



## RESEARCH ARTICLE

### ASSESSMENT OF SICKLED ERYTHROCYTE DISEASE USING THERMOGRAPHY AND ENERGY EXCHANGE PROCESSES

#<sup>1</sup>Aweda M. A. <sup>1</sup>Edi A. A. and <sup>2</sup>Kehinde M. O.

<sup>1</sup>Department of Radiation Biology and Radiotherapy, College of Medicine/Lagos University Teaching Hospital, Idi-Araba. PMB 12003, Lagos.

<sup>2</sup>Department of Medicine, College of Medicine/Lagos University Teaching Hospital, Idi-Araba. PMB 12003, Lagos.

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#### ABSTRACT

Thermography for determination of mean weighted skin temperature and physiological energy exchange processes in sickled erythrocyte patients (SS) were studied in order to determine relevance in the management of the disease. The mean skin temperature for SS patients was  $35.45 \pm 0.402$  °C while for non-sickled erythrocyte subjects (AA), it was  $35.06 \pm 0.128$  °C (101.11 % of AA). The mean oxygen consumption rate ( $VO_2$ ) in SS patients ( $130.99 \pm 21.17$  ml.s<sup>-1</sup>) was much higher than AA subjects ( $50.88 \pm 4.69$  ml.s<sup>-1</sup>), (SS mean value was 257.45 % of AA). The mean metabolic energy exchange rate (M) in SS was  $2294.87 \pm 330.95$  Jh<sup>-1</sup> while in AA subjects, it was  $916.88 \pm 111.31$  Jh<sup>-1</sup> (ie. 250.29 % of AA). The mean evaporative heat loss (E) was  $10414.70 \pm 116.105$  Jh<sup>-1</sup> in SS patients and  $10566.90 \pm 36.45$  Jh<sup>-1</sup> for AA (98.56 % of AA). The mean convective rate of heat exchange (C) in SS was  $11044.60 \pm 425.52$  Jh<sup>-1</sup> while it was  $10646.20 \pm 134.92$  Jh<sup>-1</sup> in AA (103.74 % of AA). The radiative rate of heat exchange ( $R_r$ ) in SS was  $199175 \pm 11145.10$  Jh<sup>-1</sup> while it was  $134689 \pm 3533.54$  Jh<sup>-1</sup> for AA (147.88 % of AA). The total energy balance ( $\Delta H$ ) in SS was  $225491 \pm 12729.78$  Jh<sup>-1</sup> while it was  $155003 \pm 3638.68$  Jh<sup>-1</sup> for AA (145.48 % of AA). The results showed that sickled erythrocyte patients consume more oxygen and have much higher metabolic, radiative and total energy change rates than the AA subjects.

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## INTRODUCTION

The correlation of body temperature and diseases has been known for several decades, but due to recent inventions and new technologies, it is possible to use skin temperature as a convenient and effective diagnosis. The first documented application of thermography was for early preclinical diagnosis of breast cancer [1]. Infrared thermography is a non-contact, non-invasive tool which maps the skin temperature. Physiological temperature distribution depends on complex relationships between the skin, inner tissues, local vasculature and metabolic and hormonal activities. Hence, the use of thermography as a diagnostic tool is based on the fact that pathologies would raise skin temperature due to increased metabolic activities. Thermal images have been used to quantify sensitive changes in skin temperature in relation to certain diseases [2]. Blood flow can be assessed by many methods including the washout technique, laser Doppler flowmetry and medical infrared imaging. Of all these, infrared thermography has the advantage of being non-invasive, non-traumatic, fast, reliable, non-contact, capable of producing multiple recordings at short time intervals and absolutely safe for patients and physicians. Modern techniques allow the use of multispectral imaging in which the infected regions are well delineated.

SS disease conditions are commonly associated with regional vasodilation, hyperthermia, hyperperfusion, hypermetabolism, and hypervascularization which generate a higher-temperature heat source. The heat emanating from the skin and the surrounding blood flow can be quantified by using the Pennes' bio-heat equation [3],

$$k\Delta^2 T - c_b w_b (T - T_a) + q_m = 0$$

where  $k$  is the conductivity,  $q_m$  is volumetric metabolic rate of tissue,  $T$  is the unknown tissue temperature, and  $T_a$  is the arterial temperature. Energy exchange processes take place through conduction, convection and radiation while the physiological factors influencing the net heat

metabolic heat production, radiation balance, convective transfer via sensible and latent heat, conduction and the heat loss through respiration. The curved, crescent-shaped or sickle-shaped erythrocyte is a genetic disorder that strikes the black race far more than any other population group. It is caused by a defective gene and anaemia results from abnormal hemoglobin, the oxygen-carrying component of red blood cells. The disease is often accompanied by intense pain and serious deficiencies of oxygen and other blood nutrients throughout the body. Sickle cell infection may affect skin temperature as a result of energy changes. The disease is therefore of economic and social importance. The aim of this study is to investigate the correlation of skin temperature and physiological energy exchange processes in the management of sickled erythrocyte (SS) patients. We have employed the various standard methods in the Man-ENVIRONMENT heat EXCHANGE models as developed by Blazejczyk [4] and modified in 2005 [5] for the energy balance and transfer processes. These models and their applications are based on the first fundamental law of thermodynamics. We have correlated the thermographic information with physiological parameter modifications resulting from SS infection in order to apply the results for the disease management.

## MATERIALS AND METHOD

20 subjects were recruited for this study comprising of 10 SS patients and 10 non-sickled erythrocyte (AA) subjects as control. The patients were randomly selected from the outpatient clinics, Lagos State University Teaching Hospital (LUTH), Lagos, Nigeria, following the study approval granted by the Lagos University Teaching Hospital Ethics and Research Committee. The consent of the patients was obtained after a detailed explanation of the procedure and its potential benefits. The anthropometric data obtained from each subject were age, weight, height and body mass index (BMI) was determined. The blood pressure and oxygen consumption rate were also measured. The control subjects were randomly selected among the volunteers of the staff and students of the College of Medicine of the

ranged from 15 to 55 with a mean of 24 years, while those of the control ranged from 19 to 26 with a mean of 22.5 years. All the subjects were allowed to rest in the consulting room for about 15 min, where the relative humidity and room temperature were controlled in order to achieve equilibrium of the body with the environment. Not more than three persons at a time were allowed inside the room in order to minimize the parameters that might influence temperature while the other parameters that may influence thermographic imaging were fixed. The wall-mounted air-conditioning split unit provided the required stable temperature. The skin temperatures at some specific points on the patients were obtained using the FLIR digital infrared camera model INFRACAM 1124520 (from FLIR Systems, USA). The camera is a compact light weight focal plane array-based system with a temperature resolution of 0.1 °C. A high-resolution real time image was provided on the LCD colour display on which the skin temperature is also displayed. The mean skin temperatures at specific points on the patients were determined and used to calculate the various energy exchange rates.

Skin temperatures were taken at the forehead, neck, arms, hands, feet, legs, thighs and trunk. The values were used to calculate the mean weighted skin temperature ( $T_{sk}$ ) with the equation below according to Blazejezyk (1997) [6]

$$T_{sk} = 0.071t_{fh} + 0.14 t_a + 0.05 t_{ha} + 0.07 t_f + 0.13 t_l + 0.19 t_{th} + 0.35t_t \quad \dots\dots\dots 3$$

where  $t_{fh}$  = the skin temperature of the forehead

$t_a$  = the skin temperature of the arm

$t_{ha}$  = the skin temperature of the hand

$t_f$  = the skin temperature of the foot

$t_l$  = the skin temperature of the leg

$t_{th}$  = the skin temperature of the thigh

$t_t$  = the skin temperature of the trunk

The mean weighted skin temperature is used to calculate the radiative heat exchange R and the convective heat exchange C using the formulae

$$R = 6.6 (T_w - T_{sk}) \quad \dots\dots\dots 4$$

where  $T_w$  = mean radiant temperature (°C) of the surrounding walls and

$$C = 7.0 \times V_a \times 0.6 (t_a - T_{sk}) \quad \dots\dots\dots 5$$

where  $V_a$  = velocity of air (in m/s).

The measured oxygen consumption rate was used to calculate the metabolic heat gain, the methods of

conductive C, radiative R, respiratory  $R_{es}$  and total energy balance  $\Delta H$  have been describe elsewhere [7]. The results obtained for this study was analyzed statistically and summarized as in the table 2, and compared with those of AA participants using the paired sample student T- test [8,9]

## RESULTS AND DISCUSSION

Table 1 shows the anthropometric parameters of the patients from where the BMI was calculated. There were no significant differences in the patient (SS) parameters compared with control (AA). Typical thermographs of the studied SS patients are presented in figure 1, indicated on them are the skin temperatures at (a) abdomen, (b) forehead, (c) neck and (d) chest. The  $T_{sk}$  of the SS patients was only slightly higher than that of AA, and did not vary much from patient to patient as seen figure 2.  $T_{sk}$  for AA varied from 34.28 °C to 35.60 with a mean value of  $35.06 \pm 0.128$  °C. The values varied from 34.14 °C to a maximum of 37.64 °C in SS subjects, giving a mean value of  $35.45 \pm 0.402$  °C (101.11 % of AA). Variation in  $T_{sk}$  was not significant (p-value = 0.455). The slight differences in  $T_{sk}$  indicate that SS produce a little more heat than AA. This might be due to the presence of young the erythrocytes which use the metabolic energy to maintain their structural and functional integrity.

Figures 3 indicate that the average oxygen consumption rate ( $VO_2$ ) in SS is much higher than in AA. The  $VO_2$  among the AA subjects varied from 25.71 to 69.38  $ml.s^{-1}$  with a mean value of  $50.88 \pm 4.69$   $ml.s^{-1}$ . Those of SS varied from 74.81  $ml.s^{-1}$  to 276.43  $ml.s^{-1}$  with a mean much higher than that of AA ( $130.99 \pm 21.17$   $ml.s^{-1}$ , i.e. 257.45 % of AA).  $VO_2$  is much higher in SS subjects (p-value = 0.005). This could result from the abnormal conformational changes in the molecular structures of the sickled hemoglobin [10] which causes abnormal energy expenditure of erythropoietic hyperplasia when carrying oxygen [11].

The metabolic heat production rate (M) for AA varied from 450.42  $Jh^{-1}$  to 1666.76  $Jh^{-1}$  with a mean value of  $916.88 \pm 111.31$   $Jh^{-1}$  as depicted in figure

maximum of 4842.72 Jh<sup>-1</sup> in SS subjects, giving a mean value of 2294.87 ± 330.95 Jh<sup>-1</sup> (250.29 % of AA). The difference in the mean M values for the SS compared with AA is very significant with p-value = 0.007.

Figure 5 shows the evaporative heat production rate (E) for AA which varied from 10414.66 Jh<sup>-1</sup> to 10791.83 Jh<sup>-1</sup> with a mean value of 10566.90 ± 36.45 Jh<sup>-1</sup>. The values in SS subjects varied from 9777.49 Jh<sup>-1</sup> to a maximum of 10952.32 Jh<sup>-1</sup>,

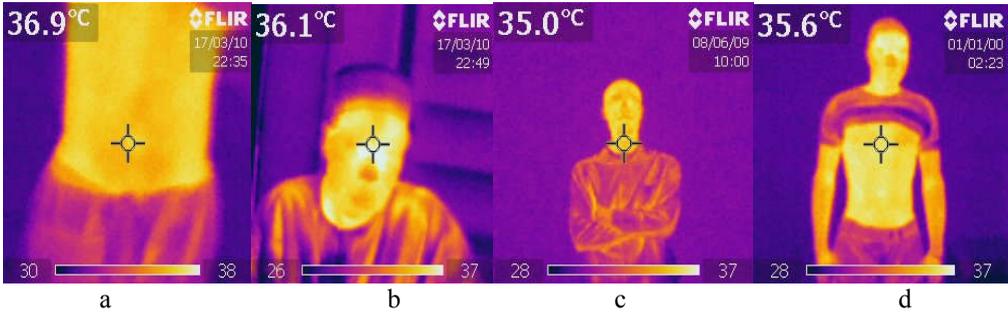


Figure 1 Typical thermographs of patients showing points of temperature measurement

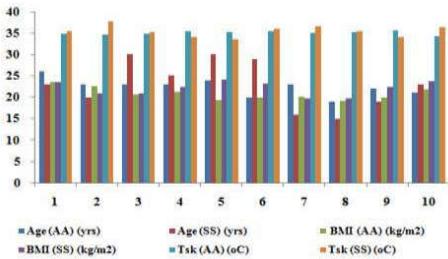


Figure 2. Mean weighted skin temperature (T<sub>sk</sub>) in SS compared with AA subjects

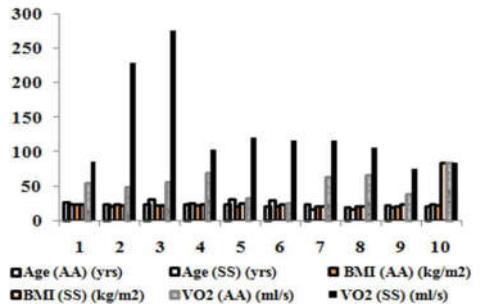


Figure 3. Mean oxygen consumption rates (VO<sub>2</sub>) in SS compared with AA subjects

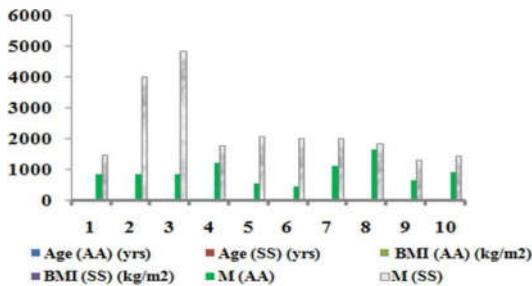


Figure 4. Mean metabolic energy change rates (M) in SS compared with AA subjects

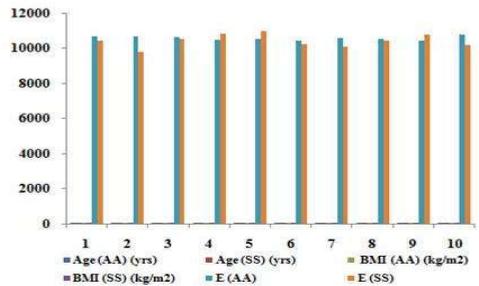


Figure 5. Mean evaporative energy change rates (E) in SS compared with AA subjects

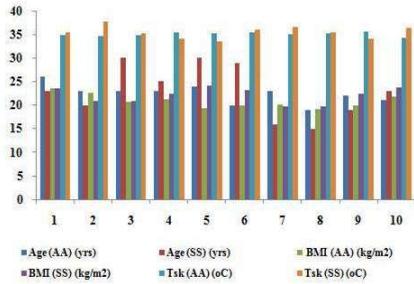


Figure 6. Mean convective energy change rates (C) in SS compared with AA subjects

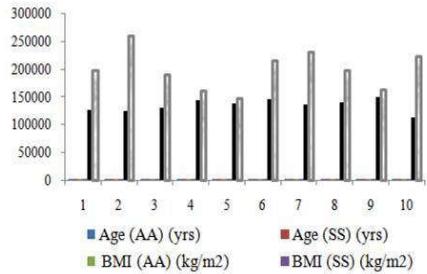


Figure 7. Mean Radiative energy change rates ( $L_r$ ) in SS compared with AA subjects

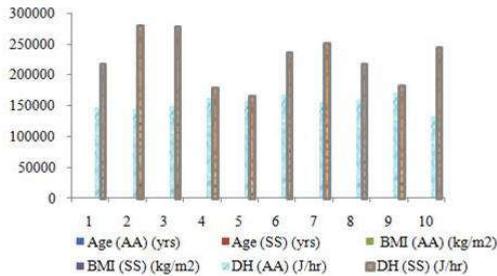


Figure 8. Mean total energy change rates ( $\Delta H$ ) in SS compared with AA subjects

**Table 1. Homozygous (AA) non- sickle cell anemia subjects as control (a) and Homozygous (SS) sickle cell anemia subjects (b)**

Ages (yr)	Weight (kg)	Height (cm)	Body mass index (kg/m <sup>2</sup> )	Body surface area (m <sup>2</sup> )	Ages (yr)	Weight (kg)	Height (cm)	Body mass index (kg/m <sup>2</sup> )	Body surface area (m <sup>2</sup> )
26	69	171	23.59	1.81	23	69	171	23.59	1.81
23	70	176	22.59	1.85	20	53	159	20.96	1.53
23	60	170	20.76	1.69	30	60.5	170	20.93	1.70
23	60	168	21.25	1.68	25	58	161	22.38	1.61
24	62	179	19.35	1.78	30	65	164	24.17	1.71
20	55	166	19.96	1.61	29	67	170	23.18	1.78
23	70	178	20.09	1.87	16	50.3	160	19.65	1.51
19	45	153	19.22	1.39	15	50	159	19.78	1.49
22	54	165	19.84	1.59	19	58	161	22.38	1.61
21	53	156	21.78	1.51	23	63	158	25.24	1.64

a

b

**Table 2. Summary of the heat exchange parameters of homozygous sickle (SS) compared with non-sickle (AA) cell subjects**

Physiological parameters	Non-sickle cell (AA) (Mean $\pm$ SEM)	Homozygous sickle cell (SS) Mean $\pm$ SEM	p- value
Oxygen consumption rate ( $\text{VO}_2$ ) (ml/s)	50.88 $\pm$ 4.69	130.99 $\pm$ 21.17	0.005
Metabolic heat production (M) (J/h)	916.88 $\pm$ 111.31	2294.87 $\pm$ 330.95	0.007
Mean skin temperature ( $T_{\text{sk}}$ ) ( $^{\circ}\text{C}$ )	35.06 $\pm$ 0.128	35.45 $\pm$ 0.402	0.455
Evaporative heat loss (E) (J/h)	-10566.90 $\pm$ 36.45	-10414.70 $\pm$ 116.105	0.300
Convective heat exchange (C) (J/h)	-10646.20 $\pm$ 134.92	-11044.60 $\pm$ 425.52	0.455
Long wave radiative heat exchange ( $L_r$ ) (J/h)	-134689 $\pm$ 3533.54	-199175 $\pm$ 11145.10	0.001
Respiratory heat loss ( $R_{\text{es}}$ ) (J/h)	-22.54 $\pm$ 0.0	-22.54 $\pm$ 0.0	N/A
Total energy production rate ( $\Delta\text{H}$ ) (J/h)	155003 $\pm$ 3638.68	225491 $\pm$ 12729.78	0.001

giving a mean value of  $10414.70 \pm 116.105 \text{ Jh}^{-1}$  (98.56 % of AA). The p-value for the variation was 0.300. The convective heat production rate (C) for AA varied from  $9821.95 \text{ Jh}^{-1}$  to  $11219.04 \text{ Jh}^{-1}$  with a mean value of  $10646.20 \pm 134.92 \text{ Jh}^{-1}$ . In figure 6, the values varied from  $9070.49 \text{ Jh}^{-1}$  to a maximum of  $13378.18 \text{ Jh}^{-1}$  in SS subjects, giving a mean value of  $11044.60 \pm 425.52 \text{ Jh}^{-1}$  (103.74 % of AA). Similar to  $T_{\text{sk}}$ , the variation was not significant; p-value was 0.455. Lee [12] in his theoretical model which assumed a wind velocity of  $0.2 \text{ m.s}^{-1}$ , predicted the ratio of convective to radiative heat loss as 1: 0.7 at a temperature of  $20^{\circ}\text{C}$ . Zhang et al. [13] have also published the work of simulation of Convective and Respiratory heat losses. The different methods developed to evaluate the heat exchange have recently been described by Buchlin [14]. de Dear et al. [15] measured radiative and convective heat loss rates and found the rates to be  $16,200 \text{ Jh}^{-1}$  and  $11880 \text{ Jh}^{-1}$  respectively. Even in the absence of wind, convection is an important mechanism in cooling human body. Convection accounts for about 33 % of the thermal loss of the human body in cool and still air. As other thermal loss mechanisms like forced convection and perspiration become more significant, radiation becomes less significant in human thermal balance [16]. The long wave radiative energy production rate ( $L_r$ ) for AA varied from  $113097.60 \text{ Jh}^{-1}$  to  $149688.00 \text{ Jh}^{-1}$  with a mean value of  $134689 \pm$

SS subjects, giving a mean value of  $199175 \pm 11145.10 \text{ Jh}^{-1}$  (147.88 % of AA). This variation was significant with a p-value of 0.001. The respiratory heat loss ( $R_{\text{es}}$ ) for AA did not vary from patient to patient. There was no difference in the value for both SS and AA, and the determined value was  $22.54 \text{ Jh}^{-1}$ . The total energy production rate ( $\Delta\text{H}$ ) for AA varied from  $132802.26 \text{ Jh}^{-1}$  to  $170680.35 \text{ Jh}^{-1}$  with a mean value of  $155003 \pm 3638.68 \text{ Jh}^{-1}$ . Figure 8 shows that the values varied from  $165415.75 \text{ Jh}^{-1}$  to a maximum of  $279456.86 \text{ Jh}^{-1}$  in SS subjects, giving a mean value of  $225491 \pm 12729.78 \text{ Jh}^{-1}$  (145.48 % of AA). The variation here was significant with p-value = 0.001. Numerous energy balance and anthropometric studies have been reported in the literature. Some of these investigated the interrelationship between energy expenditure and energy intake as they affect development of cancers [17-21]. Deficiency in energy balance is an indication of risk of developing diseases.

## CONCLUSION

The result of this study shows that the mean weighted skin temperature derived from thermography may be used to quantify various energy changes and hence for determining the variations in the different energy exchange rates between SS patients and AA subjects.

thermographicly provided the required parameter for calculating the various energy exchange rates. Significant differences in the  $\dot{V}O_2$ ,  $M$ ,  $L_r$  and  $\Delta H$  values are useful in delineating patients with sickled erythrocyte from others. Age, sex and BMI seem not to play significant role in this SS assessment method. This method provides a quick and non-invasive method of assessing the status of SS individuals based on the differences in the energy exchange rates.

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