



Serum Electrolytes and Red Blood Cell Membrane Potential of Hypertensive Patients

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Authors' contributions

This work was carried out in collaboration among all authors. Authors NC and NCI designed the study. Authors EA and Onyeabo Chimaraoke prepared the manuscript. Authors EAS and Onyejekwe Catherine interpreted the results. All the Authors read and approved the final manuscript.

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ABSTRACT

Serum electrolyte and red blood cell membrane potential of hypertensive patients in Owerri metropolis were investigated. A total of 50 volunteer subjects were used for the study. Thirty (30) of the volunteer subjects were hypertensive subjects and were used as test subjects; while the remaining 20 subjects were healthy subjects with normal blood pressure used as normotensive subjects (control). Results observed showed increased red blood cell (RBC) K^+ and Cl^- in hypertension subjects against normotensive subjects. Apart from K^+ which reduced significantly ($p < 0.05$), other electrolyte ions of the serum increased significantly ($p < 0.05$) in hypertensive subjects against normotensive subjects. However, Na^+ and Cl^- membrane potential was not significantly ($p > 0.05$) altered in hypertensive subjects against normotensive subjects while K^+ was

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significantly ($p < 0.05$) altered. The observed alterations in the parameters investigated in hypertensive subjects in the present study could be as a result of a host of derangements involving electrolyte metabolism, altered membrane transport and a possible increase in membrane fragility. This study has shown the serum electrolyte and red blood cell membrane potential of hypertensive patients in Owerri metropolis.

Keywords: *Electrolyte; hypertensive patients; membrane potential; red blood cell.*

1. INTRODUCTION

Hypertension is defined as blood pressure persistently equal to or higher than 140 (systolic) /90 (diastolic) mmHg at rest according to the World Health Organization [1]. It is a common chronic and a major global public health problem with a prevalence of 44% in Western Europe and 28% in North America [2-3]. Hypertension could be of primary or secondary types [3-4]. About 90% of patients with hypertension belong to the category of primary type with no definite cause but has been attributed to ageing, hereditary, eating habit, smoking, alcoholism, stress, fatigue, lack of exercise and obesity; while 10% of hypertensive patients suffer from the second type and such has been associated with diseases such as chronic renal disease, thyroid disease, coarctation of the aorta, amongst others [5-10]. Pressure easily returns to normal in secondary hypertensive patients when the underlying causes are treated [10]. Hypertension has also been recognized as a threat to the health of people in Africa. Kearney et al. [11] noted that by 2025, 75% of the world hypertensive population will be in developing countries.

In Nigeria, hypertension is the number one risk factor for diseases like stroke, heart failure, ischemic heart disease, and kidney failure [12]. In recent time, Nigeria has witnessed rapid increase in hypertensive patients due to a rapid increase in an adult population exposed to hypertensive risk factors. Diwe et al. [13] noted that hypertension is a very common non-communicable disease and of major public health importance in Nigeria, with a prevalence range of 8-64% depending on the study population, type of measurement and cut-off value used for defining hypertensive. However, Ajomuobi [8] reported the prevalence of hypertension at 30-45% in Nigeria. Mensah et al. [14] noted the heart, kidney, and brain as target organs in hypertension. The arterial damage blood vessels are a prime target of hypertensive damage [15]. Hypertensive target organ damage (TOD) is common in Nigeria.

Because of low awareness of hypertension in the country, hypertensive TOD is often what brings patients to healthcare facilities [15].

A lot of studies associated with hypertension have been carried out in Nigeria [8,13,15-19], but not much has been done regarding serum electrolyte and red blood cell membrane potential in hypertensive patients. The present study investigated this area and used hypertensive subjects in Owerri Municipal of Imo State, Nigeria as a case study.

2. MATERIALS AND METHODS

2.1 Description of Study Area

Owerri Metropolis consists of Owerri Municipal, Owerri North and Owerri West. It lies between coordinates 5°29'1.07" N and 7°01'59.70" E. The city of Owerri in Owerri Municipality, is assumed to be the headquarters of the metropolis because it is like the heartbeat of the metropolis. Owerri metropolis sits at the intersection of roads from Port Harcourt, Onitsha, Aba and Umuahia. It is also the trade centre for palm products, maize, yams and cassava. The metropolis house major markets such as Eke Ukwu Owerri market, New Markets, Relief markets amongst others. Inhabitants of Owerri municipal indulge in different types of works and business activities to make ends meet.

2.2 Study Population

The study population was made of fifty (50) volunteer subjects of which twenty (20) hypertensive subjects receiving treatment at Imo State Specialist Hospital, ten (10) hypertensive subjects that were not receiving treatments yet, while twenty (20) healthy subjects with normal blood pressure were used as control. Subjects selected for the present study were hypertensive men and women with systolic blood pressure of 150 mmHg or greater and diastolic blood pressure of 90 mmHg or greater ($Bp \geq 150/90$ mmHg). Apparently, healthy men and women

with normal blood pressure were selected as control. Hypertensive men and women; and healthy men/women whose informed consent was obtained.

Hypertensive subjects with diseases like diabetes, HIV, liver diseases, kidney diseases and sickle cell anemia were excluded to avoid complications in the results. Also, hypertensive subjects whose informed consent could not be obtained because they were skeptical about the research were as well excluded.

2.3 Blood Pressure Determination

The blood pressure (BP) of the subjects was taken after they had rested for about five minutes with their hands on the table and the feet on the floor. Accoson Mercury sphygmomanometer was the instrument used for the entire subjects and it was ensured that the cuff covered at least 2/3 of the upper arm. Korotkoff phases 1 and 5 were identified as corresponding to systolic and diastolic blood pressures respectively. Two readings were taken at an interval of five minutes. Systolic BP less than 140 mmHg and Diastolic BP less than 90 mmHg were taken as normal. Readings above these values were interpreted as elevated BP.

2.4 Collection of Blood Samples from the Volunteer Subjects

Blood samples were collected from the subjects with the help of syringes and placed in anticoagulant tubes. The collected blood samples were centrifuged to obtain the sera used for electrolyte estimation. The sedimented red blood cells were lysed with deionised water for estimation of intracellular electrolytes.

2.5 Determination of Test Parameters

Electrolyte concentrations for both serum and red blood cells of subjects in the present study were estimated following the instructions as directed on their diagnostic kits (Teco diagnostic kits).

2.6 Computation of Membrane Potential

This was carried out by using the Nernst equation

$$E_x = 60 \log_{10} \frac{[x^+]_0}{[x^+]_i} \text{ mv}$$

Where, E_x = membrane potential using x^0
 $[x^+]_0$ = Concentration of ion x^+ in plasma
 $[x^+]_i$ = Concentration of ion x^+ ion red blood cell

2.7 Statistical Analysis

Results were presented as mean and standard deviations of triplicate determinations. Students t-distribution was used to establish a significant difference at 5% significant levels.

3. RESULTS AND DISCUSSION

Table of mean systolic and diastolic blood pressure of normotensive and hypertensive subjects (Table 1) shows that mean systolic and diastolic blood pressure (BP) of hypertensive subjects were 170.60 ± 10.28 and 115.28 ± 6.28 respectively while systolic and diastolic blood pressure (BP) of normotensive subjects were 70.70 ± 8.39 and 93.50 ± 7.09 respectively. The systolic and diastolic blood pressure of hypertensive subjects significantly increased ($p < 0.05$) against those of normotensive subjects.

Table 2 shows the mean RBC K^+ , Na^+ , and Cl^- concentration of normotensive and hypertensive subjects. RBC K^+ , Na^+ , and Cl^- concentrations for hypertensive subjects were 92.68 ± 4.02 mEq/L, 16.97 ± 1.81 mEq/L, and 52.11 ± 3.48 mEq/L respectively against 94.51 ± 5.11 mEq/L, 16.61 ± 1.53 mEq/L, and 50.00 ± 4.59 mEq/L respectively for normotensive subjects. According to Mordecai [20], since many of the Na^+ transport mechanisms that are present in kidney cell membranes (such as $Na^+ + K^+$ co-transport and Na^+ pumps) are also present in RBC and WBC membranes, much attention has recently been devoted to the study of Na^+ transport in RBCs and WBCs. In previous studies, some hypertensive patients were found to have an unusually high RBC $[Na^+]_m$ level, even in studies in which the mean $[Na^+]_m$ values for hypertensive patients and normotensive subjects were not significantly different [21]. While many hypertensive patients have RBC $[Na^+]_m$ levels within the normal range, the RBC $(Na^+)_m$ distribution curve for the hypertensive patients appears to be skewed toward higher $[Na^+]_m$ values [21-23]. According to Giasuddin et al. [24], there are many conflicting reports about the differences in blood electrolyte levels between normotensive and hypertensive population. RBC Na^+ in hypertensive subjects increased insignificantly ($p > 0.05$) against that of normotensive subjects in the present study. RBC K^+ and Cl^- increased significantly ($p < 0.05$) in hypertensive subjects when compared to normotensive subjects.

The Framingham Heart Study found that adults with serum potassium ≥ 5.2 mEq/L had increased risk of hypertension [25]. Kesteloot et al. [26], Pikilidou et al. [27] and Rinner et al. [28] found that serum potassium level was negatively associated with blood pressure. Hu et al. [29] suggested that serum potassium level was lower in hypertension group compared with non-hypertension group. Serum K^+ level reduced significantly ($p < 0.05$) in hypertensive subjects against normotensive subjects in the present study. The observed reduction in K^+ is in line with the observation of Hu et al. [29]. The mechanism behind the observed reduction in hypertensive subjects remains unclear. Potassium and sodium play important roles in the maintenance of cellular functions, and raised or lowered serum potassium level may be harmful to health [30]. Sodium, the main extracellular cation has long been considered as the pivotal environmental factor for hypertension. Changes in serum urea, creatinine, Na^+ , and Cl^- are associated with impairment of renal function [31]. Reabsorption of filtered sodium by the renal tubules is increased in primary hypertension because of stimulation of several sodium transporters located at the luminal membrane, as well as the

sodium pump which is localized to the basolateral membrane and provides energy for transportation [32]. A pivotal luminal transporter is sodium-hydrogen exchanger type 3, which resides in the proximal tubule and thickens ascending the limb of the loop of Henle, where the bulk of filtered sodium is reabsorbed. The activity of this exchanger is increased in the kidneys of rats with hypertension [32]. The serum Na^+ increased significantly ($p < 0.05$) in hypertensive subjects when compared to normotensive subjects in the present study. The observation is inconsistent with earlier study of Hu et al. [29], who reported no significant difference in serum sodium level between hypertension and non-hypertension groups. Serum Cl^- and HCO_3^- increased significantly ($p < 0.05$) in hypertensive subjects when compared to those of normotensive subjects. Large epidemiologic studies curiously show that lower circulating levels of serum Cl^- are associated with higher cardiovascular and all-cause mortality, though the mechanism remains unclear [33]. Reduced Na^+ and increased Cl^- ions observed in hypertensive subjects in this present study are not in line with work of Giasuddin et al. [24], who reported normal levels of both ions in

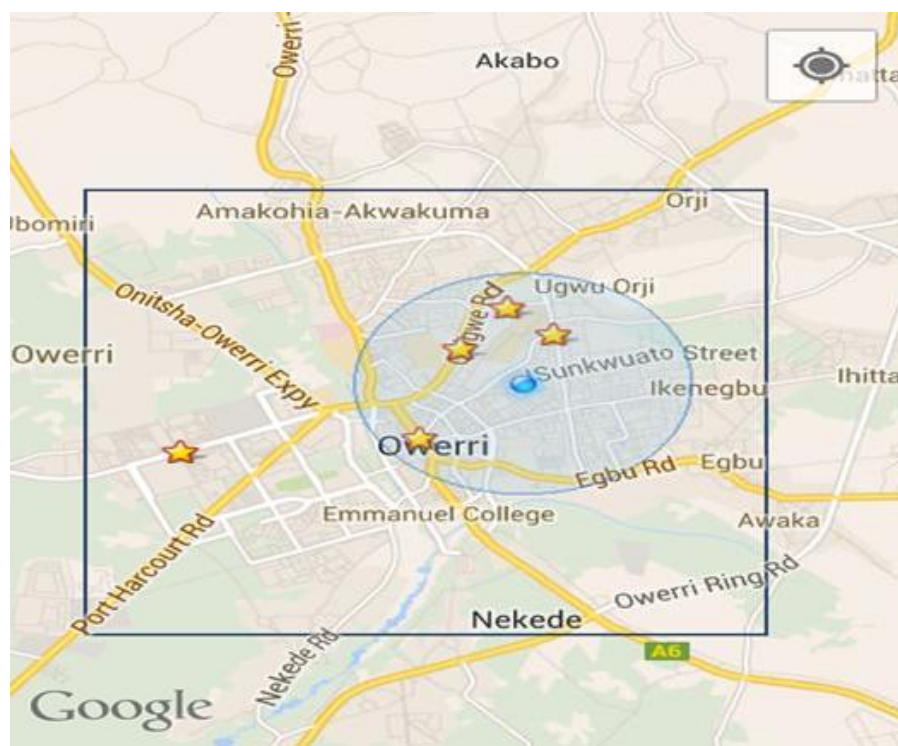


Fig. 1. Map showing the location Owerri Metropolis (Accessed from Google on 2nd March, 2019)

Table 1. Mean systolic and diastolic blood pressure of normotensive and hypertensive subjects

Parameters	Normotensive	Hypertensive	P value
Systolic BP	115.28±6.28	170.60±10.28	0.001
Diastolic BP	70.70±8.39	93.50± 7.09	0.001

Results are means and standard deviations of triplicate determinations

Table 2. Mean RBC K⁺, Na⁺ and Cl⁻ concentrations of normotensive and hypertensive subjects

Parameters	Normotensive	Hypertensive	P value
RBC K ⁺ (mEq/L)	94.51±5.11	92.68±4.02	0.05
RBC Na ⁺ (mEq/L)	16.61±1.53	16.97±1.81	0.28
RBC Cl ⁻ (mEq/L)	50.00±4.59	52.11±3.48	0.01

Results are means and standard deviations of triplicate determinations; RBC=Red Blood Cell

Table 3. Mean serum electrolyte concentration of normotensive and hypertensive subjects

Parameters	Normotensive	Hypertensive	P value
K ⁺ (mEq/L)	4.42±0.73	3.84±0.51	0.05
Na ⁺ (mEq/L)	141.12±4.16	142.77±3.61	0.036
Cl ⁻ (mEq/L)	95.78±5.50	100.44±3.61	0.001
HCO ₃ ⁻ (mEq/L)	25.73±1.77	27.24±5.50	0.001

Results are means and standard deviations of triplicate determinations

Table 4. Mean K⁺, Na⁺ and Cl⁻ membrane potential of normotensive and hypertensive subjects

Parameters	Normotensive	Hypertensive	P value
MP K ⁺	-80.11±4.43	-83.20± 3.68	0.001
MP Na ⁺	55.86±2.53	55.64±2.73	0.672
MP Cl ⁻	17.00±2.81	17.14±1.76	0.778

Results are means and standard deviations of triplicate determinations; MP membrane potential

hypertensive patients. This observation may be taken as an indication that overall renal handling of Na⁺ and Cl⁻ were abnormal in this set of hypertensive subjects. However, handling of electrolytes is modulated by a variety of substances such as aldosterone, angiotensin II, catecholamines and prostaglandins. Of these, aldosterone is the major determinant of potassium balance.

Table 4 reveals the mean K⁺, Na⁺ and Cl⁻ membrane potential of normotensive and hypertensive subjects. From the table, mean membrane potential (MP) for K⁺, Na⁺ and Cl⁻ for normotensive subjects were -80.11±4.43, 55.86±2.53 and 17.00±2.81 respectively while those of hypertensive subjects were -83.20±3.68, 55.64±2.73 and 17.14±1.76 respectively for K⁺, Na⁺ and Cl⁻. Alteration in membrane potential is indicative of membrane permeability [34]. This study revealed that K⁺ membrane potential increased significantly (p<0.05) in hypertensive subjects when compared to normotensive subjects. However, Na⁺ and Cl⁻ ion membrane potential were not significantly (p>0.05) altered in

hypertensive subjects against normotensive subjects.

4. CONCLUSION

This present study has shown that hypertension results in a host of derangements involving electrolyte metabolism, altered membrane transport and a possible increase in membrane fragility. This study also revealed that K⁺ membrane potential increased significantly (p<0.05) in hypertensive subjects when compared to normotensive subjects. However, Na⁺ and Cl⁻ ion membrane potential were not significantly (p>0.05) altered in hypertensive subjects against normotensive subjects. Changes in the red cell K⁺ and Cl⁻ concentrations as well as K⁺ and Na⁺ membrane potential may be used as markers in the assessment of hypertension, monitoring of treatment and diseases prognosis.

CONSENT

All informed consent was sought from the volunteer subjects and adequate verbal

information was provided for the subjects, which enabled them to know the essence of collecting their blood samples and the nature of the research work. Anonymity was assured as names were not required at any stage of the study. The participants benefitted from the study by having the knowledge of their blood pressure, electrolyte concentrations and blood components checked.

ETHICAL APPROVAL

The ethical clearance to conduct this study was appropriately obtained from the ethical and research committee of Imo State University/Imo State hospital management board.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

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