

zinc can be related to the lower intestinal Hg absorption due to stomach Hg retention. <sup>2</sup> is nephrotoxic, and the preventive effect of

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## Introduction

Mercury toxicity, as well as its deposition in L v H tissues depends on the chemical form (inorganic or organic), time and route of exposure [1]. The main route of exposure to mercury is oral via, in view of the large consumption of contaminated food and water [2,3]. The oxidized form of mercury, Hg<sup>2+</sup>, presents as the

## Exposure to metals

Animals were distributed in four groups (N=4/group), pretreated by gavage for five days with 0.9% NaCl (saline solution) or ZnCl<sub>2</sub> (27 mg/kg/day), and after, treated by gavage for more five subsequent days with saline or HgCl<sub>2</sub> (5 mg/kg/day). Metals were dissolved in saline solution and administered by gavage at a volume of 1 mL/kg body weight. Zn and Hg doses were selected according to previous studies performed by our research group [7-13,20].

## Tissue preparation

Twenty-four hours after the last administration of saline or HgCl<sub>2</sub>, rats were weighed and killed by decapitation. Total blood samples were collected from the body and centrifuged at 1,050 g for 10 min at 4°C to obtain the serum, which was used for determination of urea and creatinine levels and alanine aminotransferase (ALT), alanine aspartate aminotransferase (AST) and lactate dehydrogenase (LDH) activity. For the  $\delta$ -aminolevulinic acid dehydratase ( $\delta$ -ALA-D) activity assay, stomach and intestine were quickly removed, placed on ice and homogenized in 5 and 7 volumes of NaCl (150 mM, pH 7.4), respectively. The homogenate was centrifuged at 8,000 g for 30 min at 4°C and the supernatant fraction (S1) was used in the enzyme assay. Furthermore, the stomach and intestine were used in the determination of mercury levels.

## ALT, AST and LDH activity and creatinine and urea levels

Enzymes activities and creatinine and urea levels were determined by using a Labtest commercial kit as describe in Peixoto and Pereira [11].

### $\delta$ -ALA-D activity

Enzymatic activity was assayed according to Sassa [25] by measuring the rate of product (porphobilinogen - PBG) formation, as previously described by Peixoto et al. [12]. Enzyme activity was expressed as nmol PBG/h/mg protein. Protein concentration was determined by Bradford method [26] using bovine serum albumin as a standard.

## Determination of metal levels

Metal analyses were carried out using a Model AAS EA 5 atomic absorption spectrometer (Analytik Jena, Jena, Germany) equipped with a transversely heated graphite tube atomizer with pyrolytic coated tubes as described by Peixoto et al. [13] and Oliveira et al. [27].

## Statistical analysis

Results were analyzed by one-way analysis of variance (ANOVA)

the technique were considered, for statistical analysis, as containing 0.05 µg of metal/g of tissue, which was the minimum measurable quantity

## Table 2

13. Peixoto NC, Rocha LC, Moraes DP, Bebianno MJ, Dressler VL, et al. (2008) Changes in levels of essential elements in suckling rats exposed to zinc and mercury. *Chemosphere* 72: 1327-1332.
14. Oliveira CS, Joshee L, Zalups RK, Pereira ME, Bridges CC (2015) Disposition of inorganic mercury in pregnant rats and their offspring. *Toxicology* 335: 62-71.
15. Oliveira CS, Joshee L, Zalups RK, Bridges CC (2016) Compensatory renal hypertrophy and the handling of an acute nephrotoxicant in a model of aging. *Exp Gerontol* 75: 16-23.
16. Berlin M, Zalups RK, Fowler BA (2007) *Mercury: Handbook on the Toxicology of Metals* (3rd edn.). Elsevier, Amsterdam, Netherlands.
17. Agarwal R, Behari JR (2007) Effect of selenium pretreatment in chronic mercury intoxication in rats. *Bull Environ Contam Toxicol* 79: 306-310.
18. Agarwal R, Goel SK, Chandra R, Behari JR (2010) Role of vitamin E in preventing acute mercury toxicity in rat. *Environm Toxicol Pharmacol* 29: 70-78.
19. Rao MV, Chhunchha B (2010) Protective role of melatonin against the mercury induced oxidative stress in the rat thyroid. *Food Chem Toxicol* 48: 7-10.
20. Moraes-Silva L, Bueno TM, Franciscato C, Oliveira CS, Peixoto NC, et al. (2012) Mercury chloride increases hepatic alanine aminotransferase and glucose 6-phosphatase activities in newborn rats in vivo. *Cell Biol Int* 36: 561-566.
21. Saper RB, Rash R (2009) Zinc: An essential micronutrient. *Am Fam Physician* 79: 768-772.
22. Franco JL,