Exploring the Effects of Glutathione Supplementation on Liver Enzymes and Serum

Electrolyte Levels in Mice Exposed to 850-1900 MHz Mobile Phone Radiation: An

Investigation into Radiative Stress and Potential Protective Measures

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ABSTRACT

The pervasive presence of electromagnetic fields (EMF) generated by modern technologies

poses a significant threat to human health, with ionizing radiation, a byproduct of EMF,

potentially contributing to cancer development. This study explores the impact of chronic

exposure to GSM-EMFs and 900-1800 MHz-induced electromagnetic radiation (EMR) on liver

enzymes, and serum electrolytes in mice and potential mitigating effect of exogenous glutathione

administration. Thirty-five adult male mice were randomly divided into seven groups and

exposed to various modes of mobile phone radiation for five weeks, with or without glutathione

administration. Liver enzymes (ALP, ALT, AST) and serum electrolytes (sodium, potassium,

bicarbonate) were analyzed. Results showed significant increases in ALP levels in the Silent,

Ringtone + GSH, and Silent + GSH groups compared to controls, while ALT and AST levels

remained largely unchanged. Serum electrolyte concentrations did not significantly differ across

experimental groups.

Keywords: Mobile phone radiation, Glutathione, Liver enzymes, Serum electrolytes, Physiology

INTRODUCTION

The widespread presence of electromagnetic fields (EMF) generated by contemporary

technologies, encompassing common household appliances and mobile devices, presents a

palpable threat to human health. Research indicates that ionizing radiation, a byproduct of EMF,

may serve as a precursor to future cancer development, as evidenced by studies linking elevated

rates of chromosome abnormalities and micronuclei occurrence in populations exposed to

radiation from various sources such as nuclear incidents, environmental contaminants, and

medical diagnostic procedures like CT scans and x-rays [1]. Such exposure to radiation triggers

oxidative stress (OS) within bodily tissues [2], emphasize the potential health consequences

associated with prolonged EMF exposure. This phenomenon arises from an imbalance between

the levels of reactive oxygen species (ROS) and the capacity of the antioxidant system to

mitigate their effects [3]. Moreover, oxidative stress levels can escalate during different stages of

development [4]. Various physiopathological and physiological conditions, nutritional factors

[7], and exposure to external pollutants like electromagnetic radiation [5] can contribute to

heightened oxidative stress status.

Moreover, studies highlight that electromagnetic radiation intensifies the production of ROS,

culminating in various physiological repercussions, such as DNA damage. Ionizing radiation,

with its rapid ability to ionize cellular structures, presents a significant peril to both human and

animal well-being, capable of inflicting harm in an instant. Conversely, non-ionizing radiation

from device such as mobile phones, although lacking ionizing capabilities, can still pose risks

with prolonged exposure, a critical consideration in its interaction with the human body [8].

Many researchers concur that non-ionizing radiation poses significant health hazards [9]. The

potential health impacts of electromagnetic pollution and the increasing recognition of EMF

exposure spotlight the need for heightened awareness [10].

Non-ionizing radiation spans from extremely low frequency (ELF) to visible light, while

ionizing radiation encompasses higher frequencies [11]. These radiations primarily impact the

liver, kidneys, and brain due to their proximity to mobile phones during daily use [2]. The

increasing use of multi-slice CTs also further amplifies concerns, especially for children, who are

at a higher risk of radiation-induced malignancies due to their longer lifespan [12]. Additionally,

research reveals the potential adverse effects of electromagnetic frequencies emitted by mobile

phones, including DNA damage, increased cancer susceptibility, oxidative stress, chromosomal

abnormalities [13, 14, 15, 16, 17, 18], and hormonal and physiological imbalances.

The liver, the body's largest internal organ, plays a pivotal role in detoxification, ridding the

body of harmful toxins [19]. Yet, the detrimental effects of free radicals generated during hepatic

metabolism can largely be counteracted by various antioxidant enzymes and non-enzymatic

mechanisms [20, 3]. Nevertheless, a decline in the antioxidant system's response, coupled with

an overproduction of reactive oxygen species from radiation, may escalate hepatic oxidative

stress, leading to liver injury. Electromagnetic radiation emitted by 900 MHz cell phones has

been shown to impact the expression of Nrf2 protein, trigger oxidative damage, and alter the

morphology of liver cells [21].

Electrolytes are electrically charged molecules that play pivotal roles in regulating acid-base

balance, blood clotting, and the contraction of muscles and body fluids. Within the body, several

common electrolytes serve specific and vital functions, collectively maintaining the balance of

fluids between intracellular and extracellular environments. This equilibrium is crucial for

hydration, nerve impulses, muscle function, and pH levels. Among the major electrolytes are

Potassium, Sodium, Magnesium, Calcium, and Chloride. Notably, Serum Sodium, Potassium, and Chloride are fundamental determinants of the electrophysiological properties of myocardial membranes [22, 23]. In the body, serum electrolytes perform essential roles in numerous physiological and metabolic processes. Particularly among acutely ill individuals, including children, adolescents, and animals, fluctuations in serum sodium and potassium concentrations are prevalent, often leading to heightened morbidity and mortality [25]. Serum electrolyte concentrations constitute key tests for evaluating a patient's clinical condition and are closely linked to morbidity and mortality. Any deviations from the normal range of electrolyte levels in the body are termed electrolyte disorders [26]. Despite their critical importance, electrolyte disorders in the context of electromagnetic field (EMF) exposure, encompassing both ionizing and non-ionizing radiation, have received scant attention. However, these disorders hold significant potential in influencing the initiation, diagnosis, and prognosis of diseases [24].

As technology becomes increasingly integrated into our daily lives, there's a notable gap in research concerning the effects of radiation on liver enzymes and electrolyte balance, both of which are closely linked to elevated mortality and morbidity rates. Specifically, there's a lack of understanding in addressing and managing the potential health implications associated with these critical bodily parameters. Therefore, this study dig into investigating the repercussions of electromagnetic radiation exposure on liver enzymes and serum electrolytes.

The research focuses on exploring the impact of daily exposure to GSM-EMFs and 900-1800 MHz-induced electromagnetic radiation (EMR) on oxidative stress levels in mice. By examining these specific frequencies commonly encountered in modern technology, the study aims to shed light on the potential physiological consequences and implications for human health.

METHODS

Animals and grouping

The animals involved in the study were maintained and used in accordance with the Animal

Welfare Act and the Guide for the Care and Use of Laboratory animals prepared by the Ahmadu

Bello University Committee on Animal Use and Care (ABUCAUC), and ethical approval was

obtained from the ABUCAUC with the approval number of ABUCAUC/2021/005.

At the start of the experiment, thirty five (35) apparently healthy adult male Mice between the

ages of 8-12 weeks were obtained from the animal house facility of the Department of Human

Physiology, College of Medical Sciences, Ahmadu Bello University, Zaria, Nigeria. They were

housed in standard polypropylene cages and were maintained and allowed free access to feed and

water 24/7 and 12h alternate day light and darkness. The animals were allowed to acclimatize to

the environment of the behavioral laboratory for two weeks before the commencement of the

experiment.

The animals were randomly selected and divided into seven (7) groups of five (n=5) mice per

group as shown in the table below:

Table 1: Showing different irradiated group

GROUP	CATEGORY
I	Normal control group
II	Group exposed to mobile phone radiation in ringtone mode
III	Group exposed to mobile phone radiation in vibration mode
IV	Group exposed to mobile phone radiation in silent mode
V	Group exposed to mobile phone radiation in ringtone mode plus glutathione (GSH)
VI	Group exposed to mobile phone radiation in vibration mode plus glutathione and the last group
VII	Group VII were exposed to mobile phone radiation in silent mode plus glutathione

Radiation exposure

Using Itel mobile phone (it2160) with specific absorption rate (SAR) of 2W/kg and 2G Network

Bands, of frequency 850-1900 MHz, the animals in groups II to VII were all exposed to 4 hours

mobile phone radiation of an average of 300 missed calls per day for six (6) weeks. Animals in

groups V, VI and VII were administered glutathione (250 mg/kg) thirty minutes prior to mobile

phone radiation daily.

Determination of Alkaline Phospahtase

Alkaline phosphatase (ALP) activity was analyzed using the method described by Bassey *et al.*

(1946) [47] and modified by Wright *et al.* (1972) [46], with Randox kits. In this procedure, 10 µl

of the sample was mixed with 500 µl of the ALP reagent in a cuvette. The initial absorbance at

405 nm was recorded, and additional readings were taken over 3 minutes. The mean absorbance

per minute was then used to calculate ALP activity (IU/l) using the formula $2742 \times \Delta A$ 405

nm/min, where 2742 is the extinction coefficient, and ΔA 405 nm/min represents the change in

absorbance per minute for the sample.

Determination of Alanine Transaminase

Alanine transaminase (ALT) activity was measured using the method described by Shaw *et al.*

(1978) [45] with Randox kits. In this process, 50 µl of the sample and 500 µl of the ALT reagent

were mixed in a test tube. The initial absorbance at 340 nm was recorded after 1 minute, and the

timer was started simultaneously. Further absorbance readings were taken at 1, 2, and 3 minutes.

ALT activity (nm/min) was calculated using the formula $1746 \times \Delta A$ 340 nm/min, where ΔA 340

nm/min is the change in absorbance per minute for the sample, and 1746 is the extinction

coefficient.

Determination of Aspartate Aminotransferase

Aspartate transaminase (AST) activity was also measured using the method described by Shaw

et al. (1978) [45] with Randox kits. In this procedure, 50 µl of the sample and 500 µl of the AST

reagent were combined in a test tube. The initial absorbance at 340 nm was recorded after 1

minute. The timer was then started, and additional absorbance readings were taken at 1, 2, and 3

minutes. AST activity (nm/min) was calculated using the formula $1746 \times \Delta A$ 340 nm/min, where

ΔA 340 nm/min represents the change in absorbance per minute for the sample, and 1746 is the

extinction coefficient.

Determination of Sodium and Potassium ion concentration

Serum sodium and potassium concentrations were measured using the flame emission

photometry method described by Magoshes and Vallee in 1956 [49]. This technique involves the

desolvation of a solution containing these elements by a flame, which leaves behind solid salts.

These salts dissociate into neutral ground state atoms, which then become excited by the flame,

moving to a higher energy state. As the excited atoms return to the ground state, they emit light

at characteristic wavelengths (590 nm for sodium and 770 nm for potassium). This emitted light

passes through a suitable filter onto a photosensitive element, and the resulting current produced

is measured, corresponding to the amount of sodium or potassium in the original sample.

Determination of Bicarbonate Concentration

The concentration of bicarbonate ions in serum samples was determined using the titration

method described by Van Slyke and Neil in 1924 [48]. The method involves releasing carbon

174   dioxide from bicarbonate ions in the serum by adding dilute hydrochloric acid. The remaining

175   acid is then titrated with sodium hydroxide, using phenol red as an indicator.

RESULTS

LIVER ENZYMES

Serum Alkaline Phosphate (ALP)

The impact of chronic exposure to mobile phone radiation, coupled with glutathione

administration, on serum liver enzyme (ALP) levels in mice was examined, yielding the

following outcomes [Figure 1]: Control (25.97 ±3.09), Ringtone (23.37±2.04), Vibration

(35.76±4.95), Silent (141.13±12.87), Ringtone + GSH (71.09±6.38), Vibration + GSH (39.09±

6.13), Silent + GSH (80.73±17.21). Statistical analysis revealed no significant differences

($p>0.05$) between the control group and other experimental groups, except for the Silent,

Ringtone + GSH, and Silent + GSH groups, which exhibited notable increases compared to the

control group.

Figure 1: Effect of chronic exposure to mobile phone radiation and glutathione administration

on serum alkaline phosphate in mice.

Serum Alanine Aminotransferase (ALT)

The impact of chronic exposure to mobile phone radiation, alongside glutathione administration,

on serum liver enzyme levels (ALT) in mice was investigated, yielding the following outcomes

[Figure 2]: Control (111.0±10.47), Ringtone (114.5±15.57), Vibration (121.0±10.08), Silent

(124.25±11.0), Ringtone + GSH (145.5±12.10), Vibration + GSH (148.75±23.57), Silent + GSH

(69.5±5.7). Statistical analysis revealed no significant differences (p>0.05) between the control

group and the other experimental groups, except for the Ringtone + GSH and Vibration + GSH

groups, which demonstrated a notable increase.

Figure 2: Effect of Chronic Exposure to Mobile Phone Radiation and Glutathione

Administration on Serum Alanine aminotransferase in mice.

Serum Aspartate Aminotransferase (AST)

The effect of chronic exposure to mobile phone radiation and glutathione administration on

serum liver enzymes (AST) in mice gave the following results [Figure 3]; Control

(262.25±17.79), ringtone (221.0±11.08), vibration (234.25±31.497), silent (217.5±24.75),

ringtone + GSH (272.75±23.62), vibration + GSH (392.25±58.41), silent + GSH (227.5±28.0).

There was no statistical significance (p>0.05) when control group was compared to other groups.

But when the other groups are compared to each other there was statistical significance as shown

on the graph.

The impact of chronic exposure to mobile phone radiation, coupled with glutathione

administration, on serum liver enzyme levels (AST) in mice was explored, yielding the following

observations [Figure 3]: Control (262.25±17.79), Ringtone (221.0±11.08), Vibration

(234.25±31.497), Silent (217.5±24.75), Ringtone + GSH (272.75±23.62), Vibration + GSH

(392.25±58.41), Silent + GSH (227.5±28.0). Statistical analysis revealed no significant

differences (p>0.05) when comparing the control group to the other experimental groups except

the vibration+GSH group.

Figure 3: Effect of Chronic Exposure to Mobile Phone Radiation and Glutathione

Administration on Serum Aspartate aminotransferase in Mice.

Table 2: Detailed the difference of chronic exposure to mobile phone radiation and glutathione administration on notable liver enzymes.

Liver Enzymes	Control	Ringtone	Vibration	Silent	Ringtone+GSH	Vibration+GSH	Silent+GSH
Alkaline Phosphate	25.97 ±3.09	23.37±2.04	35.76±4.95	**141.13±12.87**	**71.09±6.38**	39.09± 6.13	**80.73±17.21**
Alanine Aminotransferase	111.0±10.47	114.5±15.57	121.0±10.08	124.25±11.0	**145.5±12.10**	**148.75±23.57**	69.5±5.7
Aspartate Aminotransferase	262.25±17.79	221.0±11.08	234.25±31.497	217.5±24.75	272.75±23.62	**392.25±58.41**	227.5±28.0

SERUM ELECTROLYTES

Serum Sodium ion Concentration

The impact of chronic exposure to mobile phone radiation, alongside glutathione administration, on serum sodium ion concentration was investigated, yielding the following outcomes [Figure 4]: Control (103.0 ± 4.26), Ringtone (104.94 ± 3.83), Vibration (111.28 ± 2.87), Silent (108.30 ± 2.19), Ringtone + GSH (109.60 ± 2.79), Vibration + GSH (103.10 ± 2.79), Silent + GSH (105.0 ± 3.88). Upon analysis, no significant differences were observed ($p > 0.05$) when comparing the control group with the entire experimental cohort. These findings suggest that chronic exposure to mobile phone radiation, in conjunction with glutathione administration, does not exert a substantial influence on serum sodium ion concentration. Further exploration is warranted to comprehensively understand the interplay between these factors and their potential implications for health.

Figure 4: Effect of Chronic Exposure to Mobile Radiation and Glutathione Administration on Serum Sodium Ion Concentration in Mice.

Serum Potassium ion Concentration

The impact of chronic exposure to mobile phone radiation, coupled with glutathione administration, on serum potassium ion concentration was explored, yielding the following observations [Figure 5]: Control (36.40 ± 4.26), Ringtone (34.46 ± 3.83), Vibration (28.12 ± 2.87), Silent (31.10 ± 2.19), Ringtone + GSH (29.74 ± 2.79), Vibration + GSH (36.30 ± 2.79), Silent + GSH (34.40 ± 3.88). Result revealed no statistically significant differences ($p > 0.05$) when comparing the control group with the various experimental conditions, including Ringtone mode, Silent Mode, Vibration mode, as well as their corresponding combinations with glutathione administration. These findings suggest that chronic exposure to mobile phone radiation, in tandem with glutathione administration, does not significantly influence serum potassium ion concentration. Further investigations are warranted to elucidate the nuanced interplay between these variables and their potential implications for health.

Figure 5: Effect of Chronic Exposure to Mobile Radiation and Glutathione Administration on

Serum Potassium Ion Concentration in Mice

Serum Bicarbonate Concentration

The impact of chronic exposure to mobile phone radiation, alongside glutathione administration,

on serum bicarbonate concentration was assessed, yielding the following findings [Figure 6]:

Control (24.82 ± 3.60), Ringtone (24.78 ± 3.31), Vibration (19.08 ± 2.47), Silent (24.18 ± 2.29),

Ringtone + GSH (23.32 ± 3.94), Vibration + GSH (21.98 ± 4.38), Silent + GSH (27.42 ± 8.18).

Statistical analysis revealed no significant differences ($p > 0.05$) when comparing the control

group with the various experimental conditions. These results suggest that chronic exposure to

mobile phone radiation, in conjunction with glutathione administration, does not notably alter

serum bicarbonate concentration. However, further investigations are warranted to

comprehensively understand the intricate relationship between these variables and their potential

implications for physiological health.

Figure 6: Effect of Chronic Exposure to Mobile Radiation and Glutathione Administration on Serum Bicarbonate Ion Concentration in Mice

Table 3: Detailed the difference of chronic exposure to mobile phone radiation and glutathione administration on different serum electrolytes.

Serum Electrolytes	Control	Ringtone	Vibration	Silent	Ringtone+GSH	Vibration+GSH	Silent+GSH
Na+	103.0 ± 4.26	104.94 ± 3.83	111.28 ± 2.87	108.30 ± 2.19	109.60 ± 2.79	103.10 ± 2.79	105.0 ± 3.88
K+	36.40 ± 4.26	34.46 ± 3.83	28.12 ± 2.87	31.10 ± 2.19	29.74 ± 2.79	36.30 ± 2.79	34.40 ± 3.88
$NaHCO_3$	24.82 ± 3.60	24.78 ± 3.31	19.08 ± 2.47	24.18 ± 2.29	23.32 ± 3.94	21.98 ± 4.38	27.42 ± 8.18

DISCUSSION

The utilization of mobile phones stands as one of the most rapidly advancing technological

fronts, prompting concerns regarding the interplay between electromagnetic radiation (EMR) and

liver function [27]. Amidst the different applications of low-intensity, extremely low-frequency

electromagnetic fields (EMF) in combatting ailments like malaria and cancer [28], the utilization

of radiofrequency radiation (RFR) is emerging as a promising noninvasive strategy for tumor

therapy, capitalizing on its ability to induce shock responses and apoptosis in human cancer cells

[29]. Nonetheless, findings by Costa et al. [31] suggest that even minute levels of EMF,

modulated at specific frequencies, might disrupt the growth of Hep G2 cells. However, their

study delves into the in vitro mechanisms elucidating the antiproliferative impact of such low-

intensity EMF on Hep G2 cells [30, 31]. Moreover, various studies have explored the

repercussions of EMF exposure on the delicate balance of antioxidants and oxidants in vital

organs and tissues such as the liver, heart, and lens [32, 33, 34]. Despite this wealth of research,

scant literature exists on the influence of EMF radiation on physiological parameters exposure

[35] (Unpublished report) such as serum electrolytes and liver enzymes, particularly regarding

the interplay with glutathione.

A previous study conducted by Kosanic *et al.* (2001) [38] reported the effects of vitamin C, a

potent antioxidant, on individuals exposed to Wi-Fi radiation for one day. Their findings

unveiled a significant rise in blood glucose levels and a concurrent decrease in triglyceride levels

compared to the control group. However, no notable impact was observed on cholesterol levels,

low-density lipoprotein (LDL), high-density lipoprotein (HDL), or hepatic enzyme activities

such as ALP, AST, and ALT relative to the control group. Interestingly, in the extended 5-day

tests, contrasting outcomes emerged, showcasing a significant reduction in blood sugar levels

and an elevation in cholesterol and lipoproteins among various groups compared to the control.

Nevertheless, triglyceride levels and hepatic enzyme activities remained unaffected.

This discrepancy suggests that the observed effects could potentially stem from the antioxidant

properties of vitamin C. In our study, where we investigated the influence of glutathione (GSH),

a compound closely related to vitamin C in its role as an antioxidant, on radiation exposure from

different modes of phone activity, distinct patterns were noted. Notably, a significant increase in

ALP was only noted in the Silent, Ringtone+GSH, and Silent+GSH modes, while other modes

exhibited no significant increase, as detailed in [Table 2].

Despite GSH's known physiological and metabolic roles and its capacity to shield against

radiation-induced damage by scavenging free radicals and converting them into non-radical

forms [36, 37], our findings suggest a potential surge in oxidative stress. This could arise from an

upsurge in free radical production, a decline in antioxidant defense systems, or a combination of

both factors. The lack of significant effects observed in ALP, ALT, and AST across most phone

modes may be attributed to prolonged exposure to mobile phone radiation, which has been

linked to increased serum ALT and AST levels, as reported by Li (2015) and Rahman (2022)

[5,44]. However, insights from Fatima *et al*. (2017) [40] and colleagues shed further light,

revealing that rats exposed to radiation over a two-month period, three days per week, exhibited

heightened liver enzyme activity, elevated levels of MDA and H202, and decreased antioxidant

levels such as GSH and CAT in liver tissue. While this aligns partially with our findings, it's

worth noting differences in exposure duration between our study and theirs.

Our findings also echo the conclusions drawn by Nersesova *et al*. (2014) [41], who elucidated

that despite observing changes in enzyme activity levels of liver-sensitive enzymes in both liver

tissue and blood serum upon radiation exposure, no discernible effects on hepatocyte cell

membrane permeability were evident. Instead, they noted alterations in energetic metabolism.

However, despite these observations, correlational analysis of post-radiation enzyme activity

level changes failed to reveal a clear relationship among them. Our study aligns with the findings

of Ma *et al.* (2014) [42], who reported no significant changes in serum levels of ALT, AST, and

AST/ALT in the radiated group. Notably, they observed hepatocyte nuclear atrophy and

cytoplasmic vacuolar degeneration in histological sections, which was not explored in our study.

Additionally, in Cetin *et al.*'s [43] investigation on brain and liver glutathione peroxidase

activities, along with liver vitamin A and β-carotene concentrations, they noted a decrease in

glutathione peroxidase levels in the EMF radiated groups. Conversely, brain iron, vitamin A, and

β-carotene concentrations increased in the EMR groups, a result consistent with some outcomes

from our phone model assessments on glutathione's role in mitigating EMF radiation effects on

liver enzymes.

Furthermore, our results share similarities with the findings of Nwokocha *et al*. (2012) [44]. In

their study, they observed that AST liver enzymes elevated with increasing radiation, which is

also observed in some certain phone models in our study, such as vibration+GSH, AST liver

enzyme levels elevated with increasing radiation, while others within the same category did not

exhibit statistical significance in our study. Similarly, they found that alanine transaminase

(ALT) levels increased with radiation, which is also similar with our ringtone+GSH and

vibration+GSH models, while others in this same category showed no significant changes.

Notably, our study also identified an increase in AST liver enzyme levels, particularly in the

vibration+GSH model, an aspect not explored in their research. However, the remaining phone

models in our study, including ringtone, vibration, silent, ringtone+GSH, and silent+GSH, did

not show any significant increase.

Almost similar to our findings, Nwokocha *et al.* (2012) [44] reported that serum electrolyte

concentrations remained within normal ranges following total body irradiation, this is also to

similar to our results which is detailed in [Table 3]. Specifically, sodium, potassium, and

bicarbonate levels were found to be statistically not significant ($p>0.05$) lower than the control

group after irradiation.

In conclusion, this study sheds light on the impact of chronic mobile phone radiation exposure on

liver enzymes and serum electrolytes in a Mice model. While some variations were observed in

liver enzyme levels, the overall findings suggest minimal alterations in serum electrolyte

concentrations. These results reveal the miscellaneous nature of the biological response to

electromagnetic radiation and emphasize the importance of continued research in this field to

safeguard public health exposure to mobile phone radiation..

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ETHICAL STATEMENT

This study was conducted at the Department of Human Physiology, Faculty of Basic Medical

Sciences of Ahmadu Bello University, Zaria. Approval for the project was granted by the

Ahmadu Bello University Ethical Committee on Animal Use and Care, with the assigned

approval number ABUCAUC/2021/005. All ethical protocols concerning the handling, care, and

welfare of the animals were meticulously observed throughout the study, from its inception to its

conclusion.

AUTHOR'S CONTRIBUTIONS

All authors contributed greatly to the success of this work

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CONFLICT OF INTEREST

No conflict of interest to declare in this manuscript

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