# Effect of Ginger on Oxidative Stress Induced by Hydrogen Peroxide in Male Quail

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**Abstract.** This study was conducted to investigate the effect of Ginger (*Zingiber officinale*) as an antioxidant and its ability to prevent the effect of H<sub>2</sub>O<sub>2</sub>-induced oxidative stress and its reflections on blood parameters and antioxidant status of male quail. 120 local male quail on 21 days aged were randomly distributed according to complete randomized design into 4 groups and reared until 56 days age as follows: Control group, H<sub>2</sub>O<sub>2</sub> group (tap water with 0.5% H<sub>2</sub>O<sub>2</sub>), Ginger group (1000 mg ginger/kg ration), H<sub>2</sub>O<sub>2</sub> with Ginger group (1000 mg ginger/kg ration and tap water with 0.5% H<sub>2</sub>O<sub>2</sub>). Results revealed that the H<sub>2</sub>O<sub>2</sub> induced oxidative stress retracted the blood parameters and serum antioxidant status (decrease glutathione 1.10 µmol/1 and increase malondialdehyde 1.79 nmol/ml), while the ginger enhanced the blood parameters and serum antioxidants status (elevation glutathione 2.92 µmol/1 and decline in malondialdehyde 0.41 nmol/ml) compared to the control group (1.58 µmol/1, 0.74 nmol/ml respectively). In conclusion, ginger prevented the oxidative stress effects and returned most of the biochemical values to normal values as in the control group.

Keywords: Zingiber officinale, H<sub>2</sub>O<sub>2</sub>, Malondialdehyde, Glutathione, Coturnix coturnix.

# **1** Introduction

Oxidative stress is defined as an imbalance between the production of reactive oxygen species (ROS) and the ability of defensive biological systems (body antioxidants) to remove them or repair their destructive effects [1]. Therefore, the superior production of reactive oxygen species and their free radicals leads to oxidation of poly-unsaturated fatty acids (PUFA) in cellular membranes, thus it disturbs membrane permeability due to lipid peroxidation [2], and formation of the Malondialdehyde (MDA), which is one of its most important products, and it's used as an indicator of cellular membranes damage. The antioxidant systems in the cell work to reduce/or remove the oxidative damage by scavenging those free radicals or inhibiting the process of lipid peroxidation [3].

Many plants has an antioxidant capacity that prevents or reduce the oxidative stress. Ginger (*Zingiber officinale*) is one of these plants that has antioxidant capacity [4]. The antioxidant effect of ginger is due to its active ingredient as camphene, shogaols, zingiberol, and vitamin C and E [5].

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Shanoon [6] has indicated that ginger has an antioxidant effect similar to/ or greater than vitamin C or E. Al-kattan *et al.* [7] also indicated that ginger preserves the cells that suffer from oxidative stress whether it is pathologically or chemically induced by  $H_2O_2$  with drinking water.

One of the reasons for choosing males in the study is that the males are usually subjected to high level of oxidative stress during spermatogenesis due to the high levels of polyunsaturated fatty acids of spermatozoa membrane [19, 27].

The present study aimed to evaluate the effect of crushed dry ginger roots (*Zingiber officinale*) role as an antioxidant, and its preventive ability against the hydrogen peroxide-induced oxidative stress.

# 2 Materials and Methods

This study was conducted on 120 male quail (*Coturnix coturnix*) for the period 21 to 56 days of age. The requirements of lighting, ventilation and temperature were taken into consideration according to the bird's age. The birds were distributed randomly into four groups (30 males/group) in ground cages with dimensions 2.0 x 2.5 x 3.0 m for length, width and height, respectively. Ginger roots (*Zingiber officinale*) were cleaned, dried and crushed, then it was mixed with the ration at a concentration of 1000 mg/kg, that was formed according to the poultry nutrition requirement [8] and which included a starter ration (24.6% crude protein and 2838 kcal/kg of energy). Water and ration were allowed *ad libitum* throughout the study. The study treatments were as follows:

- T<sub>1</sub>: Control group: birds were reared on standard ration and tap water.
- T<sub>2</sub>: Hydrogen peroxide group ( $H_2O_2$ ): birds were reared on a standard ration and tap water with 0.5% hydrogen peroxide.
- T<sub>3</sub>: Crushed ginger group: birds were reared on a standard ration with crushed ginger (1000 mg / kg of ration) and tap water.
- T<sub>4</sub>: Crushed ginger and hydrogen peroxide group: birds were reared on standard ration with crushed ginger (1000 mg/kg of ration) and tap water with 0.5% hydrogen peroxide.

At the end of the study (at 56 days aged), six birds were slaughtered from each group, and the blood was collected in tubes without anticoagulants, then the serum was isolated, and kept at -20°C until biochemical tests were performed, that included determination of serum glucose, cholesterol, triglycerides, Aspartate transaminase (AST), Alanine transaminase (ALT) (Biosystem Kits, Spanish made), serum glutathione GSH level, according to the modified Burtis and Ashwood method [9], and serum MDA level which was estimated by using the modified Thiobarbituric Acid Reaction Assay method [10].

Serum corticosterone hormone level was determined by AssayMax Corticosterone Elisa Kit (manufactured by AssayMax American company), using the ELISA Micro Plate Reader Type Bio-Rad Model 550 (Japanese made).

#### Statistical analysis:

Statistical analysis was carried out using the Statistical Analysis Statics program [11]. The experiment was designed according to the complete randomized design (CRD), and the

data were analyzed by using one-way analysis of variance, and to determine the differences between the means ( $P \le 0.05$ ), we used the Duncan's Multiple Range Test [12], according to researchers Steel and Torrie [13].

# **Results and Discussion**

Table 1 indicates that the addition of hydrogen peroxide with drinking water (T<sub>2</sub>) led to a significant increase in serum glucose (338.73 mg/dl) compared to the two groups of ginger T<sub>3</sub> (278.05 mg/dl) and T<sub>4</sub> (308.88 mg/dl), as well as a significant increase in the concentration of cholesterol and triglycerides compared to all study groups at (P $\leq$ 0.05). These results are in agreement with Salah [14] who said that the addition of 0.5% hydrogen peroxide in drinking water led to a significant increase in blood glucose in broiler breeder males (Arbor Acers) compared to the control group after 4 and 8 weeks of treatment.

It was found that oxidative stress increases the secretion of corticosterone from the adrenal cortex, which stimulates the gluconeogenesis process [15]. Further, reinforcing this opinion is the presence of a significant increase in the blood corticosterone level of birds of the hydrogen peroxide group (Table 3).

The elevated serum cholesterol when administering hydrogen peroxide may be due to disturbances in the metabolic processes of lipids or in the digestive processes, which led to changes in the absorption and excretion of steroid [16]. This is consistent with what Ameen Agha [17] observed of elevated cholesterol concentration in the group of chickens that consumed  $H_2O_2$  with drinking water.

Parameters Treatments	Glucose mg/dl	Cholesterol mg/dl	Triglycerides mg/dl
T <sub>1</sub> (Control)	320.00 ±4.84 AB	$208.35 \pm 8.75 \text{ B}$	163.62 ±8.31 B
T <sub>2</sub> (H <sub>2</sub> O <sub>2</sub> )	338.73 ±4.90 A	$247.68 \pm 7.49$ A	$220.25 \pm 20.70$ A
T <sub>3</sub> (Ginger)	278.05 ±12.39 C	$166.80 \pm 8.25$ C	122.55 ±12.21 B
$T_4$ (Ginger + $H_2O_2$ )	$308.88 \pm 4.75 \ B$	$221.62 \pm \! 6.12 \ B$	$147.72 \pm 13.53 \text{ B}$

 Table 1. Effect of hydrogen peroxide and crushed ginger on blood glucose, cholesterol and triglycerides of male quails at 56 days aged (Means ±SE).

- Different letters in each column indicate a significant difference at ( $P \le 0.05$ ).

-  $H_2O_2 = 0.5\%$  and Ginger = 1000 mg/kg of ration.

The high level of cholesterol in  $H_2O_2$ -stressed birds may be due to a decrease in the activity of the thyroid gland (Hypothyroidism), as the decrease of the thyroxine in stressed birds has a negative effect on the excretion of cholesterol with the bile, thus increasing its level in the blood [18].

On the other hand, the significant increase in triglycerides was in parallel with the cholesterol increase due to the positive correlation between both parameters [19].

Table 1 also shows the role of crushed ginger (T<sub>3</sub>), which led to a significant decrease ( $P \le 0.05$ ) in the glucose (278.05 mg/dl) and cholesterol (166.80 mg/dl) concentration compared to other groups, and the triglycerides (122.55 mg/dl) compared to the hydrogen peroxide (220.25 mg/dl) group of quail.

Our results were consistent with the results of Saeid *et al.* [20] who gave the ginger aqueous extract with drinking water in two concentrations 0.4 and 0.6% significantly reduced

glucose level compared to the control group, while the study did not agree with Malekizadeh *et al.* [21] who indicated in a study on laying hens that dietary addition of ginger powder in 1 and 3% did not have a significant effect on glucose concentration compared to the control group.

The ability of ginger to reduce blood glucose levels may be due to its phenolic components which classified as hypoglycemic compounds [22], or perhaps due to presence compounds that stimulate insulin secretion, which increases the entry of glucose into the cells and thus leads to a decrease in its level in the blood. This is agreed with Elshater *et al.* [23] who said that ginger had a significant hypoglycemic effect on blood glucose of alloxan-induced diabetes mellitus compared to the control group.

As for the ability of ginger to reduce the cholesterol and triglyceride levels in the blood, it is in agreement with the findings of Han *et al.* [24], who stated that it may be due to the inhibition of lipid absorption in the intestine, because the high fiber content of ginger, as well as the oleo-resin compounds [25], which works to inhibit the absorption of bile salts in the intestine, which stimulates the liver to convert cholesterol into new bile salts, which results in a decrease its level in the blood [26].

It was also noted from Table 1 that the addition of ginger with hydrogen peroxide ( $T_4$ ) led to reducing the effect of hydrogen peroxide to elevate the level of glucose (308.88 mg/dl), cholesterol (221.62 mg/dl) and triglycerides (147.72 mg/dl), and the values remained at the control group values (320.00, 208.35 and 163.62 mg/dl respectively). This is due to the antioxidant activity of ginger, which is returned to the phenolic substances in it [5].

From Table 2 it was noticed that the addition of hydrogen peroxide led to a significant increase in the level of both AST (353.73 U/L) and ALT (12.59 U/L) enzymes in the blood compared to the control group (301.87 and 9.69 U/L respectively) at ( $P \le 0.05$ ). This result was in agreement with Taha [27] when he treated broiler breeder males with hydrogen peroxide, and it also agreed with the results of Al-kattan [28], as it recorded a significant increase in both AST and ALT in the blood of laying hens when given  $H_2O_2$  in drinking water at 0.5% concentration compared to the control group.

The high level of AST and ALT in the blood may be attributed to the oxidative stress induced by hydrogen peroxide, which leads to increased cellular oxidative stress, and produce many ROS which oxidizes the polyunsaturated fatty acids in cellular membranes and damage the channels and pumps in it [29], as a result, the membrane loses its selective permeability, due to the lipid peroxidation of the cell membrane and the leaching of these enzymes to the outside of cells [30]. The validity of this assumption is further enhanced by an increase of MDA (1.79 nmol/ml) level and a decrease in GSH (1.10  $\mu$ mol/L) level in the blood of quail that were given hydrogen peroxide (T<sub>2</sub>) (Table 3).

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Parame Treatments	ters AST U/L	ALT U/L
T <sub>1</sub> (Control)	301.87 ±9.77 B	9.69 ±0,43 BC
T <sub>2</sub> (H <sub>2</sub> O <sub>2</sub> )	353.73 ±16.48 A	12.59 ±0.63 A
T <sub>3</sub> (Ginger)	246.88 ±11.85 C	8.43 ±0.39 C
$T_4$ (Ginger + $H_2O_2$ )	255.77 ±8.84 C	$10.92 \pm 1.02$ AB

**Table 2.** Effect of hydrogen peroxide and crushed ginger on blood AST and ALT of male quails at 56 days aged (Means  $\pm$ SE).

- Different letters in each column indicate a significant difference at ( $P \le 0.05$ ).

-  $H_2O_2 = 0.5\%$  and Ginger = 1000 mg/kg of ration.

As it can be seen from Table 2 that the addition of crushed dry ginger ( $T_3$ ) led to a significant decrease in the concentration of both ALT (246.88 U/L) and AST (8.43 U/L) enzymes compared to the control group (301.87 and 9.69 U/L respectively). The results of our research are in agreement with Malekizadeh *et al.* [21] who stated that the addition of ginger powder at 1 and 3% to the diet has a significant decreasing effect on AST and ALT in the blood of white Leghorn chicken compared to the control group. It also agreed with Shanoon [6] who found a significant decrease in the level of AST and ALT in the blood of broiler breeders when adding 5 and 10% of the ginger aqueous extract to drinking water compared to the control group.

This may be due to the high level of phenolic compounds in ginger, including Gingerols, Shogaols and Zingerone, which gave ginger its effective role as an antioxidant [31, 32], as well as vitamin A and C, which leads to the inhibition of the lipid peroxidation. Accordingly, the cell membranes were protected from the oxidation processes, and these membranes retained their selective permeability property, and the cells conserved their contents and not filtered out of the cell, including these enzymes, therefore their level in blood serum decreased [33].

The addition of ginger with hydrogen peroxide ( $T_4$ ) prevented the effect of peroxide stress, and this was evident from the decrease of the enzyme levels of AST (255.77 U/L) and ALT (10.92 U/L) compared to the hydrogen peroxide group (353.73 and 12.59 U/L respectively). This is due to the antioxidant effects of ginger, which that previously mentioned, as well as its lowering effect of the corticosterone level (Table 3).

It is also noticed from Table 3 that the hydrogen peroxide-induced oxidative stress, as it is clear in the significant increase of MDA (1.79 nmol/ml) and corticosterone (15.93 ng/ml) (stress hormone) level, and the significant decrease in the level of glutathione (1.10  $\mu$ mol/L) in the quail blood of H<sub>2</sub>O<sub>2</sub>- treated group (T<sub>2</sub>) compared to all the study groups.

Para	ameters C	lutathione (GSH) μmol /L	Malondialdeh (MDA) nmol /ml	yde Corticosterone ng /ml
T <sub>1</sub> (Control)	1.:	58 ±0.06 C	$0.74\pm\!\!0.04$	B 2.54 ±0.12 C
T <sub>2</sub> (H <sub>2</sub> O <sub>2</sub> )	1.	10±0.04 D	$1.79 \pm 0.12$	A 15.93 ±0.77 A
T <sub>3</sub> (Ginger)	2.9	92 ±0.13 A	$0.41 \pm 0.05$	C 1.52 ±0.18 C
$T_4$ (Ginger + H <sub>2</sub> O <sub>2</sub> )	2.0	$00 \pm 0.09$ B	$0.74 \pm 0.05$	B 4.56 ±0.57 B

 Table 3. Effect of hydrogen peroxide and crushed ginger on blood Glutathione, Malondialdehyde and Corticosterone of male quails at 56 days aged (Means ±SE).

- Different letters in each column indicate a significant difference at ( $P \le 0.05$ ).

-  $H_2O_2 = 0.5\%$  and Ginger = 1000 mg/kg of ration.

Pravda [34] and Ayed *et al.* [35] stated that hydrogen peroxide depletes the glutathione in the blood and tissues and causes a decrease in its level in the body and that its decrease is associated with an increase in the level of malondialdehyde, which is considered one of the main indicators of the occurrence of the lipid peroxidation. This finding was in line with AL-Ma'atheedi and Hassan [36] in their study on adult roosters exposed to oxidative stress, and Taha [27] who reported that the addition of hydrogen peroxide to drinking water at 0.5% of broiler breeder males led to a significant increase in malondialdehyde level, and a significant decrease in the glutathione level in testicular tissue compared to the control group, also was referring to that by Ameen Agha [17] in broilers. The elevation of corticosterone level in the  $H_2O_2$  group (T<sub>2</sub>) (Table 3) in response to stress induced by  $H_2O_2$ . This is consistent with Abdul-Majeed [15] who said that the oxidative stress induced by  $H_2O_2$  leads to an increase in the secretion of the corticosterone (Stress hormone) from the adrenal cortex in response to stress and mentioned that the hydrogen peroxide with drinking water led to a significant increase in corticosterone level of quail compared to the control group at (P≤0.05).

It is also evident from Table 3 that the dietary addition of ginger (T<sub>3</sub>) improved and significantly increased serum glutathione level (2.92  $\mu$ mol/L) compared to the control group (1.58  $\mu$ mol/L), and with the other study groups at (P $\leq$ 0.05). And when ginger was given with hydrogen peroxide (T<sub>4</sub>), it was able to prevent the effect of stress caused by hydrogen peroxide and improve the serum antioxidant status, as it raised the level of glutathione significantly (2.00  $\mu$ mol/L) compared to both groups of hydrogen peroxide and control (1.10 and 1.58  $\mu$ mol/L respectively).

And in regard to MDA, it was significantly decreased when ginger was added to the feed (T<sub>3</sub>) (0.41 nmol/ml) compared to the control group (0.74 nmol/ml) and the other study groups at (P $\leq$ 0.05), and it was at the level of its values in the control group when ginger was added with hydrogen peroxide (T<sub>4</sub>) (0.74 nmol/ml). Ginger was able to prevent the effect of stress-induced by hydrogen peroxide and returned the MDA values to the level of its values in the control group, and it was even significant with the hydrogen peroxide group at (P $\leq$ 0.05) (Table 3).

Our results are agreed with Ogbuewu *et al.* [37] who indicated that ginger acts to protect the cells because it contains gingerols and shogaols that play an essential role in protecting cells from oxidative damage resulting from the free radicals. Also, our results agree with the results of researcher Shanoon [6] who found a decrease in the level of malondialdehyde and an increase in the level of glutathione in the groups of broiler breeder chickens that consumed a diet containing 2.5 g ginger/ kg of feed compared to the control group.

The ability of the ginger plant to raise the level of glutathione and reduce the level of malondialdehyde is consistent with all of the Tohma *et al.* [38], Zidan *et al.* [39] and Zhang *et al.* [40] when they indicated that the ginger is one of the plants that possess an antioxidant capacity due to its high phenolic contents. Also, the ginger's ability in decreasing corticosterone may be related to its vitamin C content which inhibits 11-B Hydroxylase and 21-Hydroxylase, so it reduces corticosterone production by the adrenal cortex [19].

To confirm the validity of this assumption, when was given the ginger alone (T<sub>3</sub>), it reduced or decreased the level of the stress hormone (corticosterone) (1.52 ng/ml) than it was in the control group (2.54 ng/ml), and when it was added with the hydrogen peroxide (T<sub>4</sub>) (4.56 ng/ml), there was a significant decrease in its level compared to the hydrogen peroxide group (T<sub>2</sub>) (15.93 ng/ml), because it reduced and suppressed the effect of stressful hydrogen peroxide and raised the level of the corticosterone hormone (Table 3).

#### Conclusion

We conclude, our study revealed that the hydrogen peroxide was heavily impacting on all of the studied parameters of the blood, and the addition of a substance that has an antioxidant capacity as ginger (*Zingiber officinale*) prevented the negative effects of oxidative stress induced by hydrogen peroxide, also that the ginger has an ability or role to improve or enhance the antioxidant status in the body, and returned most of the parameter values and antioxidant status to their values in the control group.

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# **Conflict of Interest**

The authors declare that there are no conflicts of interest regarding the publication of this manuscript.

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