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

### Abstract:

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Background: Sickle cell anaemia (SCA) is associated with increased risks
 of multiple micronutrient deficiencies. Zinc deficiency has been observed
 in patients with Sickle cell anaemia, due to chronic haemolysis with
 subsequent loss of zinc from RBCs.

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

15 Methods: Across-sectional study was conducted from March to April 2018, 16 involving forty subjects who had been diagnosed of sickle cell anaemia 17(SCA), and had been admitted to Albuluk Hospital, in Khartoum State, as cases. Forty healthy individual with normal haemoglobin were recruited as 18 controls (age was matched in the groups ranged from 4 months to 16 19 years). Blood samples were collected and serum was separated, then the 20 21 levels of zinc and copper were measured, using atomic absorption 22 spectrophotometer. Data analysis was carried out, using SPSS version, 21 (Independent t-test was used to compare mean values in case versus 23 control group. Pearson's Correlation was done to study the relationship 24 between zinc and copper and age). 25

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27 Results: There was significant decreased in the mean levels of zinc and 28 copper in patients with sickle cell anemia when compared to control groups. The (Mean ±SD; 0.137±0.079 versus0.705±0.138ng/L, p, value= 29 0.00) for zinc (0.512±0.290 versus 0.923±0.214ng/l, p, value =0.00) for 30 copper. There was no significant difference between males and females in 31 <mark>the mean levels of zinc (p = 0.345) and copper (p = 0.656).</mark> The (Mean 32 ±SD; 0.144±0.079 versus0.127±0.080 ng/L) for zinc (0.502±0.271ng/L 33 versus 0.525±0.321 ng/L) for copper, there was no correlation between 34 levels of zinc, copper and ages, (r=0.052, p, value=0.750) for zinc, 35 (r=0.122, p, value=0.452) for copper. 36

- Conclusion: The levels of zinc and copper were decreased in patients with
   sickle cell anaemia, compared to healthy individuals. There was also
   negative correlation between the levels of zinc and copper and age of
   patients.
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42 **KEYWORDS:** Sickle cell anemia, zinc, copper, micronutrient and Sudanese.

#### 43 Introduction:

Sickle cells anaemia is a haemoglobinopathy, characterized by chronic 44haemolysis, chronic inflammation, immune deficiency, a heterogeneous 45 clinical picture and organ damage.<sup>(1)</sup> SCA is associated with increased 46 risks of multiple micro nutrient deficiencies These deficiencies may have 47a significant impact on SCA severity indices including growth retardation; 48 cell-mediated immune dysfunction, and cognitive impairment with a 49 negative impact on morbidity and mortality.<sup>(2)</sup> In addition, these nutrients 50 have a major role in the protection of the red cell membrane against 51 damage through free radical-mediated oxidation in SCA.<sup>(3)</sup> Sickle cell 52 disease is common especially in Africa and among Negroid race. In sickle 53 cell disease, there are deficiencies of some essential elements which are 54vital in maintenance of red cell integrity, body growth and development. <sup>(4)</sup> 55 The sickle haemoglobin is known to interact with diverse genes and 56 environmental factors, producing a multi-systemic disease with several 57 phenotypes. <sup>(5)</sup> 58 Minerals are inorganic substances, present in all body tissues and fluids 59

and they are necessary for the maintenance of certain physicochemical
processes which are essential to life.<sup>(6,7)</sup> They are important for human,<sup>(8,9)</sup>
so deficiencies or disturbances in the nutrition can cause a variety of
diseases, which can arise in several ways.<sup>(10)</sup> Two most common trace
metal imbalances are elevated copper and depressed zinc in SCA.
Therefore, the aim of this study was to assess the level of serum zinc and
copper among Sudanese patients with sickle cell anaemia.

### 67 Materials and methods:

68 **- Study design:** This was a cross-sectional case-control study.

-Study area and period of study: Blood samples were collected from Albuluk
Hospital, in Khartoum State, from March to April 2018.

- Study population: Forty sickle cell anaemic patients, aged between 4 months
 and 16 years, were recruited as cases and forty normal children with normal
 haemoglobin were enrolled as controls. The cases and control were age-matc
 hed; 17 of sickle cell anemic patients were females and 23 were males, and

75 16 of controls were females and 24 of them were males.

-Exclusion criteria: Any patients taking long term zinc supplements and
patients with other chronic diseases (liver disease, renal disease and heart
diseases), were excluded.

Ethical consideration: The study was approved by ethical committee of
 Medical Laboratory Science, Clinical Chemistry Department-Alneelain
 University. A verbal informed consent was obtained from the parents of each
 minor participant.

 <sup>76 -</sup>Inclusion criteria: Albuluk Hospital in-patients with sickle cell anemia who
 77 were treated with hydroxyurea, were included.

**-Data collection:** A demographic data was collected by using questionnaire.

- Sampling: About 2.5 ml of venous blood was collected from each participant,
 after formation of clot in the room temperature; the samples were centrifuged

<sup>88</sup> for 10 minutes in 3000 rpm then the serum was obtained and analyzed.

Method of assay of zinc and copper: The levels of serum zinc and copper
 was measured by using atomic absorption spectrophotometer (BUCK
 SCINTIFIC 210/211 VGP VER3.94C).

92 - Quality Control: Pathological and normal control sera were also used for the
 93 measurement of the metals, to assure accuracy and precision of results.

-Data analysis: Data was analyzed using SPSS, version 21.The results were
expressed as percentages, mean and SD. Independent t-test was used to
compare mean values in case versus control group. Pearson's correlation test
was done to study the relationship between study zinc and copper and age, pvalue less than 0.05 considered significant.

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### 100 Results:

Eighty participants were enrolled in this study; 40 patients (17 females and 2 101 102 3 males) with age mean+SD 6.67+4.16 years and 40 controls (16 female and <mark>24 males) with age mean<u>+</u>SD 6.37<u>+</u>4.16 years</mark>, age was matched in both grou 103 ps and ranged from 4 months to 16 years. Statistical analysis showed a signifi 104 105cant decrease in the levels of zinc and copper among patients with sickle cell 106 anemia (SCA), when compared to healthy individuals (table1), also statistical 107 analysis showed insignificant variation in the levels of zinc and copper among patients with sickle cell anemia when compared according to gender (table2) 108 ,and also statistical analysis showed no correlation between the level of zinc 109 and copper with ages among patients with sickle cell anemia, (figure 1 and fi 110 111<mark>gure 2</mark>).

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

	Parameters	Case (Mean±SD)	Control (Mean±SD)	P-value	
	Copper mg/l	0.512±0.290	0.923±0.214	0.000	
		<mark>(0.03-0.372)*</mark>	<mark>(0.474-1.01)*</mark>		
	Zinc <mark>mg/l</mark>	0.137±0.079	0.705±0.138	0.000	
		<mark>(0.122-1.154)*</mark>	<mark>(0.704-1.346)*</mark>		
115	P - values less than	0.05 were considered as	significant. * indicates	the range of values	
116	(lowest and highest) of copper and zinc.				
117	Table (2): Comparison the levels of copper and zinc in case group according to				
110	mandau				

118 gender.

Parameters Male (Mean±SD)	Female (Mean±SD)	P-value
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- 126 P-value less than 0.05 consider as significant:



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131 P-value less than 0.05 consider as significant:

#### 132 Discussion:

133 In sickle cell anaemia, there are deficiencies of some essential elements 134which are vital in maintenance of red cell integrity, body growth and 135 development. In the current study the levels of zinc and copper showed a significant decrease in patients with sickle cell anaemia, compared to healthy 136 137individuals (p value 0.000). This might have occurred due to chronic 138 haemolysis with subsequent loss of zinc from RBCs. Zinc deficiency can also be the result of the adverse effect of hydroxyurea which increase zinc 139 excretion. (11) 140

141 This finding was in agreement with results of some previous studies <sup>(12,13,14)</sup>

142 in Central Africa and Nigeria, which related zinc deficiency in sickle cell

143 disease to manifestations such as growth retardation, hypogonadism in males,

144 hyperammonemia, abnormal dark adaptation and cell mediated immune

145 disorder. Similarly, the biochemical evidence for zinc deficiency in patients

- 146 with SCA include low zinc concentrations in plasma, erythrocytes, hair,
- 147 lymphocytes and granulocytes.<sup>(15)</sup> This biochemical difference appears to be

148 due to various mechanisms such as chronic haemolysis, renal lost due to

149 repeated sickling leading to abnormal renal tubular reabsorption of zinc,

- 150 abnormal binding of zinc to tissue proteins, disturbed metabolism of zinc
- <sup>151</sup> metalloenzymes. <sup>(14)</sup> Also, the results revealed that there was non-significant

- 152 difference between males and females, in the levels of zinc and copper in
- 153 patients with sickle cell anaemia. Furthermore, there was negative correlation
- between levels of zinc and copper and age of the patients (r = -0.603, p = -0.603).
- 155 value= 0.000) and (r= -0.443 p- value= 0.004) respectively.

# 156 Conclusion:

157 The levels of blood zinc and copper were decreased in patients with sickle 158 cell anaemia, compared to healthy individuals. There was also negative 159 correlation between the levels of zinc and copper and zinc.

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