Capabilities of combination of ferroglukin, fosprenil and hamavit in correction of hemostasis functional state of new-born calves having iron deficit

ABSTRACT

New-born calves are still often found with iron deficit. This state negatively influences their growth and development to some extent due to hemostasiopathy development. In this connection great scientific and practical significance for veterinary science and physiology belongs to the search of approaches to effective correction of new-born calves’ hemostasiopathy in lack of iron conditions. It seemed to be perspective to evaluate the degree of influence of traditionally applied at iron deficit ferroglukin in combination with metabolism stimulators (fosprenil and hamavit) on new-born calves’ hemostasis system indices. During our research it was established that new-born calves with iron deficit also have decreased plasma antioxidant protectability, intensification of lipids’ peroxidation processes, increase of thrombocytes’ hemostatic activity and system of blood coagulation system along with decrease of vascular wall’s ability to bind it. In our study we found that combination of ferroglukin, fosprenil and hamavit given to new-born calves with iron deficit showed improved plasma antioxidant and lipid peroxidation activity. Also normalization of thromboocyte activity and positive dynamics of hemostasis vascular and plasma components was observed. Iron deficit of new-born calves can be considered as the model of hemostasiopathy abnormality. With its help we can try different means and their combinations to remove hemostasiopathy. Received results allow us to consider the usage of combination of fosprenil and hamavit on the background of ferroglukin to be unsufficient for removal of hemostasiopathy in new-born calves with iron deficit.

Key words: new-born calves; iron deficit; hemostasis system; ferroglukin; fosprenil; hamavit.

1. INTRODUCTION

Durable and planned investigations of cattle physiology are directed at acceleration of processes of its growth, increase of productivity [1], working out of modern prophylaxis of different abnormalities and creation of approaches for full removal of appeared pathology [2]. It’s very important to accumulate knowledge about different sides of calves’ physiological processes at the very beginning of their ontogenesis [3] because of special significance of the phase of birth for laying the foundation for milk and meat productivity of cattle. At present it is assumed that the leading role in support of hemostasis during the whole ontogenesis belongs to blood and its hemostatic mechanisms influencing heavily the way of individual development [4,5] with the help of hemocirculation processes. So, activity changes of hemostasis processes change activity of hemocirculation in tissues and organs, and, thus influence the common state of a body [6]. In previous researches it was found that at hemostasis deviations especially in case of a young organism there can happen quick increase of hemostasis components’ activity able to lead to microcirculation disturbance [7]. Basic works in this field were conducted on a human being [8,9,10]. Being based on their results we managed to make a conception of the presence of age-specific dynamics of hemostasis components’ activity [11,12], most vulnerable its mechanisms and the potential of different influencing variants on a body aiming at hemostatic processes optimization. Because of great social significance of thrombosis development at cardial pathology [13,14] many researchers’ attention is still devoted to haemostatic changes in given patients [15,16,17]. They investigate different aspects of hemostasiopathy pathogenesis of cardial diseases. As a result, the possibility of its correction was found. It can be done not only with the help of medicines [18,19], but also with the help of traditionally applied impacts [20,21]. It has great significance for biological investigations as it allows to minimize application of medicines.

Taking it into consideration, we have large scientific and practical interest to estimation of hemostasis state of new-born calves in conditions of different dysfunctions and search of approaches to correction of a body’s state which can positively influence the signs of hemostasiopathy. We thought to be perspective to take iron deficit as a model of hemostasis abnormality, as it is often found in new-born calves, and try to begin working out of effective way of correction of all the hemostasis system components. Given model seemed to be approved as it aims at decrease of hemoglobin content in blood and activity lowering of iron-bearing enzymes which suppress protein synthesis and activity of cellular functions [22]. In this connection iron deficit was estimated as a state accompanied by changing of the whole body’s activity. It also forms abnormalities of all the components of hemostasis system.

In this connection investigations aimed at removal of hemostasiopathy in those having iron deficit have great scientific and practical significance. Worked out at the given state variants of evidence
decrease of hemostasis disturbances can serve the basis for the following creation of correction complexes able to be effective in the field of hemostasiopathy reduction of new-born calves at many diseases. Clearing out the impact of combination of iron-bearing substance [23] and fosprenil [24] and hamavit [25] (which showed earlier their high biological activity as far as separate hemostasis components are concerned) has serious perspectives for removal of iron deficit. In this connection we put the following aim for our investigation—to find the evidence of hemostasis system activity correction of new-born calves with iron deficit with the help of ferroglukin, fosprenil and hamavit combination.

2. MATERIALS AND METHODS

2.1 Materials

The work was fulfilled with 34 newborn calves having the signs of erythrogenesis and decrease of iron content in their bodies (serum iron 12.3±0.10 umol/l, siderocytes 1.6±0.05%, hemoglobin 95.0±0.29 g/l, erythrocytes 4.1±0.13x10^12/l). The control group contained 29 healthy new-born calves. All the investigations in the present work were conducted in full correspondence with ethical norms and recommendations on humanization of work with laboratory animals containing “The European Convention on the protection of vertebrate animals used for experiments or in other scientific purposes” (Strasbourg, 1986).

2.2 Methods

The state of lipids’ peroxidation (LPO) in animals’ plasma was found out according to the quantity of thiobarbituric acid—active products in it with the help of a programme packet “Statistika for Windows v. 6.0”, (the 7th day of life). The results were processed by Student’s criterion (t). Statistical processing of received information was made with the help of a programme packet “Statistika for Windows v. 6.0”, “Microsoft Excel”. Differences in data were considered reliable in case p<0.05.

RESULTS AND DISCUSSION

Examined newborn calves with iron deficit were found to have characteristic for the given state weakness, limpness, absence of interest to the environment, paleness of rhinosecope and slime layers. These animals were noted to have increased LPO activity in plasma (acylhydroperoxide 3.46±0.012 D₃₀₀/₅₀ ml, thiobarbituric acid—active products 5.19±0.019 umol/l at value depression of blood liquid part antioxidant activity 22.0±0.23%). The values of these indices under control were equal to 1.45±0.010 D₃₀₀/₅₀ ml, 3.46±0.012 umol/l and 33.7±0.15%, correspondingly.

The state of plasma hemostasis was evaluated according to duration of activated partial thromboplastinic period (APTP), prothrombinic period and thrombinic period with the help of generally accepted methods [27].

Disaggregation capabilities of a vascular wall were defined with the help of a probe with temporal venous occlusion on the basis of visual micromethod of thrombocyte aggregation registration [10] with all the applied inductors. We calculated the value of vascular wall disaggregation activity index (VWDAI) while dividing thrombocytes aggregation period on the background of venous deadlock on the time of thrombocytes aggregation appearance without it. The index value of vascular wall anticoagulation activity of examined calves was also calculated by dividing of antithrombin III activity after venous occlusion on its value before it [27]. Vascular control over fibrinolytic blood activity was found out by calculating of index value of vascular wall fibrinolytic activity while dividing the time of euglobulinlysis before occlusion on lysis time after it [27].

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The correction of iron deficit state of new-born calves was realized by ferroglukin intramuscularly, once from the calculation of 15mg of iron on 1kg of body mass, fosprenil 0.1mg/kg intramuscularly in the morning in the scheme of liquid feeding during 6 days and hamavit—0.1ml/kg intramuscularly in the scheme of liquid feeding during 6 days beginning simultaneously with ferroglukin application. Evaluation of healthy animals’ state was made two times—at their birth and on the 7th day of life. Because of the absence of reliable differences between the results of both investigations control values of each index are presented by one figure—a simple average between them. Examination of calves having iron deficit was fulfilled twice—at their birth and on the next day after correction finish (the 7th day of life). The results were processed by Student’s criterion (t). Statistical processing of received information was made with the help of a programme packet “Statistika for Windows v. 6.0”, “Microsoft Excel”. Differences in data were considered reliable in case p<0.05.

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Intensification of activation of outer mechanism of plasma hemostasis starting and had as its basis found acceleration of prothrombin period of newborn calves with iron deficit pointed at their evident depression of anticoagulant – antithrombin III and tissue activators of plasminogen in them. Noted to have weakening of vessels’ anticoagulant and fibrinolytic capabilities because of production their walls of prostacyclin and nitric oxide molecules [38]. At the same time examined calves were at lowering of vessels’ disaggregation features. It was evidently caused by generation decrease in processes of arachidonic acid with surplus thromboxan A2 formation [37] in their blood plates. Realization of genetically defined growth and development processes of living organisms takes place at constant influence of numerous factors of environment and internal environment [28] on an organism. Physiological peculiarities of their influence are mostly expressed by the optimum of living beings’ blood content [29] especially as far as hemostasis system components’ activity is concerned [30]. Besides, any disturbances in an organism are accompanied by negative dynamics of hematological indices [31,32] including parameters of hemostasis system [33,34]. It becomes clear, that in the basis of hemostasiospathy development in case of examined newborn calves we have not only iron deficit but also found during investigation depression of plasma antioxidant protection which, as previous works showed, causes LPO activation in it. Increase of peroxidation in plasma damages structures of blood plates and vessels and affects their functions [35,36]. Found in newborn calves with iron deficit thrombocytes’ aggregation acceleration points at the increase of their receptors’ structures of blood plates and vessels and affects their functions [35,36]. Newborn calves with iron deficit having received iron preparation in combination with metabolically active means were noted to have a tendency to the index increase of vascular wall anticoagulant activity and the index rise of vascular wall fibrinolytic activity on 9.5%. Because of realization of complex application we reached APTT slowdown on 42.6% at simultaneous decrease of prothrombin period on 42.6%. It allowed them to get normalized. Besides, the value of thrombin period, defining the activity of fibrinogen transition into fibrin, of these calves increased on 13.0%. New-born calves with iron deficit were also characterized by APTT increase (on 42.3%) and prothrombin period (on 4.6%) combining with some intensification of thrombin period (on 8.7%). Realized state correction provided examined calves with iron deficit improvement of the common state and their activity, increase of their serum iron level to the control values (29.3±0.12 umol/l). On the background of ferroglukin, fosprenil and hamavit combination examined calves were found to have evident plasma content decrease of acyhydroperoxides (2.16±0.010 D233/1 ml, p<0.01) and thiorbitaric acid active products (4.12±0.014 umol/l, p<0.01) at the increase of antioxidant activity to 29.1±0.09% (p<0.01). Correction realization of animals having at the beginning iron deficit was accompanied by invariability of thrombocytes’ quantity in their blood and some slowdown of thrombocytes’ aggregation. Besides, animals’ thrombocytes most actively responded by aggregation to collagen, ADF and ristomicin, less actively – to thrombin and adrenaline addition into plasma (table). Examined calves in the result of realized application were noted to have evident VWDAI increase in relation to all the applied inductors (table). The minimum value was VWDAI value with thrombin. Other VWDAI values were a bit higher and had a tendency of approaching the control values. 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Acceleration of blood coagulation final stage pointed at the intensive change of fibrinogen into fibrin in case of examined calves [17].

Application of ferroglukin, fosprenil and hamavit combination made new-born calves with iron deficit state feel saturation of their bodies with iron, positive dynamics of red blood and common animals’ state indices. Fulfilled impact on examined calves’ bodies was accompanied by lowering of their LPO processes intensity in plasma what weakened its damaging influence on endothelium and liver thrombocytes. Found weakening of thrombocytes’ aggregation of calves with iron deficit state after getting ferroglukin, fosprenil and hamavit combination is mostly the consequence of positive impact of these means’ combination on innerthrombocyte LPO, receptor and postreceptor thrombocytes’ functioning mechanisms [38]. Developing in these conditions time increase of thrombocytes aggregation coming in response to ristomicin, pointed at lowering of adhesion cofactor – von Willebrand factor [19] in these calves’ blood.

In the result of applied impact animals having at the beginning iron deficit got some strengthening of disaggregation, anticoagulant and fibrinolytic vessels/features. In the basis of found changes there was not reached the control level of production intensification of prostacyclin, nitric oxide, antithrombin III and tissue activator plasminogen [29] in vascular endothelium of these calves.

In the result of fulfilled investigation it becomes clear that in case of application of ferroglukin, fosprenil and hamavit combination to new-born calves with iron deficit, we can provide normalization of hemocoagulation and positive dynamics of the rest hemostasis system components.

3. CONCLUSION

Iron deficit of new-born calves can be considered as the model of hemostasis abnormality. With its help we can try different means and their combinations to remove hemostasiopathy. It is caused by the facts that new-born calves having iron deficit are characterized by lowering of blood plasma antioxidant protection, intensification of LPO processes in it, increase of thrombocyte hemostatic activity and hemocoagulation at depression of vascular wall capabilities to slowdown of these processes. Because of the presence of iron deficit in new-born calves it is approved to use iron preparation. For strengthening of its impact on a body there were prescribed stimulating metabolism and anabolism means – fosprenil and hamavit. In our work we found that in case of their application to new-born calves it’s possible to strengthen antioxidant protection, weaken LPO activity, decrease thrombocyte activity, produce positive dynamics of hemostatic features of vascular wall and normalize plasma hemostasis. Received results allow us to consider the usage of combination of fosprenil and hamavit on the background of ferroglukin to be insufficient for removal of hemostasiopathy in new-born calves with iron deficit.

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40. Table. Parameters of hemostasis in newborn calves with iron deficiency treated with ferroglukin, fosprenil and hamavit

<table>
<thead>
<tr>
<th>Consider indicators</th>
<th>Calves with iron deficiency, n=34, M±m</th>
<th>control, n=25, M±m</th>
</tr>
</thead>
<tbody>
<tr>
<td>platelet aggregation with ADP, s</td>
<td>25.0±0.10</td>
<td>29.6±0.05</td>
</tr>
<tr>
<td>platelets’ aggregation with collagen, s</td>
<td>19.8±0.15</td>
<td>24.9±0.04</td>
</tr>
<tr>
<td>platelets’ aggregation with thrombin, s</td>
<td>37.9±0.21</td>
<td>46.6±0.16</td>
</tr>
<tr>
<td>platelets’ aggregation with ristomycin, s</td>
<td>22.5±0.16</td>
<td>38.6±0.07</td>
</tr>
<tr>
<td>VWDAI with ADP</td>
<td>1.44±0.003</td>
<td>1.58±0.003</td>
</tr>
<tr>
<td>VWDAI with collagen</td>
<td>1.33±0.005</td>
<td>1.46±0.008</td>
</tr>
<tr>
<td>VWDAI with thrombin</td>
<td>1.38±0.007</td>
<td>1.47±0.004</td>
</tr>
<tr>
<td>VWDAI with ristomycin</td>
<td>1.40±0.004</td>
<td>1.43±0.009</td>
</tr>
<tr>
<td>index value of vascular wall anticoagulation activity</td>
<td>1.23±0.006</td>
<td>1.28±0.005</td>
</tr>
<tr>
<td>index value of vascular wall fibrinolytic activity</td>
<td>1.23±0.009</td>
<td>1.30±0.006</td>
</tr>
<tr>
<td>activated partial thromboplastin period, s</td>
<td>27.0±0.29</td>
<td>39.8±0.33</td>
</tr>
<tr>
<td>prothrombinic period, s</td>
<td>12.2±0.25</td>
<td>17.4±0.30</td>
</tr>
<tr>
<td>Parameter</td>
<td>Control</td>
<td>Initial State</td>
</tr>
<tr>
<td>---------------------------</td>
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</tr>
<tr>
<td>Thrombinic Period, s</td>
<td>15.8±0.19</td>
<td>17.3±0.15</td>
</tr>
<tr>
<td>p&lt;0.05</td>
<td>p&lt;0.05</td>
<td></td>
</tr>
</tbody>
</table>

Legend: p - reliability of differences of indicators between the control and the initial state of the calves with iron deficiency; p<0.05 – reliability of dynamics of indicators in calves with iron deficiency on the background of correction.