Assessment of The Levels of Serum Zinc and Copper Among Sudanese Patients with Sickle Cell Anemia in Khartoum State

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7 Abstract:

8 **Background:** Sickle cell anemia (SCA) is associated with increased risks of 9 multiple micronutrient deficiencies, zinc deficiency has been observed in 10 patients with Sickle cell anemia, due to chronic hemolysis with subsequent 11 loss of zinc from RBCs.

12 **Objectives:** The aim of this study was to assess the level of zinc and copper 13 among Sudanese Patients with sickle cell anemia (SCA).

Methods: Across sectional study was conducted during the period from March to April 2018, fourty samples from diagnosed patients with sickle cell anemia (SCA), admitted to

17 Albuluk hospital in Khartoum state as cases and fourty samples from healthy18 individual

With normal hemoglobin as controls, the level of zinc and copper was measured by using atomic absorption spectrophotometer, Data analysis was carried out by SPSS version 21.

22 **Results:** There were a significant decreased in the levels of zinc and copper in patients with sickle cell anemia (SCA) with p-value = (0.000) (0.000) respectively, 23 when compared to healthy individuals. \pm SD; 0.137 \pm 0.079 24 The (Mean versus0.705±0.138ng/L) (0.512±0.290 versus 0.923±0.214ng/l) respectively. Also 25 there was insignificant variation in the levels of zinc and copper in male patients 26 27 compared to female patients, p-value (0.345) (0.656) respectively. The (Mean ±SD; 28 0.144 ± 0.079 versus 0.127 ± 0.080 ng/L) (0.502 ± 0.271 ng/L versus 0.525 ± 0.321 29 ng/L) in respectively, also there was negative correlation between levels of (zinc and copper) with duration time of disease, with (R = -0.603, p-value = 0.000) and 30 p-value= 0.004) respectively, But there was no correlation between 31 (R = -0.443)levels of (zinc and copper) with ages. 32

In Conclusion: the levels of zinc and copper were decreased in patients with sickle cell anemia when compared to healthy individuals, and also there was negative correlation between the levels of (zinc and copper) and duration time of disease.

36 **KEYWORDS:** Sickle cell anemia, zinc, copper, micronutrient and Sudanese.

37 Introduction:

38 Sickle cells anemia (SCA) is a hemoglobinopathy characterized by chronic 39 hemolysis,

40 Chronic inflammation, immune deficiency, a heterogeneous Clinical picture and 41 organ

Damage.⁽¹⁾ SCA is associated with increased risks of multiple micro nutrient 42 deficiencies These deficiencies may have a significant impact on SCA severity 43 indices including growth retardation; cell-mediated immune dysfunction, and 44 cognitive impairment with a negative impact on morbidity and mortality.⁽²⁾ In 45 addition, these nutriments have a major role in the protection of the red cell 46 membrane against stress and free radical mediated by oxidation in SCA.⁽³⁾ Sickle 47cell disease (SCD) is common especially in Africa and among Negroid race. In sickle 48 cell disease; the deficiencies of essential elements some of which are vital in red 49 cell maintenance, body growth and development have been observed.⁽⁴⁾ The sickle 50 hemoglobin is known to interact with diverse gene and environmental factors 51 producing a multisystemic disease with several phenotypes. ⁽⁵⁾ 52

Minerals are inorganic substances, with chemical constituent present in all body 53 tissues and fluids and they are necessary for the maintenance of certain 54 physicochemical processes which are essential to life.^(6,7) They are important for 55 human,^(8,9) so deficiencies or disturbances in the nutrition can cause a variety of 56 diseases and can arise in several ways.⁽¹⁰⁾ Nutrient metals from diet are 57 incorporated into blood if blood levels are depleted, transported into cells if cellular 58 levels are inadequate, or excreted if blood and cell levels are sufficient or 59 overloaded. When this system fails to function properly, abnormal levels and ratios 60 of trace metals can develop. One of the most common trace-metal imbalances is 61 62 elevated copper and depressed zinc; however the aim of this study was to assess the level of serum zinc and copper among Sudanese patients with sickle cell anemia. 63 Materials and methods: 64

- Study design: this was a Cross sectional case control study.

-Study area and period: Blood samples were collected from Albuluk hospital,
in Khartoum state, during the period from March to April 2018.

- Study population: forty Patients with sickle cell anemia (SCA) as a case and
forty sample from normal individual with normal hemoglobin as control,
gender and ages was matched (case group with age mean+SD (6.67+4.16)
years and control group with age mean+SD 6.37+4.16 years, sickle cell
anemic patients were 43% females and 57% males, and normal healthy
individual (control) were 40% female and 60% male.

74 -Inclusion criteria: patients with sickle cell anemia (SCA), were included

75 -Exclusion criteria: any patients take long term zinc supplements and

patients with other chronic diseases (liver disease, renal disease and heartdiseases), were excluded.

- Ethical consideration: The study was approved by ethical committee of
 Medical Laboratory Science, Clinical Chemistry Department –Alneelain
 University. Subjects involved in this study were informed by the aims of the
 study and its importance, and verbal informed consent was obtained from
 each participant.

- 83 **-Data collection:** by using questionnaire.
- Sampling: blood samples were collected and serum was separated.
- **-Method:** The levels of serum zinc and copper was measured by using atomic
- 86 absorption spectrophotometer.

- Quality Control: Pathological and Normal control sera were measured, to
 assure accuracy and precision of results.

-Data analysis: Data was analyzed using SPSS version 21.The results were
 expressed as percentage, Mean and SD. Independent T-test was obtained to
 compare the study parameters in case versus control group. Correlation was
 done to study the relationship between study parameters and study variables,
 p-value less than 0.05 considered significant.

94 Results:

95 Statistical analysis showed a significant decrease in the levels of zinc and copper among patients with sickle cell anemia (SCA), when compared to 96 healthy individuals (table1), also statistical analysis showed insignificant 97 variation in the levels of zinc and copper among patients with sickle cell 98 anemia when compared according to gender (table2), also statistical analysis 99 showed negative correlation between the levels of (zinc and copper) and 100 101 duration time of disease (SCA), (figure 1 and figure 2) respectively, and also 102 statistical analysis showed no correlation between the level of zinc and 103 copper with ages among patients with sickle cell anemia, (figure 3 and figure 104 4) respectively.

105 Table (1): Comparison the levels of zinc and copper in case versus control:

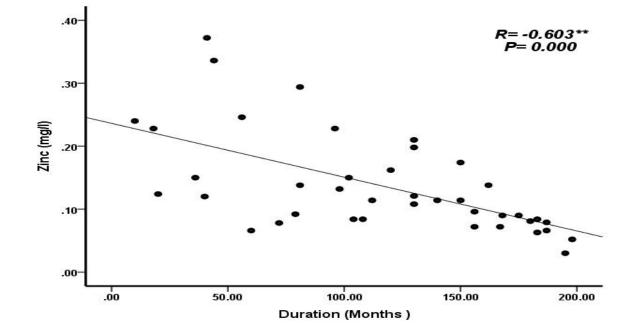
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Parameters	Case (Mean±SD)	Control (Mean±SD)	P-value
Copper	0.512±0.290	0.923±0.214	0.000
Zinc	0.137±0.079	0.705±0.138	0.000
	Copper	Copper 0.512±0.290	ParametersCase (Mean±SD)Control (Mean±SD)Copper0.512±0.2900.923±0.214

106 P-value less than 0.05 consider as significant

Table (2): Comparison the levels of copper and zinc in case group according to gender.

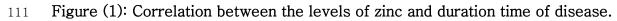
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Parameters	Male (Mean±SD)	Female (Mean±SD)	P-value
Copper	0.502±0.271	0.525±0.321	0.656
Zinc	0.144 ± 0.079	0.127±0.080	0.345

UNDER PEER REVIEW



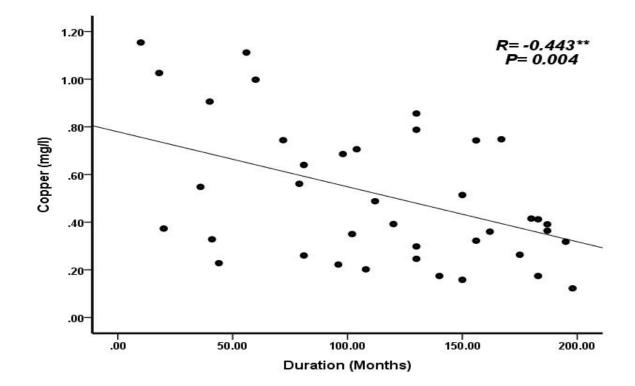
109 P-value less than 0.05 consider as significant





112 P-value less than 0.05 consider as significant

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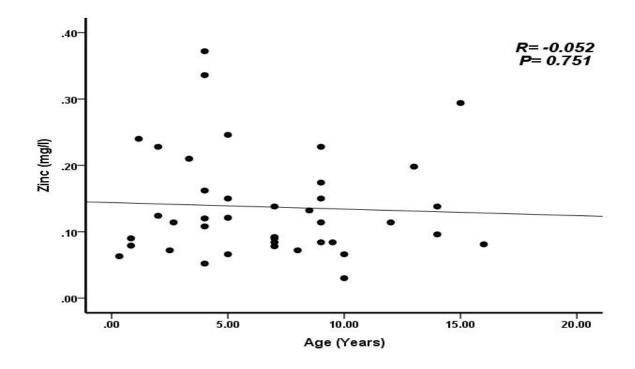


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- 117 Figure (2): Correlation between duration time of disease and copper levels.
- 118 P-value less than 0.05 consider as significant:
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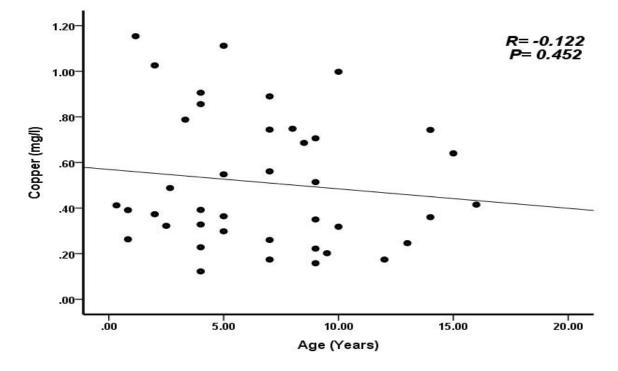
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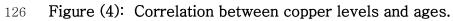
122 Figure (3): Correlation between zinc levels and ages.

123 P-value less than 0.05 consider as significant:

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UNDER PEER REVIEW





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128 P-value less than 0.05 consider as significant:

129 Discussion:

130 Sickle cell disease (SCD) is common especially in Africa and among Negroid race, 131 In sickle cell disease, the deficiencies of essential elements some of which are vital 132 in red cell maintenance, body growth and development have been observed, in the 133 current study the levels of zinc and copper showed a significant decrease in 134patients with sickle cell anemia (SCA) when compared to healthy individuals with pvalue (0.000), that might be occur due to chronic hemolysis with subsequent loss of 135 zinc from RBCs, Zinc deficiency can also be the result of the adverse effect of 136 hydrourea which increase zinc excretion.⁽¹¹⁾ This finding was agreement with 137 results of previous study done by (tagney and Philips, 1993; Parad, 2002; Idonij et 138139 al., 2011), which related zinc deficiency in sickle cell disease to manifestations such as growth retardation, hypogonadism in males, hyperammonemia, abnormal dark 140 adaptation and cell mediated immune disorder. Similarly, the biochemical evidence 141 for zinc deficiency in patients with SCD includes low zinc concentrations in plasma, 142erythrocytes, hair lymphocytes and granulocytes,⁽¹²⁾ also there was insignificant 143 variation in the levels of zinc and copper in patients with sickle cell anemia when 144 compared according to gender the p-value was (0.345 and 0.656) respectively, and 145 146 also there was negative correlation between levels of (zinc and copper) with duration time of disease, with (R = -0.603, p-value = 0.000) and (R = -0.443)147p148 value= 0.004) respectively, But there was no correlation between levels of (zinc and 149 copper) with ages.

150 Conclusions:

The levels of zinc and copper were decreased in patients with sickle cell anemia when compared to healthy individuals, and also there was negative correlation between the levels of (zinc and copper) and duration time of disease.

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