The Impact of Antihypertensive Agents on Health-Related Quality of Life of Hypertensive Patients

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Abstract

Chronic diseases and their treatment regimen plays a greater role in determining the patient related health outcomes of which one is Health Related Quality of Life. Hypertension itself along with its pharmacotherapy impacts HRQoL of hypertensive patients. The primary objective of the study was to determine the impact of hypertension medications on the health-related quality of life among hypertensive patients with the secondary objective to assess which medication significantly affects health related quality of life of hypertensive patients. A quantitative questionnaire based cross-sectional survey was undertaken in outpatient departments of Sandeman Provisional Hospital (SPH) Quetta using an EQ-5D-3L to determine the Health-Related Quality of Life. Convenience Sampling technique was used for data collection. Majority of respondents (8.7%) had no problem in first three domains while having some problem in pain and anxiety domains. Following them 7.6% of patients had some problems in all domains of the EQ5D tool. Moreover, the ACE Inhibitors and Diuretics impacts HRQoL of patients of current study. A total of 263 participated in the study of which (25%) of the respondents were of age group 48-75 years old and were married 73.4% mostly. Majority (8.7%) had no problem in first three domains while having some problem in pain and anxiety domains. ACE Inhibitors and Diuretics significantly influence the health status of hypertensive patients acquiring pharmacotherapy. It was concluded that antihypertensive agents have a significant impact on HRQoL of hypertensive patients especially Angiotensin Converting Enzymes and Diuretics, which show significant impact on HRQoL of hypertensive patients. However, it is also suggested to assess the impact of antihypertensive therapies given in combination must also be studied in order to provide health professionals a better understanding of their prescribed regimen to improve patient related health outcomes.

Keywords: Hypertension, Health Related Quality of Life, Quetta, ACE Inhibitors, Diuretics, EQ5D-3L

1. Introduction

Hypertension is a continuing health problem of immense public health concern globally especially for developing world [1]. In 2010, 1.39 billion individuals globally were living with increased blood pressure, and out of those, 1.04 billion were from Lower Middle-Income Countries and 349 million in Higher Income Countries [2]. It was projected that in the year 2025, the number may increase to 60% to a total of 1.56 billion [3].

Globally, half of the total death toll caused by stroke and heart disease is due to hypertension. It is the most important risk factor for cardiovascular diseases [3]. Number of interventions, pharmacological and non-pharmacological, have been designed for the purpose of treating hypertension. Despite of having antihypertensives’ influence on lowering the risk of stroke, coronary heart disease, congestive heart failure, and mortality they also impose a greater effect on an individual’s life style and their Health related quality of life [4, 5]. The pharmacological treatment along with the disease itself has been reported to have side effects that moderately have an adverse effect on an individual’s Health-related quality of life especially the psychological domain [4, 6], however, the improvement in health related quality of life has also been evidenced by the studies [4]. The consequence of their adverse effect on health related quality of life was reported as poor compliance to antihypertensive treatment and deterioration of symptoms [6].

Health Related Quality of Life therefore can be defined as individuals perceived quality of life representing satisfaction in those areas of life likely to be affected by health status [7-9]. "Health-related quality of life" (HRQoL) integrates aspects of physical, psychological, and social health, as defined by the World Health Organization in 1948 [8]. Patients with chronic conditions often suffer from prolonged physical and psychological distress, which
deteriorates quality of life and leads to decreased life expectancy [10, 11]. The development of chronic conditions with established decreased life expectancies is very disturbing for the patients. The complex nature of diseases and sometimes their treatments has stressful effect on social and economic status of the patients. HRQoL can be altered by both the immediate and long-term treatment effects, especially in the chronic disease setting. Even in the controlled status the feeling of being ill heavily imbalances the quality of life (QoL) [7].

There has been considerable number of studies conducted on determining HRQoL of hypertensive patients but hardly any studies have been reported so far which has determined HRQoL of hypertensive patients who have been on the hypertensive pharmacotherapy in Pakistan and especially in the city of Quetta. Therefore, the primary objective of the study was to determine the impact of hypotensive medications on the health-related quality of life among hypertensive patients with the secondary objective to assess which medication significantly affects health-related quality of life of hypertensive patients.

Understanding the quality of life of hypertensive subjects as well as aspects that pharmacological and non-pharmacological interventions affect them may contribute health care professionals in evolving and applying interventions targeted at improving anti-hypertensive care which would ultimately lead to improved or better health-related quality of life [12, 13].

2. Methodology
2.1 Study Design and Setting
A quantitative questionnaire based cross-sectional survey was undertaken in outpatient departments of Sandeman Provisional Hospital (SPH) Quetta. The hospital provides services to majority of the city and provincial population.

2.2 Participants and Sampling Criteria
The estimated prevalence of hypertension in Balochistan has never been reported for which the sample size cannot be determined on the basis of prevalence. Addressing the current situation convenience data sampling technique was used and therefore 400 hypertensive patients were approached for the data collection.

Patients of age 18 or above having a documented diagnosis of hypertension for past 6 months or longer and were prescribed one or more antihypertensive medications were included in the study. Moreover, those hypertensive patients who understand Urdu language and were conscious with no mental incapacity and willing to participate were also included in the study. Patients having co-morbidities, critically ill or hospitalized were not included in the study. Pregnant ladies were also excluded from the study as these all conditions influence HRQOL along with hypertension and that would make the results biased.

2.3 Ethical Consideration
The national bioethical committee guidelines for researches involving humans were followed and therefore was reviewed and approved by Research Committee from Department of Pharmacy Practice, Faculty of Pharmacy and Health Sciences, University of Balochistan. Furthermore, approval was also granted from Medical Superintendent of the Hospital. Patients consent was also given a priority and confidentiality of their information was maintained.

2.4 Study Instrument
Health Related Quality of Life was assessed by EuroQoL-5D-3L, which is a generic instrument for the measurement of health outcomes such as HRQoL (EuroQol-Group, 1990). This HRQoL determining tool consists of two parts, of which the first part contains five domains (mobility, self-care, usual activities, pain/discomfort, anxiety/depression), and the second part consists of 20cm health meter called as EQ-VAS (Visual Analogue Scale). Each of 5 domains of first part of EQ-5D are measured by three levels of severity i.e. no problems, some or moderate problems and extreme problems. Responses to these five items are used to obtain the weighted EQ-5D index score, which demonstrate health state with a possible range from 0.594 to 1.0. The EQ-VAS (Visual Analogue Scale) consists of two distinct endpoints, the best imaginable health state (score of 100) and the worst imaginable health state (score of zero). EQ-5D was scored in accordance with the criteria designed by EuroQoL.

2.5 Data Analysis
The coded data was entered and were analyzed by using SPSS v. 20. Descriptive analysis was used for the patients’ characteristics and EQ-5D health status. Chi-square test was applied as an inferential statistical test to determine the impact of antihypertensives on the health-related quality of life of study participants.
3. Results

3.1 Study Population Characteristics

Out of 400 patients of hypertension 357 consented to participate in study out of which, 263 patients fulfilled the inclusion criteria. Patient characteristics demonstrates that majority (25%) of the respondents were of age group 48-75 years old and were married 73.4%. Most of the patients 37.3% (n=98) had no income while 24% (n=63) of patients had monthly income above 30,000 Pakistani Rupees as displayed in Table 1.

Table 1. Characteristics of Study Participants

| Characteristics       | Frequency n = 263 | Percentage (%) |
|-----------------------|-------------------|----------------|
| **Age group**         |                   |                |
| 18-27                 | 56                | 21.3           |
| 28-37                 | 42                | 16             |
| 38-47                 | 63                | 24             |
| 48-57                 | 68                | 25             |
| ≥58                   | 34                | 12             |
| **Gender**            |                   |                |
| Male                  | 138               | 52.5           |
| Female                | 125               | 47.5           |
| **Marital status**    |                   |                |
| Single                | 70                | 26.6           |
| Married               | 193               | 73.4           |
| **Ethnicity**         |                   |                |
| Punjabi               | 69                | 26.2           |
| Baloch                | 79                | 30             |
| Pathans               | 77                | 29.3           |
| Sindhi                | 17                | 6.5            |
| Others                | 21                | 8.0            |
| **Education**         |                   |                |
| Religious education only | 39            | 14.8           |
| Primary               | 10                | 3.8            |
| Matric                | 32                | 12.2           |
| FA/FSc                | 33                | 12.5           |
| B. A/B.Sc             | 63                | 24.0           |
| Higher education      | 86                | 32.7           |
| **Occupation**        |                   |                |
| Unemployed            | 53                | 22.2           |
| Self-employed         | 32                | 12.2           |
| Govt. servant         | 61                | 23.2           |
| Private servant       | 21                | 8.0            |
| Housewife             | 63                | 24             |
| Student               | 33                | 12.5           |
| **Income**            |                   |                |
| No income             | 98                | 37.3           |
| Not want to disclose  | 52                | 19.8           |
| Less than 10,000      | 13                | 4.9            |
| 10,000-20,000         | 16                | 6.1            |
| 21,000-30,000         | 21                | 8.0            |
| Above 30,000          | 63                | 24             |
| **Location**          |                   |                |
| Urban                 | 170               | 64.6           |
| Rural                 | 93                | 35.4           |
3.2 EQ5D Health Status

The hypertensive patients receiving pharmacotherapy have reported more than 60 different EQ-5D health states out of which 26 are mentioned in the Table 2 with bulk of respondents (8.7%) had no problem in first three domains while having some problem in pain and anxiety domains. Following them 7.6% of patients had some problems in all domains of the EQ5D tool.

Table 3 shows that majority of the study participants showed no problems in mobility (52.9%, n=139) and in taking care of themselves (65%, n=171). The table also demonstrates that most of the participants have some problems in performing their usual activities (47.9%, n=126). Additionally, bulk of study respondents had moderate ache (n=169, 64.3%) and were having moderate level of anxiety and/or depression (46.4%, n=122).

### Table 2. Self-Reported EQ5D Health States

| S. No. | Health status | Frequency n=263 | Percentage (%) |
|--------|---------------|----------------|----------------|
| 1      | 11122         | 23             | 8.7            |
| 2      | 22222         | 20             | 7.6            |
| 3      | 11111         | 18             | 6.8            |
| 4      | 11121         | 16             | 6.1            |
| 5      | 11222         | 13             | 4.9            |
| 6      | 11112         | 9              | 3.4            |
| 7      | 11223         | 9              | 3.4            |
| 8      | 12221         | 9              | 3.4            |
| 9      | 21122         | 9              | 3.4            |
| 10     | 21121         | 6              | 2.3            |
| 11     | 21123         | 6              | 2.3            |
| 12     | 22232         | 4              | 1.5            |
| 13     | 12112         | 3              | 1.1            |
| 14     | 12211         | 3              | 1.1            |
| 15     | 11123         | 3              | 1.1            |
| 16     | 11211         | 3              | 1.1            |
| 17     | 32232         | 3              | 1.1            |
| 18     | 11212         | 3              | 1.1            |
| 19     | 22233         | 3              | 1.1            |
| 20     | 31333         | 3              | 1.1            |
| 21     | 12212         | 3              | 1.1            |
| 22     | 21112         | 3              | 1.1            |
| 23     | 12222         | 3              | 1.1            |
| 24     | 12323         | 3              | 1.1            |
| 25     | 11212         | 2              | 0.8            |
| 26     | 11113         | 2              | 0.8            |

### Table 3. EQ5D Domains

| EQ5D Domain                  | Frequency n=263 | Percentage |
|------------------------------|-----------------|------------|
| **Mobility**                 |                 |            |
| I have no problems in walking about | 139             | 52.9       |
| I have some problems in walking about | 111             | 42.2       |
| I am confined to bed          | 13              | 4.9        |
| **Self-Care**                |                 |            |
| I have no problems with self-care | 171             | 65         |
| I have some problems washing or dressing myself | 82               | 31.2       |
| I am unable to wash or dress myself | 10              | 3.8        |
| **Usual Activities**         |                 |            |
| I have no problems with performing my usual activities | 109             | 41.4       |
| I have some problems with performing my usual activities | 126             | 47.9       |
| I am unable to perform my usual activities | 28              | 10.6       |
Pain / Discomfort
I have no pain or discomfort 59 22.4
I have moderate pain or discomfort 169 64.3
I have extreme pain or discomfort 35 13.3

Anxiety / Depression
I am not anxious or depressed 80 30.4
I am moderately anxious or depressed 122 46.4
I am extremely anxious or depressed 61 23.2

3.3 Impact of Antihypertensive Agents on Health Status

The Table 4 of the impact of drugs on HRQoL shows ACE Inhibitors and Diuretics significantly influence the health status of hypertensive patients acquiring pharmacotherapy.

Table 4. Association of Antihypertensive agents with Health Status and VAS Score

| Drugs used      | Frequency | EQ5D-score Mean | SD± | P-value | VAS-score Mean | SD | P-value |
|-----------------|-----------|-----------------|-----|---------|----------------|----|---------|
| ACE Inhibitors  | 100       | 0.436           | 0.409| 0.008   | 63.60          | 13.65| 0.012   |
| ARB             | 47        | 0.566           | 0.327| 0.198   | 62.45          | 13.14| 0.255   |
| DIU             | 53        | 0.370           | 0.440| 0.005   | 61.04          | 16.50| 0.004   |
| BB              | 136       | 0.476           | 0.364| 0.082   | 62.99          | 12.4 | 0.114   |
| CCB             | 115       | 0.498           | 0.345| 0.318   | 64.70          | 12.1 | 0.816   |
| VD              | 71        | 0.514           | 0.329| 0.768   | 65.86          | 11.03| 0.290   |
| LOP             | 149       | 0.470           | 0.389| 0.061   | 63.87          | 12.5 | 0.099   |
| STA             | 101       | 0.451           | 0.388| 0.013   | 61.83          | 13.1 | 0.012   |
| SSRI            | 6         | 0.611           | 0.422| 0.380   | 71.67          | 13.2 | 0.290   |

4. Discussion

The influence of treatment of hypertension on quality of life has been scarcely investigated in population-based surveys. Yan et al found significant improvements in HRQoL during BP control therapies [14]. Similar results were obtained that drug interventions would considerably relieve symptoms, comfort patients’ anxiety, and improved their self-reported quality of life. [14,15]. Other studies, in contrast, have reported no or even adverse influences of BP lowering therapies on HRQoL [14,15]. Some studies have shown that changes in HRQoL vary by antihypertensive treatment [16].

The study determined that health related quality of life was affected in participants with hypertension under antihypertensive treatment. The association of drug treatment with health-related quality of life has been attributed pharmacological and adverse effects of BP-lowering drugs. The short-term impact of antihypertensive therapy on quality of life can be appraised [14]. Erickson et al and Trivisol reported lower health related quality of life due to symptoms associated with drug treatment [17,18]. Chung et al compared three antihypertensive therapies in a randomized clinical trial, while the three treatments showed no difference in the primary outcome, the results showed major differences in health-related quality of life [11]. On the other hand, Antihypertensive treatment has a positive effect on the health-related quality of life indicators, a positive effect that may be due to the drug effect or to the placebo effect [19].

The current study used a universally accepted tool for the assessment of quality of life, which is essential for demonstrating the overall trend and it was found in the present study that antihypertensives have an influence on health-related quality of life. The association of drug treatment with lower quality of life has not been attributed to adverse effects of BP-lowering drugs as a whole or to a group of agents. The TOHMS trial, which followed more than 1000 individuals randomized to four classes of blood agents and placebo for four years, did not find any difference in several scores of quality of life between participants who received active drugs or placebo [18,20].

From the findings of the present study, two of the antihypertensive drug groups i.e., diuretics and angiotensin converting enzymes ACE’s, showed significant association with health-related quality of life. In support of significant effect of diuretics on health-related quality of life most of the studies indicate negative effect of diuretics.
on health-related quality of life which may be due to altering urinary frequency, erectile disorder, muscle cramps and fatigue or inadequate symptomatic relief with diuretics drugs [21].

Croog et al evaluated the effect of thiazide diuretic alone or in combination with other antihypertensives lead to decreased health related quality of life scores [22].

However, much of the criticism on quality of life with thiazides is based on the use of (too) high doses [20, 23]. Age dependent significance of diuretic was reported in older patients with better health related quality of life [24]. The studies that have examined the quality-of-life aspects of ACE inhibitors have usually reported favorable results, although most of these studies were not necessarily directed towards patients with hypertension [25]. ACE inhibitors seem to be effective in maintaining or even improving cognitive function through mechanisms other than blood pressure reduction [26]. In particular, patients taking captopril experienced less sexual dysfunction and had higher scores on measures of general well-being and life satisfaction, work performance and cognitive function [27].

In case of supportive therapy statins showed significant effects, Discontinuation of statin treatment was associated with better quality of life [28]. On the other side Statin treatment has beneficial effects on vascular function [29], and generally low frequency of real adverse effects [30], all of which would be mechanisms to affect health related quality of life. One of the octogenarian studies, community-dwelling men showed no significant difference in the health-related quality of life between statin users and nonusers [31].

The TOHMS trial, which followed more than 1000 individuals randomized to four classes of blood agents and placebo for four years, did not find any difference in several scores of quality of life between participants who received active drugs or placebo [18], [20].

5. Conclusion

It was concluded by the present study that antihypertensive agents have a significant impact on HRQoL of hypertensive patients. The Angiotensin Converting Enzyme Inhibitors and Diuretics show significant impact on HRQoL of hypertensive patients. However, it is also suggested to assess the impact of antihypertensive therapies given in combination must also be studied in order to provide health professionals a better understanding of their prescribed regimen to improve patient related health outcomes.

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