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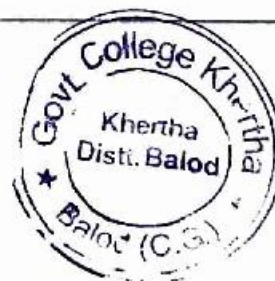
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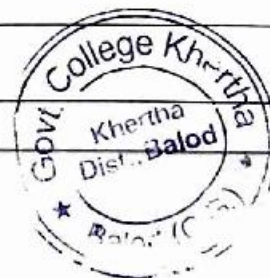
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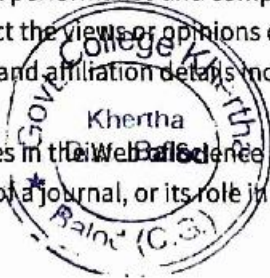
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Impact of Heavy Metals Consumption on Human Health: A Literature Review

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Author's contribution

The sole author designed, analyzed, interpreted and prepared the manuscript.

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Review Article

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ABSTRACT

While some heavy metals are essential trace components, many are bio toxic in human biochemistry. As a result, a full grasp of underlying systems is required. For supporting life and minimizing environmental damage, we must grasp their sources, liquidation techniques, chemical modifications, and deposition patterns. These metals are discharged into the environment by both natural and human activity, including mining, industrial operations, and vehicular emissions. Soils and groundwater are contaminated when they leak into subsurface fluids and eventually into the aquifer. The world's commerce and coordinating systems frequently generate environmental toxicity and poisoning. Swallowed, they generate bio toxic compounds, lose structural integrity, and block bio reactions. This study's goal is to learn more about heavy metals and their bio toxic effects on humans.

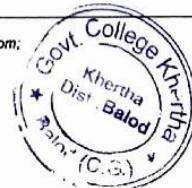
Keywords: Heavy metal; environment; human health; illness.

1. INTRODUCTION

Environmentally important metallic elements in the environment, they are tough to eradicate.

Metal is an important class of harmful chemicals in various occupational and environmental circumstances. The ubiquity of exposure today affects these harmful compounds to human

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health. The increased use of the wide range of metals in business and our daily lives has resulted in serious worries due to dangerous metal contamination in our environment.

Metals are a natural component of the ecosystem. They are highly conductive substances that voluntarily release their electrons into cations. In biodiversity, metals such as air, crusts, water bodies, including plants and animals, can also develop worldwide. 23 of the present 35 natural metals contain high specifications and atomic densities above 5 g/cm³, commonly referred to as heavy metals [1-2]. Heavy metals include antimony, tellurium, steel, platinum, silver, manganese, cobalt, cadmium, steel, platinum, and zinc. The heavy-metal high-density class is known not only for its negative impact on ecosystems and organisms [3]. Heavy metals which may produce inappropriate diseases and syndromes, if inappropriate, but which are of acute or chronic toxicity at high doses.

The Heavy Metals can be distributed by various natural processes including volcanic eruptions, spring water, erotic and bacterial activity as well as by the use of fossil fuels, industrial processes, agriculture and feed [4]. These heavy metals have diverse effects on living species and humans. Metallic macromolecules bind and hinder cell function. Thus, heavy metal poisoning in humans has numerous effects. The CNS can cause harmful blood molecules in the lungs, liver, and kidneys [5]. Physical, musculoskeletal, and neurological functions are slowing down, mimicking Parkinson's and Alzheimer's [6]. Acids, mutations, hormone disorders, and cancer may ultimately be damaged by long-term contact or by heavy metal compounds [5].

2. MECHANISM OF HEAVY METAL TOXICITY

Some heavy metals release free radicals that can harm other cells or induce oxidative stress. The free radical mechanism for generating heavy metals is particular (Engwa et al., 2019).

Arsenic: Bacteria, algae, fungi, and humans use methyl-hazardous compounds containing MMA and DMA (DMA). It's the end product of persistent arsenic poisoning.

iAs (V) → iAs (III) → MMA (V) → MMA (III) → DMA (V)

Inorganic arsenic methylation eliminates urine as a bioindication of chronic arsenic exposure via MMA (V) and DMA end products (V). However, MMA (III) remains inside the cell as an intermediary product.

MMA III is a hazardous intermediate product of arsenic that may be responsible for arsenic-induced cancer [7].

Iron: It has biological components like haemoglobin and physiological activities. The human body needs iron. However, as the Fenton reaction illustrates, iron is one of the heavy metals commonly known as radical hydroxyl [8].

$Fe^{3+} + O_2 \rightarrow Fe^{2+} + O_2$
 $Fe^{2+} + H_2O_2 \rightarrow Fe^{3+} + OH \cdot + OH^-$ (Fenton reaction)
 $Fe^{2+} + H_2O_2 \rightarrow Fe^{3+} + OH \cdot + OH^-$ Fenton reaction

Net reaction (Haber-Weiss reaction):

$O_2 \cdot + H_2O_2 \rightarrow OH \cdot + OH^- + O_2$
 $O_2 \cdot + H_2O_2 \rightarrow OH \cdot + O_2$

Besides the aforementioned reactions, the following reactions can occur:

$OH \cdot + H_2O_2 \rightarrow H_2O + H^+ + O_2 \cdot -$
 $OH \cdot + H_2O_2 \rightarrow H_2O + H^+ + O_2 \cdot -$
 $OH \cdot + Fe^{2+} \rightarrow Fe^{3+} + OH^- + Fe^{2+} \rightarrow Fe^{3+} + OH^-$
 $LOOH + Fe^{2+} \rightarrow Fe^{3+} + LO \cdot + OH^-$
 $LOOH + Fe^{2+} \rightarrow Fe^{3+} + LO \cdot + OH^-$

Oxidative iron oxidation occurs frequently. OH[•] can oxidise proteins, lipids, or DNA. FAPy-G is formed when OH[•] interacts with guanine [9].

MOX can harm PUPL. Inculcated. They found OH[•]iron can oxidise lipid membranes (lipid peroxidation).

He hypothesised: Lipoperoxidase:

Initiation: Lipid + R[•] / OH[•] → Lipid[•] Initiation: Lipid + R[•] / OH[•] → Lipid[•]

Propagation: Lipid + O₂ → Lipid-OO[•] Propagation: Lipid + O₂ → Lipid-OO[•]

Lipid-OO[•] + Lipid → Lipid-OOH + Lipid-Lipid-OO[•] + Lipid[•] → Lipid-OOH + Lipid[•]

Termination:

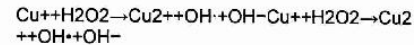
Lipid[•] + Lipid[•] → Lipid-Lipid Termination: Lipid[•] + Lipid[•] → Lipid-Lipid

Lipid-OO[•] + Lipid[•] → Lipid-OO-Lipid Lipid-OO[•] + Lipid[•] → Lipid-OO-Lipid



Early phase radical (R•)/OH• impacts lipid membrane to form radical lipid. This radical lipid promotes radical peroxy lipid formation via dioxygen or lipid reactions. This process also breaks down lipid molecules. Finally, two radical lipid molecules or a radical peroxy lipid molecule engage a stable lipid molecule. Malondialdehyde is the major lipid peroxidation aldehyde and an indication [10]. Proteolytic enzymes require transition metals to break down proteins. This leads to the formation of R•, alcohoxyle (RO•), alkyl peroxy (ROO•), protein-centred alkaline radicals (PCARs), oxidation, loss of itstidin residue, introduction and development of R• [8].

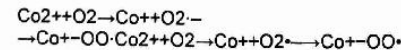
Copper: For cupric (Cu²⁺) and cupric (Cu¹⁺) reactive oxygen (ROS) forms that may lead to oxidation and reduction processes were identified. CU²⁺ can be reduced to Cu⁺ by accessible biological reducers such as Glutathione (GSH) and ascorbic acid which can catalyse H₂O₂ in OH• through the Fenton reaction [11].



The radical OH• generated might react with different biomolecules. Experimental studies have shown that copper can also promote DNA rupture and base oxidation with oxygen-free radicals [5]. Although in vivo study has demonstrated that copper-induced low-density lipoprotein (LDL), in vitro research has demonstrated a definite lack of copper-induced LDL oxidation [12].

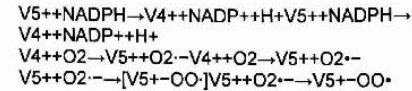
Chromium: Free H₂O₂ radicals are produced by chromium (Cr), particularly Cr⁴⁺ [13]. Chromium has also been demonstrated to identify free radicals in the liver and blood of animals. One electron was reduced to find the intermediates of Cr⁵⁺.

Cobalt: As a result of the H₂O₂ collapse, Cobalt (Co), in particular Co²⁺ has been found to create superoxide (•O₂⁻) [14].

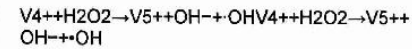


• Vanadium

Vanadium is a heavy metal that produces free radical compounds in various oxidation states. NADPH and ascorbic vanadium (IV) antioxidants diminish vanadium (V) in plasma binding plasma proteins [15].



V(IV) may also form OH• from the breakdown of H₂O₂ under physiological conditions at around 7 pH in accordance with the Fenton reaction.



3. EFFECTS OF HEAVY METAL ON HUMAN HEALTH

Metal, methane, gold, iron, plaster, manganese, mercury, and zinc are heavy metals [16]. A frequent environmental and food heavy metal. Small amounts are necessary for healthy health, while large amounts can be poisonous or hazardous. Intoxication by heavy metals can affect the brain and lungs. Long-term disease exposure, such as MS, Parkinson's, Alzheimer's, and Muscle Dystrophies, may cause physical, muscle, and neurological alterations. Long-term exposure to several metals and chemicals can cause cancer [17]. Some heavy metals are naturally hazardous in the environment. Thorough awareness of heavy metals is essential to avoid excessive interaction [18].

Arsenic: Arsenic contamination happened owing to both natural and human geological processes. Anthropogenic sources of arsenic include mining and processing. Smelting can produce arsenic from both old and new air and soil [19]. These sources may discharge or runoff into surface waters. Groundwater can also be contaminated by arsenic minerals. The sedimentary and meta-sedimentary basis [20]. The element is found in nearly every type of chemical and medication. Some insecticides, fertilisers, and feeds can leach arsenic into the environment. Human health is harmed by inorganic arsenic forms like arsenite and arsenate. It can cause lung, liver, bladder, and skin cancer. Arsenic is found in the air, food, and water. Arsenic is a major source of toxicity in over 30 nations [21]. Where arsenic concentrations in groundwater are above the WHO standard (10 g/L) [22]. Misuse of arsenic compounds, pesticides, or minerals can contaminate water. Chronic toxicity refers to arsenic toxicity. It is based on skin disease diagnosis. Keratosis and pigmentation are distinct skin disorders implying arsenic exposure [23].



Chart 1. Description of heavy metals

Name	Description
Arsenic	Arsenic is a major heavy metal for the environment and humans. It is available as oxides or sulphides of iron, sodium, calcium, copper, etc. Inorganic forms of arsenic, Arsenite and arsenate are toxic to the environment and living things. Natural, industrial, and accidental sources of arsenic in humans. Arsenic insecticides, natural mineral reserves, and inappropriate chemical disposal can contaminate drinking water. Suicidal or accidental intake of arsenic can cause acute poisoning in youngsters. Arsenic is a protoplasmic toxin that targets the sulfuryl group, enzymes, and mitosis.
Lead	Lead is a particularly hazardous metal that has been widely employed in a variety of parts of the world, causing significant environmental and health hazards. Lead is a somewhat blue silver metal in a dry atmosphere. It begins to tarnish upon contact with air and so forms a variable chemical mix depending on the conditions. Industrial processes, food and tobacco consumption, as well as water and household use, are primarily helpful. Petrol and house paint were created by combining tanks, battery storage devices, toys, and mowers.
Mercury	Metallic mercury is a naturally occurring metal that is capable of warming colourless and odourless gas, as well as bright white silver and odourless liquid. Mercury is a highly bio accumulative and toxic substance. Mercury's presence has a detrimental effect on the marine environment, which is why numerous studies focus on the dispersion of mercury in the ocean. Mercury is mostly produced by human activities such as agriculture, municipal wastewater disposal, mining, and cremation.
Cadmium	Cadmium is an ATSDR-category 6 heavy metal. It is a by-product of zinc production that can be consumed by humans and animals. This metal develops in the body following human absorption. Initially, during WWI, the metal was used to replace pigment and tin in paint. It is also used in rechargeable batteries, metal manufacturing, and smoke production. In alkaline batteries, cadmium is utilised for electrode coating, coloration, location, and even plastic stability. This metal is usually inhaled and consumed, causing chronic and acute exposure. In the ecosystem, cadmium stays in soil and sediment for decades. These metals are progressively taken into the human body by plants.
Chromium	The tenth biggest element is chromium. The most common Cr-6 is trivalent Cr-3. Combining oil and coal, ferrochrome oil refractory, pigment oxidants, catalysts, chromium stainless steel, fertilisers, oil wells, and boiling metal tannery. Wastewater and fertiliser discharge chromium into the environment.
Copper	It's a heavy metal used to build copper pipes, wires, and other products. It is also used for making copper and birth control pills intrauterine. Copper sulphate is added to drinking water and pools. Because of anthropogenic and industrial activity it can colonise and absorb soils. Copper is found in almonds, avocados, sprouts and bran.
Manganese	This metal is added to gasoline in the form of Methylcyclopentadienyl manganese tricarbonyl (MMT), and as a result, the petroleum vapours contain a manganese compound that is extremely hazardous.
Nickel	Applied in battery production, nickel plating, machine components, metal placement, steel production, smoking, wire, electric parts, etc. Nickel can cause contact dermatitis in certain persons. Although working with nickel alloys, like stainless steel, is completely safe. But nickel compounds and even metallic nickel must be handled with extreme care as they may cause cancer.
Aluminium	Aluminium is the world's third-largest crust element. It is found in the environment. Aluminium mining and processing is more eco-friendly. Aluminium may be a major cause of human, animal, and plant ailments.
Iron	The second richest metal is Iron. He's the twenty-sixth leader. Iron is crucial to the growth and survival of all living things. It is found in algae, cytochromes, catalases, and oxygen proteins including haemoglobin and myoglobin.



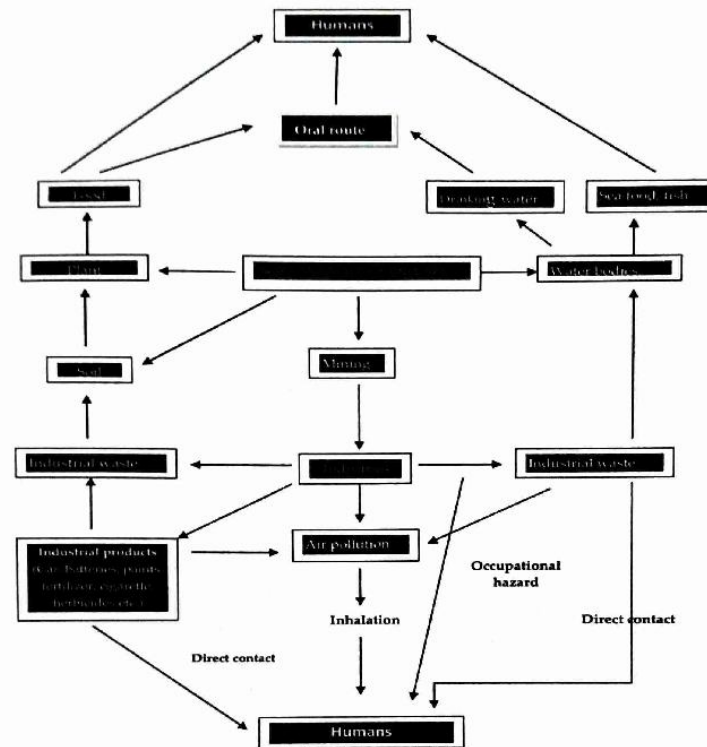


Fig. 1. Heavy metal intoxication mechanisms in humans (source: Engwa et al., 2019)

Blood vessel damage, nose and vomiting, reduced erythrocytes and leucocytes, irregular heartbeat, pricking hands and legs. Long-term contact can cause skin lesions, internal malignancies, neurological difficulties, lung, peripheral vascular, cardiovascular, and diabetes issues [24]. Chronic arsenic poisoning alters permanent organs and increases mortality. While this condition is dangerous, there is no effective cure [25].

Lead: Fossil fuel mining, production, and use have exacerbated air, water, and soil pollution. Batteries, cosmetics, metals, solders, pipelines, etc [23]. Sadly, due to its toxicity, plum is used in paints, gasoline, and other products only to a limited PVC pipes were exposed to purple paint, polluted soil, and industrial [28]. Toxic lead poisoning caused symptoms in the CNS and GI tracts of children and adults [27]. Plum water

toxicity Components of plumbing and water discharge [28]. Lead is toxic (EPA). The body has a huge impact on numerous parts. Feathers are deposited as insoluble phosphate into the skeleton bone by blood flow [29]. Acute or chronic lead toxicity Acute exposure can cause nausea, vomiting, diarrhoea, renal failure, fatigue, insomnia, arthritis, and vertigo. Acute exposure occurs in various workplaces. Feather exposure causes mental retardation and neurodegeneration (brain and kidney damage) [23]. Plum intoxication is dangerous for most organs. When the blood brain block is exposed to more fluid [30]. The main cause is feather ions from the environment and household, although appropriate avoidance may lessen the threat of toxicity [31].

Mercury: Mercury's heaviest. It's a poisoning of mercury. Many industries are damaging the

environment with mercury [32]. Organic and inorganic mercury can be generated by metals. Quecksilver toxicity can harm the foetus [33]. Methyl mercury is high in seafood from 1 to 50 g/. Lipophilicity of fatty and maize liver [34]. Methylene mercury is microbial. Mercury Chloride and Methyl Mercury Carcinogenicity Mercury produces disaster in addition to these symptoms. mercury poisoning causes fright and loss of memory. Inflammation of the lung and heart may occur. Organic mercury causes tiredness, depression and loss of memory.

Cadmium: Cadmium and its compounds affect human health in numerous ways. Caution should be taken when handling cadmium due to its toxicity. Cadmium is reabsorbed by the kidneys. Consumption of cadmium can cause vomiting and GID, while current exposure can cause substantial lung damage and respiratory pain. Causing bone and lung deposition, long-term cadmium exposure Lead exposure causes bone and lung damage. According to studies on both animals and humans, cadmium can cause osteoporosis (skeletal deterioration). There is a bone fracture outbreak in Japan because of the "Itai-itai" disease [35]. Women had more bone fractures, had poorer bone density, and lost less height than males, hence their cadmium toxicity was higher. Exposure to cadmium can cause renal failure. Exposure to cadmium may disrupt calcium metabolism, the production of kidney stones and hypercalciuria. Cadmium is a human category 1 carcinogen in toxicology. Smokers absorb cadmium more than non-smokers since the major source is tobacco [36]. Cadmium may cause atrophy of the testicles and raise prostate cancer risk.

Chromium: Chemists use chromium to make dyes, pigments, It is solid, liquid, or gaseous. Chromium compounds are very stable in water. It can be divalent, 4-valent, 5-valent, or hexavalent. The most stable forms are Cr(VI) and Cr(III), and only human exposure matters [37]. Like calcium chromate, zinc, strontium, and feather chromate, chromium (VI) is a carcinogen. Conversely, chromium (III), an essential dietary mineral for both animals and humans, aids glucose metabolism. Hexavalent chromium compounds are absorbed faster than trivalent chromium compounds. A wide variety of chromium products are available [23]. Schroeder et al. (1970) found 390 g/kg Cr in cigarettes, but no data on chromium inhalation. When chromium compounds encounter broken skin, they create a deep penetrating hole. Exposure to chromium

compounds can cause and treat monthly ulcers. Erythrocyte glutathione reductase inhibitors limit the ability to convert methemoglobin to haemoglobin [38]. Many in vitro and in vivo studies have shown that chromato chemicals can induce DNA damage including DNA adducts, chromosome abnormalities, chromatid exchange for sisters, replication alterations, and DNA transcriptions.

Aluminium: Aluminium is the third most abundant element on Earth. Inhalation, ingestion, and skin contact are all present in one oxidation (3+). Aluminium is found in water, food, drinks, and medications. Of course, food contains aluminium. Aluminium and its compounds are slightly absorbed in humans, however the rate is unknown. The symptoms of a high aluminium content in the body are nausea, mouth ulcers, skin ulcers and arthritic pain. But these are mild and transient effects [39]. The WHO hypothesised in 1997 that aluminium exposure increases the risk of Alzheimer's disease in people. Contact dermatitis and irritating dermatitis have been reported in workers exposed to aluminium. Aluminum has harmed the brain, causing memory loss, balance issues, and coordination issues [40]. Aluminum is difficult to eliminate from the body and might cause bone or brain damage. Aluminum poisoning can be caused by dusty conditions, long-term intravenous feeding, impaired renal function, hemodialysis, or ingestion of high-aluminum compounds. Aluminum can be found in polluted dialysates and phosphate binders. Aluminum exposure can cause secondary hyperparathyroidism, adynamic bone disorder, and osteomalacia [41]. Aluminum toxicity is linked to respiratory issues, anaemia, poor iron absorption, and neurological issues.

Iron: Iron is the most plentiful transition metal. Many essential proteins and enzymes in living beings require a biological cofactor. It aids most aerobic species' breathing. Radiation can interact with biomolecules, cells, tissues, and the entire body if not appropriately segregated. Doctors have long been fascinated by iron toxicity. Many iron-containing products are used on children, causing iron poisoning [42]. Cyclisme de l'ironThe initial stage of iron therapy involves gastrointestinal symptoms like vomiting and diarrhoea [43]. A six-to-twenty-four-hour overdose is termed latent. Between 12 and 96 hours follow the initial clinical manifestation. Lethargy, tachycardia, liver necrosis, improper metabolism, and mortality [44]. Stages 4 and 5



have iron overload. Ulcers and constipation are prevalent. Excess iron consumption boosts cancer risk in wealthy meat-eating cultures. Workers at risk of asbestosis are the second leading cause of lung cancer [45]. Asbestos-related cancer is linked to free radicals. Iron can harm a cell's DNA. Iron oxidises DNA, causing cancer [46]. Iron sulphate, monohydrate, and heptahydrate cause no immediate harm whether consumed orally, cutaneously, or inhaled. The FDA deems iron salts safe, with little negative effects. Iron toxicity [47]. Free radicals create friends like hydrogen peroxide and superoxide in pathological metabolism [48]. Surplus superoxides and hydrogen peroxides react aggressively with free iron, causing oxidative damage [49]. Hydroxyl radicals can destroy enzymes, damage DNA, and oxidise lipids and polysaccharides. It may cause cell death [50-51].

4. CONCLUSION

Heavy metals enter the body via food, water, air pollution, skin contact, and work. Heavy metals like iron and manganese are essential for specialised biochemistry and physiology, but too much can be harmful. The majority of other heavy metals have little to no effect on the body. Heavy metals cause oxidative stress, biomolecule damage (e.g., enzymes, proteins, lipids, and nucleic acids), cancer-causing DNA, and neurotoxicity. Toxicology advances have improved our understanding of the medical consequences of hazardous chemical exposure in humans. Developmental reversals, malignancies, kidney damage, endocrine, immunological, neurological, and other illnesses discovered through research are examples. The current focus of research is on the unique biochemical and molecular processes that contribute to human illness. We looked at how heavy metals like arsenic, lead, and mercury affect the environment and living things like humans. Metals studied included cadmium, chromium, aluminium, and iron. Clear rules, identification of high-concentration heavy metal locations, and adequate laws and regulations are required to protect public health. Heavy metal exposure must be managed to avoid future issues. Technical methods can help limit heavy metal exposure. Keeping track of your exposure to heavy metals in the environment and in yourself can help prevent exposure. Preventing heavy metal toxicity requires international and national cooperation.

CONSENT

It is not applicable.

ETHICAL APPROVAL

It is not applicable.

COMPETING INTERESTS

Author has declared that no competing interests exist.

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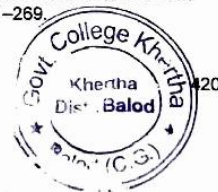
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The requirement for ferric in the initiation of lipid peroxidation by chelated ferrous

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