



## Effects of Cadmium on the Olfactory Bulb of Albino Rat Fed with Zinc Deficient Diet

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### ABSTRACT

*Cadmium, an environmental pollutant, is reported to have toxic effects on nearly all systems of human body though its toxic effects nervous system are less well documented. Its pathophysiological actions are related to body stores of  $Ca^{++}$ ,  $Zn^{++}$  and  $Na^{++}$ . This investigation was conducted to explore the toxic effects of  $Cd^{++}$  on the olfactory bulb of albino rats fed with the  $Zn^{++}$  deficient diet, to have an idea about its corresponding toxic effects on human beings. 16 Charles foster strain rats were taken. 8 rats (control group) and the remaining 8 rats (experimental group) were kept for 30 days on the stock ration and the  $Zn^{++}$  deficient diets, respectively. Experimental group rats were given  $Cd^{++}$  injections. Than  $10\mu$  thick sections of the olfactory bulb were stained with H&E, Thionine and Gleese silver stains. On microscopic observation, various layers of the olfactory bulb (of experimental group rats) showed degenerative changes*

**KEYWORDS :** Cadmium toxicity, Zinc deficiency, Olfactory bulb, Albino rats

### Introduction

Exposure to toxic environmental pollutants with or without nutritional deficiencies is being increasingly recognized as a serious health hazard. Cadmium, a heavy metal is an environmental pollutant.  $Cd^{++}$  is used in industries like plastics, stabilizers, pigments, electroplating alloys, batteries and chemicals and as a component of insecticides, pesticides and fertilizers.  $Cd^{++}$  is toxic to every organ. It is mainly toxic to kidneys and liver. Effects of  $Cd^{++}$  on human nervous system are less well documented. Though CNS dysfunction and anosmia has been reported.<sup>1&2</sup>. Average intake of  $Cd^{++}$  is estimated as 20-50  $\mu$  g/day.

Zinc is required for DNA, RNA and protein synthesis<sup>3</sup> and gene expression<sup>4</sup> for activity of many enzymes. Anencephaly and hydrocephaly have been reported following 'in utero' zinc deficiency in rats<sup>5</sup>. Many pathophysiological actions of  $Cd^{++}$  are related to interaction of  $Cd^{++}$  with body stores of essential ions, especially,  $Ca^{++}$ ,  $Zn^{++}$  &  $Na^{++}$ . Long term  $Cd^{++}$  administration results in increase amount of  $Zn^{++}$  associated with metallothionein<sup>6</sup> resulting into a deficiency of available  $Zn^{++}$  in the body.

Since no experimental studies were available regarding  $Cd^{++}$  neurotoxicity in zinc deficiency states, the aim of this study is to possibly fill the lacuna in the knowledge of this subject.

### Material and Methods

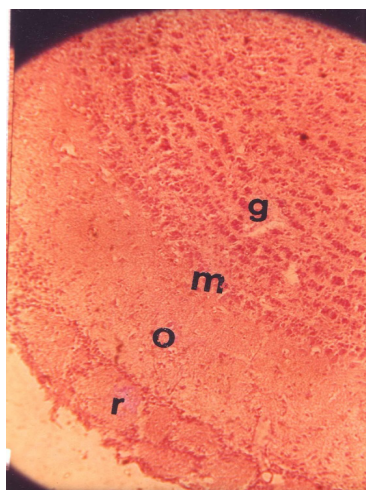
#### Animals

16 Charles' foster strain rats, 8 males & 8 females were taken. They are divided into control and experimental groups, each group containing 8 rats (4 males and 4 females). Control group rats and the experimental group rats were kept for 30 days on stock ration diet and zinc deficient diet, respectively. Experimental groups rats received intraperitoneal  $CdCl_2$  (99% pure anhydrous) injection in the dose of 2mg/kg body weight daily. 24 hours after the last injection, rats were sacrificed. Intra venous infusion was used as perfusion apparatus. 10% formalin solution in normal saline was used as perfusion fluid. In perfusion fixation, rats were anaesthetized by intra peritoneal injection of Nembutal (35 mg/kg body weight). Than thorax was opened, 18 gauge needle was introduced into ascending aorta through the left ventricle. Right atrium was widely opened and perfusion was done by formal saline at the pressure of 5 feet of water pressure. Perfusion was stopped when head & tail stiffness got pronounced and there was oozing of perfusion fluid on cutting the snout with the scissors. After this brain was removed, next olfactory bulbs were cut and put in a fixative (formal saline) over night. Then after usual procedures,  $10\mu$  thick sections were cut and stained with H & E, thionine and glees silver stains.

### Observation and Results

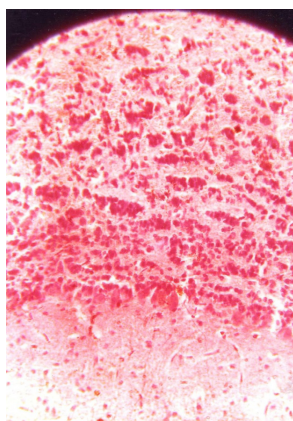
Histological sections of experimental group rats showed histopathological changes in general, but some layers of the olfactory bulb were more conspicuously affected. Mitral cell layer showed the presence of

oedma, clumping of mitral cells and spaces. Lamina fibrosa showed loosening and thickening with infiltration by periglomerular cells. In the lamina glomerulosa, the olfactory glomeruli showed distortion, oedema and spaces while while the periglomerular cells showed necrosis and disorganization.



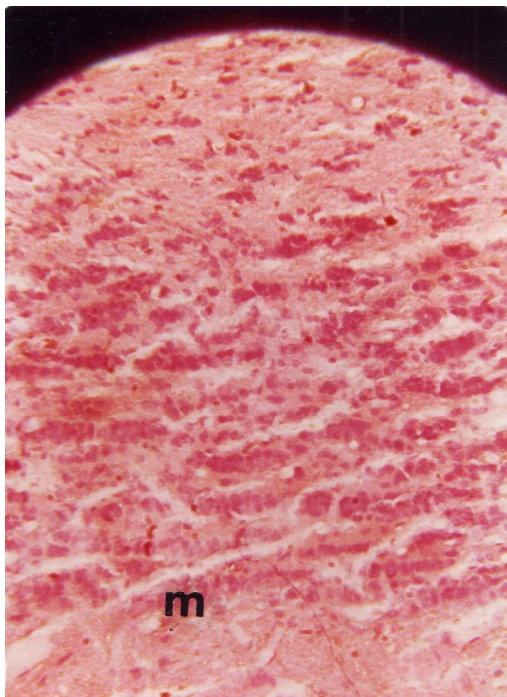
**Fig. C-1. H & E x 200**

Photomicrograph (control) of the olfactory bulb of an albino rat on normal diet and without cadmium exposure.



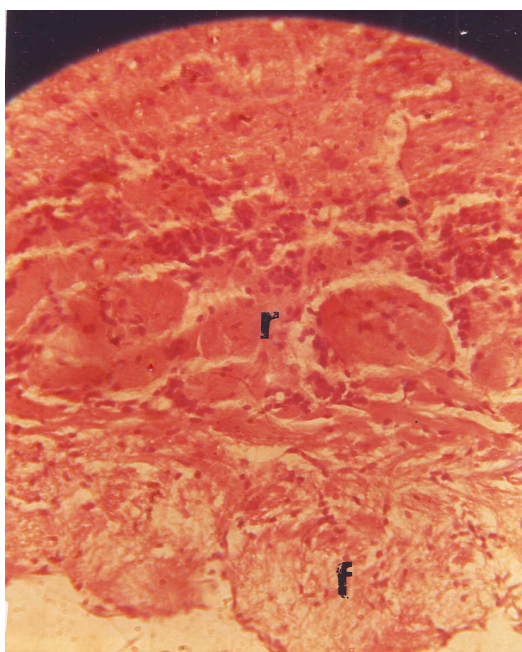
**Fig.C-2: H & E x 400**

Photomicrograph (control) of the olfactory bulb of an albino rat on normal diet and without cadmium exposure.



**Fig.E-1: H & Ex 400**

Photomicrograph (experimental) Showing degeneration of mitral cells and presence of oedema, cell clumping & spaces in the mitral layer



**Fig.E-2: Gleese silver x 400**

Photomicrograph (experimental) Lamina Fibrosa (f) shows loosening, thickening and infiltration by periglomerular cells. The olfactory glo-

meruli (r) shows distortion oedema & spaces. The periglomerular cells show necrosis and disorganization.

## DISCUSSION

Both  $Cd^{++}$  and  $Zn^{++}$  belong to the same group, i.e., II b of the d-block-elements (transitional elements) of the periodic table. Most of the zinc ores contain  $Cd^{++}$  (like calamine). Very little material is available about effects of  $Cd^{++}$  on C.N.S. Reports about  $Cd^{++}$  toxicity on the olfactory bulb are all the more scanty.  $Zn^{++}$  deficiency may result from  $Cd^{++}$  administration<sup>6</sup>. No report is available regarding histological alteration of olfactory bulb secondary to  $Zn^{++}$  deficiency, although disturbance of smell is documented<sup>7</sup>. In the present study, the dose of 2mg/kg body weight of  $CdCl_2$  was chosen in view of the reported 10% mortality with 4mg/kg/body wt. doses of cadmium<sup>8</sup> and reported 14<sup>th</sup> Day  $LD_{50}$  value of a single administration of  $Cd^{++}$  to be 3.55mg/kg/ body weight by the intra peritoneal route<sup>9</sup>.

## Microscopic Findings

Structural alteration in the olfactory bulb was observed following  $Cd^{++}$  intoxication of albino rats fed on  $Zn^{++}$  deficient diet. Olfactory bulbs were found more vulnerable to  $Cd^{++}$  exposure, than other parts of the brain<sup>10</sup>, any specific site of lesion in the olfactory bulb has not been reported by him. No worker has observed the effect of  $Cd^{++}$  exposure on the olfactory bulb of mature rats, neither with concomitant administration of  $Zn^{++}$  deficient diet to the animals, which has been taken under in this study. In the present study, atrophic changes were seen in the lamina fibrosa (loosening, thickening of nerve fibres and eodematous changes). The glomeruli showed distortions, oedematous changes and disorganization of periglomerular cells (appeared irregularly scattered).. Mitral cell layer appeared to be greatly affected and showed clumping of mitral cells, oematous changes and spaces. All these changes under light microscope are possibly due to selective accumulation of  $Cd^{++}$  in the olfactory bulb

## Conclusion

From the above study it was concluded that olfactory bulb is vulnerable to toxicity of Cadmium in zinc deficient state similar to the other parts of the brain and the histopathological changes mainly included oedema, clumping, distortion and other degenerative changes in it's various layers.

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