

## **Zinc and Copper Levels among Hypertensive Subjects in Port Harcourt, Nigeria**

### **Abstract**

Hypertension is a serious medical condition that is increasingly affecting increasing number of individuals globally. This study investigated the levels of copper and zinc, and the copper/zinc ratio among hypertensive subjects in Port Harcourt. A total of 200 subjects, aged between 30 and 70 years, were involved in the study, comprising 100 hypertensive subjects and 100 apparently healthy control subjects. Five (5ml) of venous blood were collected from each subject and put into plain bottle for the determination of copper and zinc using atomic absorption spectrometry (AAS). The data generated were analyzed using SPSS version 23. The mean copper and zinc levels were significantly higher in the hypertensive subjects compared to the control, as well as the systolic and diastolic blood pressure values. The levels of these parameters were significantly higher in male hypertensives compared to the female subjects. The results from this study show that there is an increased level of copper and zinc in hypertension, and these increases are more in men hypertensive subjects than in female hypertensive subjects. The assessment of the levels of copper and zinc, and copper/zinc ratio in hypertension is recommended in order to help in the management of the condition.

**Keywords:** Zinc, Copper, copper/zinc ratio, hypertension, Port Harcourt, Nigeria

### **Introduction**

Hypertension (sometimes referred to as high blood pressure), is generally defined as systolic blood pressure of 140 mmHg or higher and/or diastolic blood pressure of 90 mmHg or higher [1]. When the heart pumps blood for circulation, blood pushes against the walls of the artery through which they flow. This push creates a pressure on the arterial walls.

Blood pressure is usually measured as systolic and diastolic blood pressures. Systolic blood pressure is the pressure in the arteries when the heart beats, while diastolic pressure is the pressure in the arteries when the heart rests [2]. Primary hypertension (also called Essential hypertension) is the most prevalent type, and affects 90-95% of hypertensive patients, while secondary hypertension makes up about 5-10% of the hypertensive patients [1]. The pathogenesis of primary hypertension is multifactorial and complex; Genetic factors, environmental factors, such as sedentary lifestyle, stress, smoking, obesity, salt (sodium) sensitivity, and alcohol intake, are significant factors. Hyperactivity of the renin-angiotensin-

aldosterone system and sympathetic nervous system, abnormal production of natriuretic peptides, and deficiency in endothelial vasodilatation substances are also contributory factors. Secondary hypertension is hypertension that is caused by an underlying condition with a known cause, such as chronic kidney disease, aortic or kidney artery stenosis, or endocrine problems including elevated levels of aldosterone, cortisol, or catecholamine [3]. It is important to recognize secondary hypertension from primary hypertension, because it is treated differently to essential hypertension, by treating the underlying cause of the elevated blood pressure [1]. Generally, hypertension is a chronic condition that is marked by elevated blood pressure [4]. It is often called ‘the silent killer’ because it shows no early symptoms [1].

Trace elements are necessary for the optimal physiological processes in human health; excess or deficiency may lead to various diseases [5]. Therefore, changes in the levels or concentrations of these trace elements in the human body can trigger abnormal metabolic processes which can lead to life-threatening disease conditions. Copper and zinc are two important trace elements that are part of the antioxidant enzymes. It is therefore necessary to study the levels of these elements in hypertension.

This study evaluated the levels of copper and zinc among subjects with hypertension.

## **B. Methodology**

### **(i) Study Design**

A total of 200 subjects, aged between 30 and 70 years, were involved in this study, involving 100 subjects with hypertension (who were attending clinic) and 100 apparently healthy control subjects. The sample size was determined using GPower version 3.1.9.2.

For each subject, blood pressures were recorded twice, and the mean of two readings was calculated for each patient. The age, weight, and height, were also measured, and the body mass index (BMI) calculated for each subject.

## **(ii) Sample Collection**

Five (5) millilitres of venous whole blood was collected with minimum stasis from each subject into vacuum tubes. Special care was taken to avoid haemolysis of the blood samples during handling and the blood separated immediately from whole blood by centrifuging to get the serum. The serum was put into a clean plain bottle.

## **(iii) Sample Analysis**

Samples were digested according to the method adopted from a earlier study [6]. Briefly, 1ml each of the serum, was taken into Pyrex flask separately. Then, 3ml of freshly prepared mixture of concentrated nitric acid and hydrogen peroxide ( $\text{HNO}_3$ -  $\text{H}_2\text{O}_2$ ) (2:1 v/v) were added the sample and stood for 10minutes. Small amount of anti-bumping was also added. The mixture was heated following a one – stage digestion programmed at 80% of the total power in a sharp light- up (Dial) domestic microwave oven with maximum heating power of 800 w for 3 minutes after this the digestion flasks were cooled. It was heated until the volume reduced to two third of its original volume and its colour became clear and turned yellowish and all its components digested.

The resulting solutions were evaporated to semidried mass to remove excess acid. The worked-up samples were allowed to cool. The solution was carefully filtered using whattman filter paper of 0.2mm. After which the digested sample was introduced into 50ml volumetric flask and make it up to the mark with distilled water. The sample is stored in a sample bottle and kept below 4°C before analysis. A blank extraction was carried out through the complete procedure using triply distilled water. Digested samples and reagent blank were analyzed for copper (Cu) and zinc (Zn) by Atomic absorption Spectrophotometer (AAs) using acetylene flame at wave length of 217nm. The atomic Absorption spectrophotometer was calibrated using zinc standard solution and copper standard solution before measurement.

### Calculation

$$\text{Concentration} = \frac{\text{Absorbance of test}}{\text{Absorbance of standard}} \times \text{concentration of standard}$$

### (iv) Statistical Analysis

Data from this study were analyzed using Statistical Package for Social Sciences (SPSS) version 23. Comparison of means was done using t-test and ANOVA, with p-values less than or equal to 0.05 being considered statistically significant. Results were expressed as mean  $\pm$  Standard deviation.

## C. RESULTS

**Table 1: Demographic Information**

The data from this study indicate that demographic data shown below:

| Male | Female | Age (yrs) |
|------|--------|-----------|
|------|--------|-----------|

|               |    |    |        |
|---------------|----|----|--------|
| Hypertensives | 58 | 42 | 52± 3  |
| Control       | 64 | 36 | 54+ _2 |

**Table 2: Comparison of Biochemical parameters**

The table below shows the comparison of the levels of copper and zinc among the subjects. The hypertensives had significantly higher copper levels and significantly lower zinc levels. The copper/zinc ratio is also significantly raised in the hypertensive subjects.

|                        | <b>Copper<br/>(µg/ml)</b> | <b>Zinc<br/>(µg/ml)</b> | <b>Cu/Zn</b> | <b>SP<br/>(mmHg)</b> | <b>DP<br/>(mmHg)</b> |
|------------------------|---------------------------|-------------------------|--------------|----------------------|----------------------|
| Hypertensives Subjects | 1.34± 0.09                | 0.85± 0.08              | 1.58± 0.14   | 159.83± 16.37        | 92.17± 6.25          |
| Control Subjects       | 1.16± 0.07                | 1.08± 0.07              | 1.08± 0.06   | 107.80± 7.97         | 72.85± 4.48          |
| p-value                | <0.001                    | <0.001                  | <0.001       | <0.001               | <0.001               |
| t-value                | 10.31                     | 14.910                  | 21.460       | 18.634               | 16.829               |

**Table 3: Comparison of Biochemical Parameters According to Sex**

The male hypertensives had significantly higher copper levels compared to the female subjects. However, there were no significant differences in zinc levels and the cu/zn ratio.

|         | <b>Copper</b><br><b>(µg/ml)</b> | <b>Zinc</b><br><b>(µg/ml)</b> | <b>Cu/Zn</b> | <b>SP</b><br><b>(mmHg)</b> | <b>DP</b><br><b>(mmHg)</b> |
|---------|---------------------------------|-------------------------------|--------------|----------------------------|----------------------------|
| Female  | 1.31± 0.08                      | 0.83± 0.06                    | 1.58± 0.14   | 159.29± 12.30              | 92.25± 6.46                |
| Male    | 1.37± 0.09                      | 0.87± 0.08                    | 1.59± 0.14   | 160.32± 19.53              | 92.10± 6.16                |
| p-value | 0.014                           | 0.103                         | 0.778        | 0.810                      | 0.926                      |
| t-value | 2.530                           | 1.658                         | 0.283        | 0.241                      | 0.093                      |

#### **D. Discussion**

This study evaluated the levels of copper and zinc, as well as the copper/zinc ratio among hypertensive subjects.

The blood pressures of the hypertensives were significantly higher than the control subjects. The significantly raised systolic and diastolic blood pressures of the hypertensive subjects is understandable because of the consistently higher than normal pressure of blood flowing in their system, which is a major characteristic of hypertension [7].

The mean serum zinc and copper levels were significantly higher in hypertensive subjects compared to control subjects. This finding agrees with an earlier study [8], which reported a similar observation among patients with hypertension. Increased levels of zinc favours the activity of the enzyme, carbonic anhydrase which eventually predisposes to the development of hypertension [9].

The hypertensive subjects in this study had significantly higher levels of copper compared to the control subjects. It has been reported that copper levels are raised in conditions of inflammation [10]. Increased levels of free radicals and oxidative stress are and known contributors to the pathogenesis of hypertension [11]. Raised levels of copper increase the levels of free radicals, which can lead to vascular resistance and hypertension [9].

Both elements, copper and Zinc, are important factors in the function of enzyme superoxide dismutase, an essential antioxidant enzyme (SOD) [8]. Therefore, alterations in the levels of these elements may negatively affect the activity of SOD, which may in turn cause vasculo-endothelial dysfunction in hypertension.

Also, the hypertensive subjects had significantly higher copper/zinc ratio. Elevated copper/zinc levels have been associated with higher risks of cardiovascular mortality [12].

These findings of raised copper and copper/zinc ratio, and significantly low zinc levels in hypertensive subjects attending clinic in our study agree with the work of another study [13], which reported a similar finding.

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