Histological, Scanning And Transmission Electron Microscopic Studies On The Possible Protective Role Of Ginger Extract Against Acrylamide Induced Intestinal Damage In Mice

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Abstract

**Objective:** This study was carried out to evaluate the protective effect of ginger *Zingeber officinale* extract (ZOE) against the acrylamide (AC) which is an industrial chemical used in water treatment and it is synthesized during cooking of starch food at high temperature.

**Method:** Thirty adult male albino mice, each weighs 20-25 g were divided into three groups (10 mice/group): (I) control group. (II) acrylamide treated group. (III) acrylamide & ginger group. Acrylamide was given to experimental animals in the drinking water at a non-lethal dose of 200 p.p.m for 10 weeks (3 days/week). Ginger extract was orally administrated at 50 mg/L (~5 ml/day) for 10 weeks (3 days/week). The ileum samples were collected for light microscope study and for scanning and transmission electron microscope examination.

**Results:** This study revealed that acrylamide induces pathological changes of the ileum of the treated mice specially the absorptive epithelial cells. The scanning electron microscopic study revealed damage of the ileal villi, some red blood corpuscles appeared at the site of damage. The transmission electron microscopic examination clearly demonstrated degeneration of most cell organelles as mitochondria, deterioration and degranulation of the rough endoplasmic reticulum, dilatation of Golgi apparatus.

**Conclusion:** The administration of ginger extract decreased the histological alterations and ensuring the anti-inflammatory, and antitoxic effects of ZOE at its chosen dosage level.

**Key words:** Acrylamide – ginger- ileum.

Introduction

Acrylamide is an industrial chemical used in the synthesis of polyacrylamide and has multiple applications as additive for sewage and water treatment (Smith and Oehme, 1991). Individuals can be exposed to acrylamide either in their workplace or in the environment (Marsh et al., 1999). Recent findings of the presence of acrylamide in starch foods cooked at high temperature have refocused worldwide attention on its carcinogenicity (Tareke et al., 2002). In addition, acrylamide is used extensively in molecular laboratories for gel chromatography (LoPachin, 2004).

According to WHO (2002), the average intake of 70µg of acrylamide per day for an adult would be associated with a lifetime cancer risk . Studies in animal models and humans are necessary to better understand the toxicity of acrylamide .It was shown that acrylamide might be formed through the Maillard reaction from amino acids (e.g asparagines) and reducing sugars (e.g glucose) (Mottram et al., 2002).

Glycidamide, a metabolite of acrylamide, binds to DNA and can cause genetic damage. Prolonged exposure to acrylamide has induced tumours in rats, but cancer in man has not been convincingly shown. The International Agency for Research on Cancer (IARC) has classified acrylamide as "probably carcinogenic to humans" (IARC, 1994). Barber et al. (2001) reported that the rate of acrylamide conversion to its epoxide metabolite glycidamide is higher during subchronic dosing conditions.

On the other hand the ginger has been widely studied for its pharmacological activities and has been reported to exhibit anti-inflammatory, antipyretic, antimicro-bial, hypoglycemic, antimigraine, antiox-idant, hepatoprotective, diuretic, hypocho-lesteremia (Langner et al.,1998; Mascolo et al.,1989) and antihypertensive activities (Ghayur and Gilani., 2005). Phytochemical studies showed the presence of pungent
principles, such as gingerol, shogoal, zingerone and paradol (Connell and McLachan., 1972).

The rhizome of the plant Zingiber officinale roscoe, commonly known as ginger, has been commonly used as a food additive and spice as well as phytomedicine since ancient times. The typical use of ginger in the kitchens as a condiment began in the 13th century; which enhanced the importance of this rhizome in european markets (Langner et al.,1998). More commonly, ginger has been traditionally used in disorders of the gastrointestinal tract, as a stomach laxative, sialogogue, gastric emptying enhancer, appetizer, antiemetic and antidyspeptic and at the same time as an anti diarrheal and anticolic agent (Ghayur and Gilani., 2005; Nadkarni, 1976).

Several studies conducted in animals (Yamahara et al., 1990; Qian and Liu., 1992) and humans (Mowerey and Clayson., 1982; Sharma and Gupta., 1998) showed the prokinetic action of ginger; however, the precise mechanism of its action is not yet clear. On the contrary, some studies also reported the inability of ginger to impart any stimulant effect on the bowel (Stewart et al., 1991; Phillips et al.,1993); while others showed that ginger exhibits a spasmolytic action but the precise mode of action remains to be elucidated. However, no study pointed out the presence of a combination of stimulatory and inhibitory activities in ginger, because of scarcity of information on acrylamide and its metabolites on the intestinal tissue. This has opened up new avenues for understanding the pathogenesis of intestinal degeneration induced by acrylamide and the role of ginger in minimizing the toxicity induced by it.

**Material And Methods**

**Materials**

1-Acrylamide (AC) is patented by P.S Park scientific limited, Northhampton, United Kingdom. (AC) is a chemical intermediate (monomer) used in synthesis of polyacrylamides.

Acrylamide monomer is a white crystalline form, soluble in water, ethanol, methanol and acetone. Its chemical formula is CH$_2$CHCONH$_2$. Synonyms of acrylamide are: 2-propenamide, ethylene carboxamide, acrylamide and vinylamide.

2-Ginger or Zingerber officinale roscoe (Family Zingeberaceae). Ginger ethanolic extract was prepared from conc-entrated pure ginger powder. The stock solution of the ginger extract at concen-tration 1g/l was prepared in 22% alcohol (200 mg of ginger extract was dissolved in 44 ml of ethanol and the volume was then adjusted to 200 ml with water). Drinking solution was prepared freshly every 3 days, by dilution of 25 ml of the ginger extract stock solution into 500 ml of water, resulting in final concentration of 50 mg/l of ginger extract in 1.1 % alcohol according to Bianca et al. (2000).

**Experimental design**

This study was carried out on 30 male adult CD-1 mice, each weighs 20-25g; they were divided into three groups, 10 mice each. The first group served as control and received 1.1% alcohol (11 ml of alcohol in 1 l water) for 10 weeks and fed ad libitum. The second group was given acrylamide at a dose 200 p.p.m in drinking water for 10 weeks (3 times/week) according to Ko et al. (1999). The third group received ginger (ZOE) at 50 mg/L (~5 ml) 1 hour prior to the administration of acrylamide (200 p.p.m) in drinking water for 10 weeks (3 days/week) according to Bianca et al. (2000).

**Methods**

Acrylamide was administered in drinking water, and after 10 weeks (3 times /week) the animals were decapitated; small parts of the ileum were immediately excised and fixed in alcoholic Bouin for light microscopic study. Specimens were dehydrated, cleared and embedded in paraffin wax. Sections of 5 μm in thickness were stained with hematoxylin and eosin (Chayen et al., 1973). The cytoplasm appeared reddish-pink and the nuclei acquired a blue colour.

Other pieces of the ileum were fixed in 2% glutaraldehyde in 0.1 phosphate buffer, postfixed in 1% osmium tetra oxide for 2 hours at 4°C, dehydrated and embedded in epon. The semithin sections were stained with toluidine blue and the ultrathin
sections stained with uranyl acetate and lead citrate and examined on transmission electron microscope (JEOL-Ex 1010 transmission electron microscope at Al-Azhar University).

For scanning electron microscopic examination (SEM), small pieces of ileum were washed several times in distilled water. They were fixed in phosphate buffered 2.5% glutaraldehyde for 3-4 hr and postfixed in phosphate buffered 1% osmium tetra oxide for 1 hr. The specimens were dehydrated in a graded series of ethanol and dried at a critical point using liquid CO₂. The dried specimens were attached to the stubs and then coated with gold by coating apparatus JFC-1100E Ion sputter. These specimens were examined on JEOL 1200EX II electron microscope at 28 KV at the Faculty of Science, Ain Shams University, Cairo, Egypt.

Results

General observations

Mice treated with acrylamide exhibited restlessness and a marked increase of body weight.

Light microscopic examination

a- Control animals

The ileal mucosa of the control mice is built up of numerous folds forming the villi, through which the connective tissue of the lamina propria containing the simple tubular glands i.e; the crypts of Lieberkühn is found (Fig.1).

The lining epithelium of the villi is composed of many cell types, such as the absorptive columnar epithelial cells (enterocytes), goblet cells and Paneth cell. The predominant cell type is the enterocytes; they have striated borders, finely granular cytoplasm and oval basally located nuclei (Figs.2&3). The goblet cells are scattered between the enterocytes, they have heavily chromatinated basal nuclei (Fig.2).

The epithelial layer lining the crypts of Lieberkühn is continuous with that of the villi (Fig.4). The lamina propria of the mucosa is formed of fine connective tissue containing lymphocytes, fibroblasts and blood capillaries (Fig.5).

b- Acrylamide treated animals

Mice treated with acrylamide for 10 weeks revealed marked histopathological alterations. The lining epithelium of the mucosa showed marked discontinuity (Figs. 6&11). The absorptive columnar cells displayed signs of necrosis characterized by distinct vacuolations and pyknotic nuclei (Figs.6&7).

The mucus-secreting goblet cells were few (Figs. 6, 7&9). These changes resulted in the formation of large clear spaces in this material. In addition, lymphocytic infiltration was prominent and numerous enlarged lymph nodules appeared in the lamina propria of the villi (Fig.8).

The epithelial cells of the crypts of Lieberkühn were faintly stained with ill-defined cell boundaries, vacuolated cytoplasm and pyknotic nuclei (Figs. 9&10).

c- Animals received ginger together with acrylamide

The histological structure of the ileal mucosa appeared more or less normal (Figs. 12&13). The absorptive columnar epithelial cells lining the crypts of Lieberkühn and the villi were clearly demineralized and were more or less similar to that of the control group (Figs.14&15). The lamina propria showed less damage and alterations (Fig.16).

Scanning electron microscopic examination

The filiform and foliate ileal villi of the control animals appeared normal, with non-damaged cover. Few mucous secretions were noticed, and normal submucosal layer appeared as a base for the villi (Figs. 17&18).

Administration of acrylamide (AC) for 10 weeks resulted in apparent changes appeared in some surface areas of the villi (Figs.19&20). Also, mild bleeding appeared as red blood corpuscles in the intervillar spaces and facing the lumen, this bleeding was accompanied with severe damage of the villi (Fig.21).

On the other hand, animal group which had received the ginger (ZOE) before (AC), showed that the villi appeared more or less normal. Also, some of mucous secretions were observed but no bleeding or blood corpuscles were shown (Figs. 22, 23 & 24).

Transmission electron microscope examination

a. Control animals

Electron microscopic examination of the ileal mucosa revealed that the ileal
mucosal epithelium is built up of one layer containing many types of cells resting on a basement membrane.

The absorptive columnar cells have ovoid basally located nuclei (Fig.25). Each nucleus had a prominent nucleolus and heterochromatin clumps mostly adjacent to the inner surface of the nuclear envelope. The cells are provided with microvilli at their luminal surfaces. The microvilli appeared as closely packed, long, parallel projections on the apical surfaces of the epithelial cells (Fig. 28).

The cytoplasm of the absorptive cells contained mitochondria of various shapes with well developed cristae (Figs.26&27). Rough and smooth endoplasmic reticulum (ER) were seen in the cytoplasm (Fig.27). The endoplasmic reticulum ER is formed of a continuous network of canaliculi and saccules throughout the cell especially near the nucleus (Fig.26). The reticulum was predominantly rough (carrying ribosomes), but smooth ER was also found especially towards the apex (Fig.29).

At the cell periphery, especially near the apex, junctional complexes bound the adjacent cells together (Fig.27). The lateral cell interfaces showed interdigitations of adjacent plasmalemmeae. The Paneth cell (Fig.30) appeared with oval, basal nucleus and prominent nucleolus. Abundant rough endoplasmic reticulum and large secretory granules were clearly shown.

Goblet cells were few among the absorptive cells in the ileum, without brush border. They contained electron-lucent mucus secreting granules and basal nuclei (Figs.31&32).

b. Acrylamide treated animals

The ileal epithelial cells of mice received acrylamide for 10 weeks, displayed several changes in their ultrastructure. In some cells, especially those in the villus base, the microvilli forming the brush border were partially degenerated and showed few plebs (Fig.33). In other cells, especially those located at and near the villus tip, the microvilli appeared only slightly affected (Fig.35). Severe destructive changes in the absorptive columnar epithelial cells were observed. Most nuclei of the cells of the treated animals showed marked alterations (Fig.33); some of them displayed few chromatin and others seemed pyknotic and had highly irregular outlines, internal cloudy appearance with the disappearance of a prominent dense nucleolus (Fig.35). Moreover, in some highly injured cells, the nuclei were degenerated (Figs. 35, 36&37).

The Golgi apparatus of acrylamide-treated animals cells seemed hypertrophied, so its cisternae were dilated (Fig. 34). The arrangement of the rough ER cisternae was somewhat disturbed; in some regions of the cytoplasm these cisternae seemed fragmented (Fig.35). Many cells were severely injured (Fig.37). Figure 36 represents two cells in their way to degenerate. They appeared undergoing cloudy degeneration, and the ER lumen was dilated and degenerated.

Acrylamide administration for 10 weeks revealed that the majority of the mitochondria were slightly swollen, and the cristae of some of them had lost their normal appearance and became difficult to be distinguished (Figs. 34&35).

The goblet cells became few, tall, more globular and many vacuoles were found within the cytoplasm. Their nuclei showed pyknosis and became more basally located. Smooth endoplasmic reticulum succuli became very dilated (Fig.38).

Figures 39 and 40 showed damage of the lamina propria in the core of the ileal villus. This damage was accompanied with the presence of many lymphocytes.

c. Animals received ginger together with acrylamide

The electron microscopic examination ileum of mice that received ginger (ZOE) extract 1 hour before the acrylamide administration showed that the general architecture of the ileal tissue was more or less similar to that of the control group. The nuclei of the absorptive cells appeared with normal basal position surrounded by abundant cytoplasm, the microvilli of the brush border were projecting in a regular and normal manner (Fig.41). Some cisternae of Golgi apparatus, and many lysosomes were prominent (Fig.42). In addition, a plenty of mitochondria and the rough endoplasmic reticulum were observed (Fig.44). Different types of leukocytes with normal nuclei and cytoplasm were clearly noticed (Fig.43).
Figures 1-5: Photomicrographs of transverse sections of the ileum of control mice.

Fig. 1: Photomicrograph of transverse section of ileum of control mice showing the mucosal villi, and the crypts of Lieberkühn (arrows). (H&E)

Figures 2, 3 & 5: A magnified regions, illustrating the structure of ileal villi showing the simple columnar epithelial cells with the brush border (arrow head) interspersed with few goblet cells (G) and interepithelial lymphocytes (arrows). (Figs. 2 & 5 H&E) (Fig. 3. TB)

Fig. 4: Showing the structure of the crypts of Lieberkühn lined with columnar epithelial cells and few goblet cells. (TB)
Figures 6-11: Photomicrographs of the ileum of mice treated with acrylamide for 10 weeks.

Figures 6, 7 & 11: Showing vacuolation and degeneration of lamina propria, apparent hyperplasia of the surface columnar cells which appeared to be separated from each other at many sites (thin arrow). (Figs. 6 & 7 H&E, Fig. 11 TB)

Fig. 8: A magnified part of the mucosal villus of ileum of mice revealed the loss of villus architecture and increase of interepithelial lymphocytes with few goblet cells. (H&E)

Fig. 9: A magnified part of the villus apical region, the cytoplasm showing vacuolation. (H&E)

Fig. 10: Illustrating the crypts of Lieberkühn, faintly stained epithelial cells, Pyknotic nuclei and vacuolation of the cytoplasm (arrow heads). (TB)

Fig. 11: Showing damage and vacuolation of lamina propria (right upper arrow), also notice discontinuity of surface columnar cells (left lower arrow). (TB)
Figures 12-16: Photomicrographs of the ileum of mice treated with acrylamide after receiving the ginger extract ZOE.

**Figures 12&13**: Showing normal histological structure of the columnar epithelial cells of the ileal villi and lining of the crypts. (H&X)

**Figures 14&15**: Showing a magnified part of the inner regions of the crypts and columnar cells which facing the lumen, the villi regained their normal structure, lamina propria appeared with more or less normal structure in the core of the villi. (H&E)

**Fig.16**: Showing the mucosal villi with more or less normal lamina propria. Notice that the villi become more thinner than of control villi, and very few goblet cells (G) are observed. (TB)
Figures 17&18: Scanning electron micrographs of the dorsal surfaces of different types of mucosal villi (filiform & foliate) of ileum of control mice.

Figures 19-21: Scanning electron micrographs of the ileal villi of mice treated with the acrylamide, showing loss of the normal structure of the villi accompanied with damage and bleeding which is shown in Fig.21 at the sites of red blood corpuscles.
Figures 22-24: Scanning electron micrographs of the ileal villi of treated mice with acrylamide and previously received ginger, showing the normal structure of the villi with presence of mucous secretions (arrows) on the surface.
Figures 25-32: Transmission electron micrographs of the ileum of control mice.

Fig.25: Showing the columnar (absorptive) cells (c) with microvilli (arrow heads), different types of cells such as endothelial cells (arrow), and lymphocytes (L) can be shown. (X 3000)

Fig.26: Showing a magnified part of absorptive cell illustrating the nucleus (N), different forms of mitochondria (M) with well developed cristae. Note the smooth endoplasmic reticulum (SER) and the free ribosomes (R). (X 20000)

Fig.27: Showing the cellular junctions between the adjacent cells (arrows). Notice the rough endoplasmic reticulum (RER). (X10000)

Fig.28: Showing the microvilli of control ileum. (X25000)
Fig.29: Showing intercellular junctions. Notice infoldings of the plasma membrane and the microvilli (arrows). The rough endoplasmic reticulum (arrow heads) surrounding the mitochondria. SER at the apical regions is observed. X10000

Fig.30: Showing the normal paneth cell (PC) with basal nucleus, prominent nucleolus, abundant RER and large secretory granules. X13000

Fig.31: Showing a part of goblet cell (G) with its mucous secretion. X 10000

Fig.32: Showing the goblet cell (G) with oval nucleus (n) and many cisternae of RER (arrows) were observed in the dark cytoplasm. X8000
Figures 33-40: Electron micrographs of ileum of mice treated with acrylamide for 10 weeks at dose 200 p.p.m.

Fig.33: Showing highly damaged columnar absorptive cells, degeneration of the microvilli and formation of blebs (arrow heads), pyknotic irregular nuclei (arrow). Note the hypertrophy of Golgi elements (GO). X6000

Fig.34: Showing a magnified part of an absorptive columnar epithelial cell with dilated cisternae of Golgi apparatus (arrows). Notice the pyknotic nucleus (N). X20000

Fig.35: Showing highly damaged and degenerated epithelial cells. Notice the pyknotic nucleus (N), fragmentation of RER, complete deterioration of the ground cytoplasm and mitochondria (M). X15000

Fig.36: Showing complete degeneration and dissolution of the cytoplasm resulting in formation of large vacuoles. Notice the karyolytic cells (arrows). X8000
Fig.37: Showing a necrotic cell with complete degeneration of its nucleus. Notice the dilatation of Golgi apparatus cisternae (GO).

Fig.38: Showing atrophy and degeneration of the goblet cell. Notice the SER with dilatation of their succuli and disappearance of most cell organelles.

Fig.39: Showing invasion of many lymphocytes.

Fig.40: Showing the core of an ileal villus with many lymphocytes, notice edematous lamina propria.
Figures 41-45: Electron micrographs of the ileum of a mouse treated with acrylamide and received the ginger (ZOE) 1hr prior the acrylamide administration.

**Fig.41:** Showing the absorptive columnar epithelial cells appeared more or less normal. Lysosomes (LY) were also noticed and mild dilatation of Golgi apparatus cisternae. (GO). X15000

**Fig.42:** Showing a magnified part of the absorptive columnar cell with dilated cisternae of Golgi apparatus. The cytoplasm contains more or less normal cell organelles. X 15000

**Fig.43:** Showing different types of leukocytes and the lamina propria displaying less damage and edema. X5000

**Fig.44:** Showing normal distribution of the mitochondria (M) normal appearance of RER which carries many ribosome (arrow heads). X20000
Discussion

It is evident from the present study that the light, scanning and transmission electron microscopic studies of ileal epithelial cells of the mice adversely responded to the administration of the monomer crystalline substance acrylamide. Acrylamide is a water soluble organic substance; it is absorbed by all routes and easily distributed throughout the body organs (Barber et al., 2001). In the present study, the mice were chosen for these experiments due to their sensitivity to the toxic substances.

Paulsson et al. (2002) reported neurotoxic and carcinogenic effect of acrylamide in animals, and revealed the lower sensitivity of the rat than that of the mouse to the carcinogenic action of this substance. In 1995 Friedman and his associates concluded that the administration of acrylamide in drinking water increased the incidence of mammary tumours.

The gastrointestinal tract is lined with a single layer of epithelium, which forms a highly selection barrier designed to allow the efficient transport of nutrients and water, while preventing the entry of potentially toxic luminal pathogenic organisms and their products or any toxic substances (Adam et al., 2006). According to those authors, (as an interpretation of our finding), acrylamide may have the ability to destroy this intestinal barrier.

At the beginning of the present study after two to three weeks of administration of acrylamide (200 p.p.m) in the drinking water, a marked increase of weight was observed (Marsh et al., 2007). Histopathologically the lymphocytes infiltration were noted, this finding come in agree with the physiological explanation which based on the pharmacokinetic model for acrylamide and it was found that it is distributed within five compartments (arterial blood, venous blood, liver, lung and all other tissues lumped together) (Kirman et al., 2003). The present results showed that the superficial luminal cells of the ileal villi of the treated animals were affected; this agree with studies on aspirin which have shown to induce similar damage (Ivey et al., 1980). Vacuolar degeneration, nuclear pyknosis, cloudy swelling and necrosis were of the most marked signs of tissue impairment in the present study. These destructive changes were similarly reported in the colon, nervous and reproductive systems and produce tumours in certain hormonally responsive tissues following treatment with acrylamide (Barber et al., 2001).

In the present study, the surface mucosa of the ileum was studied by using scanning electron microscope. After treatment with acrylamide, the cell surfaces were damaged in some areas. Similar results were observed by Specian and Neutra (1980) who investigated the epithelial topography of the surface crypt cell in rabbit and monkey colon. In addition, Bonvicini et al. (1985) reported a clinical investigation of different gastrointestinal diseases by using (SEM); they found that the mucosal surface of the involved mucosa showed certain lesions as bleeding and enlargement of the crypts. These results agree with the present findings where the bleeding appeared and was indicated by the presence of the red blood corpuscles.

Kamel et al. (1985) investigated the structural changes in mouse intestinal villi following lower body heating by using SEM. Some villi were severely damaged at their tips; these are similar to the damage described in the present study after acrylamide treatment.

The ultrastructural study showed that the microvilli of the absorptive columnar cells exhibited signs of degeneration. A similar observation has been described in the small intestine of Tilapia nilotica (Sakr, 1993), where signs of damage and necrosis of the columnar epithelial cells appeared, and then the cells revealed morphological alterations. These results suggested that the columnar cells with their brush borders are the main target cells and act as endocrine disruptors after treatment with chemicals that manufacture plastic products (Besaratinia and Pfeifer, 2004) who concluded that the mutagenicity of acrylamide in human and mouse cells is based on the capacity of its epoxide.
metabolite glycidamide to form DNA adducts. The mitochondrial destruction in the absorptive columnar cells of the ileum of the mice was one of the important results which has been illustrated under various pathological conditions (Adams et al., 1977; EL-Beih et al., 1993).

The present results also showed that the rough endoplasmic reticulum was broken-down, degranulated and fragmented into small structures after treatment with Acrylamide. Such damage is considered to be one of the essential factors responsible for the lowered activity of the cell (Elewa et al., 1999).

The Golgi apparatus of the cells of treated animals displayed damage. These results confirm the findings of other research workers in different pathological instances (Moussa et al., 1987; Winton and Flaks, 1988; EL-Beih et al., 1994). The observed increase of lysosomes in the cells of animals received ginger together with acrylamide was confirmed by the reports presented by Helman et al. (1985) which revealed that the lysosomes undergo proliferation, abundance and finally rupture at later stages of adverse conditions, with the consequent release of their digestive hydrolases into the cytosol could speculatively account for a considerable proportion of damage produced in the cells as observed in the present work. Moreover, the present study showed that the mucous goblet cells were very few, became elongated but with degenerated nucleus and cytoplasmic organelles; also the Paneth cells were hardly detected. Furthermore, many lymphocytes were seen in the lamina propria. These results explained the potent immunologic function of the ileum, and agree with the study that was carried out by Mohamed et al. (2006).

In the present study, the administration of ginger (ZOE) 1 hr prior to acrylamide in drinking water decreased the histopathological alterations which induced by acrylamide. Where, the ethanolic ginger extract was selected to apply in this study according to Mascolo et al. (1989) who found that the ethanolic ginger extract (100-300 mg/kg) contains compounds which inhibit prostaglandin release by leukocytes, which may be responsible for ginger's antipyretic activity and anti-inflammatory effects.

Abdel-Ghaffar (2006) reported that the animals treated with ZOE alone did not show any alteration in serum glucose. So, this could ensure the protective role of ginger against the oxidative hyperglycemia. As far as we can throw more light on the beneficial effect of ginger which is often touted for health benefits and how it was able to reduce the ultrastructural changes caused by acrylamide in the ileal mucosa of mice. Cotran et al. (1994) reported that the mitochondrial dysfunction resulting from acrylamide treatment that would create lack of ATP needed for normal function due to its toxic effect {histotoxic hypoxia} and the mitochondria were significantly protected after incorporating ginger.

In consistency with the result of the present study Ghayur and Gilani (2006) found that the extract of ginger exhibits species spasmogenicity in gut tissues, along with a dormant no effect, mediated via the blockade of voltage-dependent Ca$^{2+}$ channels. Thereby, it maintains the integrity of the plasma membrane and protects it against apoptosis.

Zhongguo (1992) reported that ginger juice exhibits anticholinergic and antihistaminic action. Ginger juice produces antimotion sickness action by central and peripheral anticholinergic and antihistaminic effects. Since it was found that the acrylamide substance is classified as carcinogenic and mutagenic substance that can induce DNA damage in the cell (Barber et al., 2001).

Nakamura and Yamamoto. (1982& 1983) reported that ginger juice contains antimutagenic components that suppress \( \text{6}-\text{gigerol} \) which is a potent mutagen. So, it could induce DNA repair in damaged cells and induce their apoptosis. In addition, they revealed that the ginger has been repeatedly proven effective in the gastrointestinal distress and is indicated as safe to use in moderate amounts. Overall, studies have proved it to have some effect on motion sickness and post-operative nausea. However, the studies failed to indicate any CNS involvement.

In conclusion, the histopathological changes of ileum may contribute to toxic effect of acrylamide, DNA changes which lead to pyknosis of the nuclei and necrosis of the cells. Hence, the application of electron microscopy in the present study has the merit of bringing into vision numerous details on the ginger protective role which could be considered a good remedy in gastrointestinal problems that counteracts the histopathological impairments caused by acrylamide and provides improvement of the general health.

References


دراسات هستولوجية و تركيبية دقيقة للتأثير المحتمل لمستخلص الزنجبيل المضاد للإلكتروليد المحدث لتلف الأمعاء الدقيقة في الفئران البيضاء

هالة جلال محمذ الطنطبوى
قسم علم الحيوان – كلية العلوم- جامعة عين شمس

الأكريلاميد مادة كيميائية صناعية تستخدم في تغليف مادة البولي أكريلاميد وكذلك
تستخدم في العديد من التطبيقات الصناعية. كما أن البولي أكريلاميد تستخدم لتنقيه مياه
الشرب. وقد تؤدي الجرعات المتراكمة من مادة الأكريلاميد غير الملممة إلى حدوث أضرار
جسيمة للأعضاء المختلفة بجسم الإنسان. من أجل ذلك تم إجراء هذه الدراسة بغرض عمل
دراسات هستولوجية بالمجهر الضوئي وكل من المجهر الإلكتروني الماسح والمجهر
الإلكتروني النافذ على الأمعاء الدقيقة للفاير الأبيض ثم دراسة الدور الوقائي لمستخلص
الزنجبيل.

تم تقسيم الفئران إلى ثلاث مجموعات: الأولى ضابطة معالجة بـ 1.1 محلول كحولي،
والثانية عولجت بالأكريلاميد في مياه الشرب (200 جزء في المليون) لمدة 10 أسابيع
(ثلاثة أيام في الأسبوع) والمجموعة الثالثة وهي المجموعة المعالجة بالأكريلاميد و
الزنجبيل أُعطيت مستخلص كحولي للزنجبيل (50 ملليجرام/لتر) حيث تم إعطاء الحيوانات هذا
المستخلص قبل الأكريلاميد بساعة واحدة.

وقد وجد أن الأكريلاميد تسبب في حدوث تلفا في الأمعاء الدقيقة على شكل تلف في
الخلايا الطائتة للعمليات المعوية إلى جانب حدوث تلف في طبقة الأنسجة الضامة داخل
العمليات المعوية. أيضاً لوحظ تدمير وترسبات لتركيبات الدقيقة للخلايا الطائتة. اضطراب
التخليص العملي للنسيج وظهرت فيه الخلايا الالتهابية الارتياحية وتكونت تجاويف سبتيزيمية
في الخلايا وظهرت علامات التحلل الشامل هذا على مستوى الفحص بالمجهر الضوئي. أما
المجهر الإلكتروني الماسح فقد أظهر أن الخلايا متضررة من تدمير بعض النزف والمهرو
النافذ أظهر تغيرات ملحوظة في عضيات الخلايا، الأغشية البلازمية. تحدثت الميكروكوندريا
بغير التركيب الدقيق لجهاز حولي وظهر أيضاً أضمحل ومحدث نقص في الشبكة
الإنبوتازيمية كما حدث تلف في أونية الخلايا مما غير شكلها وظهر نقص في المادة
الكروماتينية بداخلاها وظهور علامات التدمير الكامل.

وأوضحنا المعاملة بمستخلص الزنجبيل إلى جانب الأكريلاميد تحسنًا ملحوظاً في نسيج
الأمعاء الدقيقة. وقد أوضحنا الدراسة دور الزنجبيل كمشروع طبيعي وقائي فعال ضد
التأثير الضار لهذه المادة و التي يمكن أن تنتج من طريقة طبخ بعض الأطعمة التشوية.