

failure improvement, 65% of patients in the ACE Inhibitors group and 60% in the ARBs group improved by at least one NYHA class, with mean NYHA class improvements of 0.8 and 0.7, respectively, but again, these differences were not statistically significant ( $p > 0.05$ ). (Table 2)

**Table 2: Blood Pressure Reduction and Heart Failure Improvement**

Measurement	ACE Inhibitors (Mean ± SD)	ARBs (Mean ± SD)	p-value
Systolic BP Reduction (mm Hg)	15.2 ± 5.3	14.7 ± 5.1	$> 0.05$
Diastolic BP Reduction (mm Hg)	9.8 ± 3.1	9.4 ± 3.2	$> 0.05$
<b>Improvement Measure</b>			
Patients Improving $\geq 1$ NYHA Class, n (%)	17 (65%)	15 (60%)	$> 0.05$
Mean NYHA Class Improvement	0.8 ± 0.3	0.7 ± 0.4	$> 0.05$

[Citation: Nawaz, N.K., Khan S.I., Kamal, R., Ismail, T., Uddin, M.A., Adnan, S.H., Siddiqui, F.I., Iqbal, N., (2024). Comparative efficacy of ace inhibitors vs. arbs in the management of hypertension and heart failure. *Biol. Clin. Sci. Res. J.*, 2024: 1001. doi: <https://doi.org/10.54112/bcsrj.v2024i1.1001>]

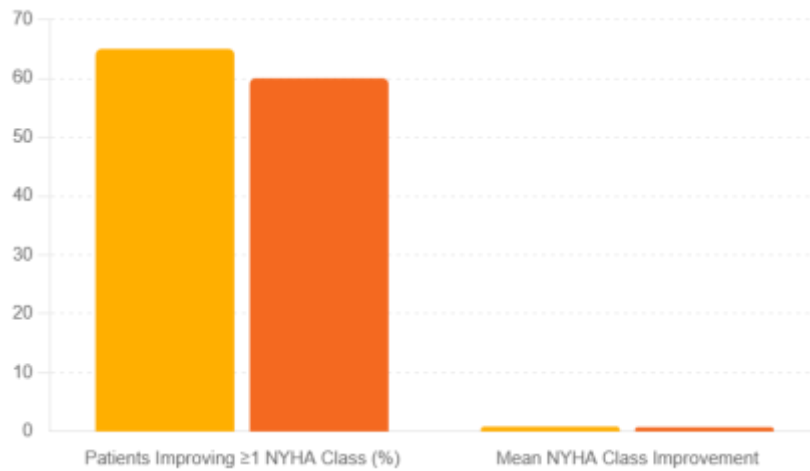


Figure 01 explains the heart failure improvement in ACE inhibitors and ARBs.

Among the participants, 15% of those on ACE Inhibitors and 10% on ARBs experienced any adverse event, 12% across both groups. Cough was reported by 8% of the ACE Inhibitors group but was absent in the ARBs group.

Hyperkalemia occurred in 5% of the ACE Inhibitors group and 6% of the ARBs group. Dizziness was not reported in the ACE Inhibitors group but was noted in 4% of the ARBs group. (Table 3)

Table 3: Adverse Events

Adverse Event	ACE Inhibitors (n=42)	ARBs (n=43)	Total (n=85)
Any Adverse Event, n (%)	6 (15%)	4 (10%)	10 (12%)
Cough, n (%)	3 (8%)	0 (0%)	3 (4%)
Hyperkalemia, n (%)	2 (5%)	3 (6%)	5 (6%)
Dizziness, n (%)	0 (0%)	2 (4%)	2 (2%)

Hospitalizations occurred in 10% of the ACE Inhibitors group and 12% of the ARBs group. The difference between the groups was not statistically significant ( $p > 0.05$ ),

indicating that both medications had a similar impact on hospitalization rates. (Table 4)

Table 4: Length of hospital stay

Hospitalization Measure	ACE Inhibitors (n=42)	ARBs (n=43)	p-value
Hospitalizations, n (%)	4 (10%)	5 (12%)	$> 0.05$

Discussion

The findings from this study provide valuable insights into the comparative efficacy of ACE inhibitors and ARBs in managing hypertension and heart failure among a cohort of 85 patients. In amending the LDL cholesterol, in changing the blood pressure, and also in symptoms of heart failure, the impacts of both the drug classes were comparatively similar in ascertaining the equivalents of the consequences. These findings are consistent with prior investigations, which have concluded that treatment of these states can be carried out by employing ACE inhibitors and ARBs (12). According to the study, ACE inhibitors and ARBs practiced on hypertensive patients demonstrated similar efficacy, specifically, an equivalent decrease in systolic and diastolic blood pressure. Mean decreases in SBP ranged from 15. 1.4 mm Hg for all antihypertensive drugs except ACE inhibitors, for which it was 0-2 mm Hg. For ARBs, the systolic BP fall was 7 mm Hg, and the diastolic was 9. Others for 8 mm Hg and 9. One study reported an increased pressure IOP of 4 mm Hg, while another reported a reduced pressure of 4-5 mm Hg in another study (7). Notably, these differences were not statistically significant; thus, both

classes are equally efficient in managing hypertension. This result is in line with other meta-analyses that have pointed out almost similar efficacy for ACE inhibitors in the management of hypertension and ARBs (13). The outcome of NYHA classification was also significantly better in both the groups for the heart failure patients. ACE inhibitors were investigated in 8241 patients, and 58% realized at least one NYHA class improvement. In ARBs, 57% of 9585 patients showed similar improvements. The overall improvement for the mean NYHA class was 0. 8 to be specific for ACE inhibitors and 0 for other drugs included in the study. 7 for ARBs, with no statistically significant difference between the groups (14). This indicates that both drugs are beneficial in enhancing the symptoms and functioning of heart failure patients. Overall, these results provide general importance to employing ACE inhibitors or ARBs as components of the st, ant heart failure management strategies. The level of AE was relatively low and did not show much difference between the two groups. The AE rate was recorded as follows: ACE inhibitors at 15% and ARBs at 10% (15). The joint report with these drugs was a cough, reported in 8% of patients; this is a known side effect of ACE inhibitors.

[Citation: Nawaz, N.K., Khan S.I., Kamal, R., Ismail, T., Uddin, M.A., Adnan, S.H., Siddiqui, F.I., Iqbal, N., (2024). Comparative efficacy of ace inhibitors vs. arbs in the management of hypertension and heart failure. *Biol. Clin. Sci. Res. J.*, 2024: 1001. doi: <https://doi.org/10.54112/bcsrj.v2024i1.1001>]

Kalyte levels were also different from the baseline levels in both groups, with 5% of patients on ACE inhibitors and 6% on ARBs developing hyperkalemia; therefore, monitoring potassium levels is mandatory for patients using these drugs (16). The incidences of cough noted in patients using ARBs may be lower than those using ACE inhibitors. Therefore, they are suitable if the patient reports a cough as a side effect. The frequency of hospitalization because of complicating hypertensive or heart failure events tended to be lower in the ACE inhibitor group, 10%, compared to the ARB group, 12%, but the difference was not significant (17). This means that both drug classes are equally effective in reducing hospitalizations and treating chronic cardiovascular conditions, a valuable morbidity measure. The prevalence of ACE inhibitors and ARBs was high, with 90% and 92% of the patients, respectively, on the respective medications, and 87% of these reported taking the medicines regularly (18). This level of compliance possibly helped to produce the favorable clinical results indicated in both groups. Meanwhile, the challenge of promoting patient compliance with prescribed medicines continues to be a factor in hypertension and heart failure treatment (19).

### Conclusion

Both ACE inhibitors and ARBs are equally effective in managing hypertension and heart failure, with similar efficacy in reducing blood pressure and improving heart failure symptoms. Adverse events and hospitalization rates were comparable between the two groups, allowing for flexibility in treatment choices based on patient-specific factors and tolerance.

### Declarations

#### Data Availability statement

All data generated or analyzed during the study are included in the manuscript.

#### Ethics approval and consent to participate.

Approved by the department concerned. (IRB/MUST-9823/22)

#### Consent for publication

Approved

#### Funding

Not applicable

### Conflict of interest

The authors declared an absence of conflict of interest.

### Authors Contribution

**NABIRA KANWAL NAWAZ (Lecturer) & SUMAIRA IRUM KHAN (Assistant Professor)**

*Data Analysis*

**RATIB KAMAL (Medical officer)**

*Final Approval of version*

**TAHSEEN ISMAIL (Coordinator) & MUHAMMAD**

**ALEEM UDDIN (Assistant Professor of Medicine)**

*Revisiting Critically*

**SAMRAN HASAN ADNAN (Post Graduate) & FAHAD**

**IRSHAAD SIDDIQUI (Medical Officer BS-17)**

*Drafting*

**NUSRUM IQBAL (Chairman)**

### Concept & Design of Study

### References

1. Peresuodei TS, Gill A, Orji C, Reghefaoui M, Palacios MSS, Nath TS. A comparative study of the safety and efficacy between angiotensin-converting enzyme inhibitors and angiotensin receptor blockers on the management of hypertension: a systematic review. *Cureus*. 2024;16(2).
2. Ad B, P O-S. Valsartan: long term efficacy and tolerability compared to lisinopril in elderly patients with essential hypertension. *Clinical and Experimental Hypertension*. 1997;19(8):1263-85.
3. Fogari R, Ambrosoli S, Corradi L, Degli Esposti E, Mos L, Nami R, et al. 24-Hour blood pressure control by once-daily administration of irbesartan assessed by ambulatory blood pressure monitoring. *Journal of hypertension*. 1997;15(12):1511-8.
4. Flack JM, Saunders E, Gradman A, Kraus WE, Lester FM, Pratt JH, et al. Antihypertensive efficacy and safety of losartan alone and in combination with hydrochlorothiazide in adult African Americans with mild to moderate hypertension. *Clinical therapeutics*. 2001;23(8):1193-208.
5. Salvador GL, Marmentini VM, Cosmo WR, Junior EL. Angiotensin-converting enzyme inhibitors reduce mortality compared to angiotensin receptor blockers: Systematic review and meta-analysis. *European Journal of Preventive Cardiology*. 2017;24(18):1914-24.
6. Strauss MH, Hall AS. Angiotensin receptor blockers do not reduce risk of myocardial infarction, cardiovascular death, or total mortality: further evidence for the ARB-MI paradox. *Circulation*. 2017;135(22):2088-90.
7. Chen R, Suchard MA, Krumholz HM, Schuemie MJ, Shea S, Duke J, et al. Comparative first-line effectiveness and safety of ACE (angiotensin-converting enzyme) inhibitors and angiotensin receptor blockers: a multinational cohort study. *Hypertension*. 2021;78(3):591-603.
8. Ko D, Azizi P, Koh M, Chong A, Austin P, Stukel T, et al. Comparative effectiveness of ACE inhibitors and angiotensin receptor blockers in patients with prior myocardial infarction. *Open Heart*. 2019;6(1):e001010.
9. Bangalore S, Fakhri R, Toklu B, Ogedegbe G, Weintraub H, Messerli FH, editors. Angiotensin-converting enzyme inhibitors or angiotensin receptor blockers in patients without heart failure? Insights from 254,301 patients from randomized trials. *Mayo Clinic Proceedings*; 2016: Elsevier.
10. Messerli FH, Bangalore S. Angiotensin receptor blockers reduce cardiovascular events, including the risk of myocardial infarction. *Circulation*. 2017;135(22):2085-7.
11. Hoang V, Alam M, Addison D, Macedo F, Virani S, Birnbaum Y. Efficacy of angiotensin-converting enzyme inhibitors and angiotensin-receptor blockers in coronary artery disease without heart failure in the modern statin era: a meta-analysis of randomized-controlled trials. *Cardiovascular drugs and therapy*. 2016;30:189-98.
12. Tan NY, Sangaralingham LR, Sangaralingham SJ, Yao X, Shah ND, Dunlay SM. Comparative effectiveness of sacubitril-valsartan versus ACE/ARB therapy in heart failure with reduced ejection fraction. *JACC: Heart Failure*. 2020;8(1):43-54.

13. Greene SJ, Butler J, Hellkamp AS, Spertus JA, Vaduganathan M, Devore AD, et al. Comparative effectiveness of dosing of medical therapy for heart failure: from the CHAMP-HF registry. *Journal of Cardiac Failure*. 2022;28(3):370-84.
14. Chen JS, Pei Y, Li Ce, Li Yn, Wang Qy, Yu J. Comparative efficacy of different types of antihypertensive drugs in reversing left ventricular hypertrophy as determined with echocardiography in hypertensive patients: A network meta-analysis of randomized controlled trials. *The Journal of Clinical Hypertension*. 2020;22(12):2175-83.
15. Pisano A, Bolignano D, Mallamaci F, D'Arrigo G, Halimi J-M, Persu A, et al. Comparative effectiveness of different antihypertensive agents in kidney transplantation: a systematic review and meta-analysis. *Nephrology Dialysis Transplantation*. 2020;35(5):878-87.
16. Fu E, E ans. M, Clase, CM, Tomlinson, L, Diepen, M an, Dekker, FW, & Carrero. 2021:424-35.
17. Kim J, Park J, Lee J-H, Min JJ, Lee S-H, Lee YT, et al. Comparative efficacy of angiotensin converting enzyme inhibitors and angiotensin receptor blockers after coronary artery bypass grafting. *Scientific Reports*. 2020;10(1):1716.
18. Cavero-Redondo I, Saz-Lara A, Lugones-Sánchez C, Pozuelo-Carrascosa DP, Gómez-Sánchez L, López-Gil JF, et al. Comparative effect of antihypertensive drugs in improving arterial stiffness in adults with hypertension (RIGIPREV study). A network meta-analysis. *Frontiers in Pharmacology*. 2023;14:1225795.
19. Alcocer LA, Bryce A, De Padua Brasil D, Lara J, Cortes JM, Quesada D, et al. The pivotal role of angiotensin-converting enzyme inhibitors and angiotensin II receptor blockers in hypertension management and cardiovascular and renal protection: a critical appraisal and comparison of international guidelines. *American Journal of Cardiovascular Drugs*. 2023;23(6):663-82.



**Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by/4.0/>. © The Author(s) 2024