

ERYTHRON RESPONSE TO PCB INTOXICATION

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Introduction

Dioxins are a class of highly toxic and broadly dispersed environmental contaminants. Dioxins include 75 dibenzodioxins, 135 dibenzofurans and 20 biphenils. According to their chemical structure polychlorinated biphenyls (PCB) belong to a class of polyhalogenated cyclic hydrocarbons. PCB are widely used in agriculture and family life as dielectric liquid in transformers and condensers. PCB can enter the human body through the skin and gastrointestinal tract (V.N.Maistrenko,1998).

PCB are lipophilic; they are easily absorbed in a lipid tissue and a liver. Sometimes more toxic metabolites can appear. The intensity of peroxidic lipid oxidation processes is increased, antioxidant activity of blood and cells is decreased; it leads to the erythrocytes membrane destruction during affected by PCB (A.F.Kaioumova,1996). An increased erythrocytes hemolysis brings about tissue hypoxia, immunological distress, free radicals formation and peroxide lipid oxidation activation.

Methods and Materials

The aim of this investigation is to analyse of the variable character in the erythron central chain when exposed to PCB in 0,1 and 0,05 LD₅₀ dose. 100 white non pedigree male rats (150-200 gr.) were at different doses of PCB. 0,1 LD₅₀ dose of PCB was introduced into the stomach during 28 days (in the form of olive oil) in group 1. The group 2 of animals was also subjected to PCB by the probe during the same time and the total dose was 0,05LD₅₀. The control group consumed the equivalent amount of olive oil. All the animals were killed on the 1-st and 28-th day of an observation. Peripheral blood findings examination was carried out by means of common methods after 24 hours of the beginning of toxicants infusing and on the 28-th day. Distribution calculation and classification of erythroblastic islands (EI) into 5 classes was done according to Y. M. Zakharov (1990). Class 1 contains 2-8 erythroid cells in the "crown" since CFUe differentiation into proerythroblast and the subsequent division have been performed. Class 2 - from 9 to 16 cells; class 3 - more than 16 nuclear erythroid cells. Normoblasts and reticulocytes, unable to be fissioned, are concentrated in "crown" in involution class. A class of EI reconstruction possesses a division capable cells (pro- and basophilic erythroblasts) and undividable cells. CFUe involvement estimation into the process of proerythroblasts differentiation, a chance of reinvolvement of central macrophage (CMPh) into a new wave of erythropoiesis were carried out according to Yu. M. Zakharov and L.V.Vorgova (1990).

Results and Discussion.

A statistical decrease of erythrocytes, hemoglobin and hematocrit was noted when exposed PCB. Erythrocyte amount in group 1 was $3,76 \pm 0,19 \times 10^{12}/l$ ($p < 0,001$) by the end of the first day, group 2 - $4,24 \pm 0,56 \times 10^{12}/l$ ($p < 0,01$), while the control parameter was $5,48 \pm 0,6 \times 10^{12} /l$. Group hemoglobin content decreased up to $132,0 \pm 8,0$ gr/l ($p < 0,001$), group 2 - $140 \pm 8,04$ gr/l ($p < 0,001$),

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while the control data was $175 \pm 9,09$ gr/l. Hematocrit was also under chance: it was a tendency to be decreased up to 41 % in group 1 and 45 % in group 2, while the control index was 49 % ($p < 0,01$).

Erythrocytes destruction changing values being exposed to the inserted preparations and their peripheral blood metabolites during 24 hours PCB injection can be explained by the fact, that erythrocytes are being heavily destroyed under the influence of the injected preparations and their metabolites.

According to A. Kaioumova (1996), toxicants and metabolites have both direct and indirect influence on erythrocytes membrane. Their impact concerns the erythrocyte membrane phospholipids peroxidation increasing. The fact is that the lipid oxidation products are accumulated in the body (when being intoxicated), antioxidant activity of tissue and blood decreases, erythrocyte life is decreased, and the time of their destruction increases.

On day 28 we could observe the restoration of the peripheral blood indices in both groups. Peripheral blood analysis has confirmed this fact. Erythrocytes amount in group 1 was $6,18 \pm 0,36 \cdot 10^{12}/l$. The second group: $5,5 \pm 0,66 \cdot 10^{12}/l$ ($p < 0,05$); hemoglobin respectively – $150 \pm 12,0$ gr/l and $151 \pm 12,21$ gr/l ($p < 0,05$).

The obtained indices can be explained by the adaptation of an erythron to hemolytic action of PCB and their metabolites, marrow erythrocytes reproduction increase which is stable to hemolytic action. The accelerated process of erythrocytes, destruction is accompanied with their marrow reproduction disfunction when affected by PCB. This fact is confirmed by the following data: EI/femur bone absolute index is decreased; an impeded CFUe differentiation into proerythroblasts, according to the EI first class and newly reconstructed EI; impeding of the amplification wave and erythroblasts maturing in the proliferative EI classes. EI erythropoiesis impeded reconstruction was combined with marrow erythropoietin inactive MF functional activity impediment: their impediment to a repeated complexity with CFUe and their lysosomal laminar complex. The above mentioned changings were evidently shown in group 1 (Table 1,2).

Ecotoxicant effect expressed in erythrocyte suppression in EI possibly reflects their property to suppress intracellular interaction mechanisms. Increasing activity of proteolytic enzymes free radical oxidation and lipid peroxids damaging the formation of EI erythropoiesis is likely to be the underlying cause intracellular interaction damage in EI exposed to PCB.

References

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Table 1. Distribution of erythroblastic islands of different bone marrow maturity class (%) in different groups of rats (M±m, p)

| Groups of Rats | EI ×10 ³ / femur | Distribution of erythroblastic islands (%) | | | | |
|----------------|-----------------------------------|--|----------------------|-----------------------|-----------------------|-----------------------|
| | | EI1 | EI2 | EI3 | EIrec | EIinv |
| Control | 120±42.1 | 4±2.84 | 12±3.17 | 25±0.89 | 15±2.39 | 44±3.39 |
| Group 1 | 95±16.4 p < 0,05 | 6.4±1.6 p < 0,05 | 20.4±5.8 p < 0,05 | 18.4±4.3 p < 0,05 | 24.4±1.02 p < 0,05 | 30.4±5.4 p < 0,05 |
| Group 2 | 124±22.1 p < 0,05 | 0 | 0 | 50.6±5.7 p < 0,001 | 6.4±1.11 p < 0,01 | 42.6±3.11 p < 0,05 |

Note: p is calculated with regard to control indexes at each period of study

Table 2. Functional indexes of erythropoiesis in EI in rats in different groups (M ±m).

| Index | Total CFU-E number involved in EI differentiation EI ₁ + EI ₂ + EI ₃ + EIinv + 2EIrec × 10 ³ /femur | CFUe involvement amount in erythroid differentiation EI ₁ + EIrec (×10 ³ /femur) | Index of repeated EI CM in Volvment in erythropoiesis: EIrec/Eiinv |
|---------|---|--|--|
| Control | 138,2 ± 2,86 | 23,1 ± 0,86 | 0,36 ± 0,06 |
| Group1 | 123,9 ± 2,5 p<0,05 | 35,6 ± 0,4 p<0,05 | 1,7 ± 0,05 p<0,05 |
| Group2 | 131,94 ± 1,16 p<0,001 | 7,94 ± 1,16 p<0,001 | 0,18 ± 0,05 p<0,001 |

Note: p is calculated with regard to the control indexes at each period of study.

