ABNORMAL FINDINGS IN HAEMOGRAMS OF DACHSHUND PUPPIES: PRESUMPTIVE (IMMUNODEFICIENCY) FAMILIAL DISEASE

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1. INTRODUCTION

1.1. Case Presentation

In a litter of seven Dachshund puppies, one was stillborn and six presented with diffuse skin infections characterized by dermatitis, abscessation, pustules, crusts and ulcers. Two of the puppies were referred for further evaluation. One male puppy was referred at four months of age and a sister litter mate was referred two months later. A complete blood count, biochemical and histological examination were performed on these puppies to identify the pathologic process. Clinical, biochemical, haematological and histological evaluation of subjects. This report characterizes quantitative and qualitative haematological abnormalities in two puppies that resulted in a diagnosis of dysmyelopoiesis. The existence of a familial immunodeficiency syndrome was speculated. This is the first report of such a syndrome in Dachshund puppies.

Keywords: Dysmyelopoiesis, Blood Smear, Neutrophil Function Disorder, Dachshund

ABSTRACT

In a litter of seven Dachshund puppies, one subject was stillborn and six presented with diffuse skin infections characterized by dermatitis, abscessation, pustules, crusts and ulcers. Two of the puppies were referred for further evaluation. One male puppy was referred at four months of age and a sister litter mate was referred two months later. A complete blood count, biochemical and histological examination were performed on these puppies to identify the pathologic process. Clinical, biochemical, haematological and histological evaluation of subjects. This report characterizes quantitative and qualitative haematological abnormalities in two puppies that resulted in a diagnosis of dysmyelopoiesis. The existence of a familial immunodeficiency syndrome was speculated. This is the first report of such a syndrome in Dachshund puppies.

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1. INTRODUCTION

1.1. Case Presentation

In a litter of seven Dachshund puppies, one was stillborn and six presented with diffuse skin infections characterized by dermatitis, abscessation, pustules, crusts and ulcers. Two of the puppies were referred for further evaluation.

1.2. Case 1

A four-month old male presented with fever, lethargy, anorexia, vomiting, diarrhea, dermatitis and conjunctivitis. A Complete Blood Count (CBC) showed: mild normocytic hypochromic non-regenerative anemia (RBC 4.11×10¹²/L, Reference Interval (RI) 6-9×10¹²/L; hemoglobin 72 g L⁻¹, RI 150-190 g L⁻¹; hematocrit 26%, RI 38-55%; MCV 63 fL, RI 60-77 fL; MCHC 28 g dL⁻¹, RI 31-34 g dL⁻¹; reticulocytes 41400 µL⁻¹ RI >60.000), mild leukocytosis (15.6×10⁹ L⁻¹ RI 6-12×10⁹ L⁻¹) and mild thrombocytopenia (113×10⁹ L⁻¹ RI 150-500×10⁹ L⁻¹). The differential leukocyte count was: segmented neutrophils 2652 µL⁻¹ RI 3000-10000 µL⁻¹, eosinophils 624 µL⁻¹ RI 0-600 µL⁻¹, lymphocytes 3900 µL⁻¹ RI 1000-4000 µL⁻¹ and atypical cells 8586 µL⁻¹ RI 0 µL⁻¹.

Microscopic evaluation of the blood smear revealed the presence of keratocytes, schistocytes, anulocytes and severe signs of dysplasia in both neutrophils and monocytes. The dysplasia was characterized by abnormal nuclear shape, hypersegmented and/or hyposegmented nuclei, irregular chromatin pattern and atypical cytoplasmic granules. The platelets were dysplastic to an extent that they were initially mistaken for blood parasites, because most of them had a tail-like structure (Fig. 1 and 2). Clinical signs of altered primary hemostasis were not evident. For this reason, PCR screening for Trypanosoma spp. was performed, but it was negative. Cytological diagnosis of dysmyelopoiesis was made.

The biochemistry panel was characterized by mild hypoalbuminemia (25 g L⁻¹ RI 32-46 g L⁻¹) and mild hyperglobulinemia (38 g L⁻¹ RI 15-35 g L⁻¹) with increase in acute phase protein fractions and a mild increase in alkaline phosphatase (364 U L⁻¹ RI < 81 g L⁻¹) and creatine phosphokinase (526 U L⁻¹ RI < 180 U L⁻¹). The mild hypoalbuminemia and hyperglobulinemia
with increase in acute phase protein fractions was attributed to the persistent infection.

The dog died three days after referral and before bone marrow sampling could be attempted. Necropsy and histopathological examinations were carried out and the results showed: multifocal, necrotizing to suppurative hepatitis, neutrophilic lymphadenitis of the mesenteric lymph nodes, ulcerative enteritis of the small intestine and extramedullary hematopoiesis in the spleen. The histological findings were suggestive of septicemia.

### 1.3. Case 2

A six-month-old female presented with fever, lethargy, anorexia, vomiting, diarrhea and diffuse skin granuloma. The CBC showed: mild normocytic, hypochromic non regenerative anemia (RBC 4.04×10¹² L⁻¹, RI 6-9×10¹² L⁻¹; hemoglobin 82 g L⁻¹, RI 150-190 g L⁻¹; hematocrit 31%, RI 38-55%; MCV 76 fl, RI 60-77 fl; MCHC 26 g dL⁻¹, RI 31-34 g dL⁻¹; reticulocytes 38000 μL⁻¹, RI >60.000) and moderate leukocytosis (35.3×10⁹ L⁻¹, RI 6-12×10⁹ L⁻¹). The platelet count was normal (210×10⁹ L⁻¹, RI 150-500×10⁹ L⁻¹). The differential leukocyte count was: eosinophils 706 μL⁻¹, RI 0-600 μL⁻¹, lymphocytes 10590 μL⁻¹ RI 1000-4000 μL⁻¹, atypical myelo-monocyte cells 22592 μL⁻¹ RI 0 μL⁻¹ and blasts 1412 μL⁻¹ RI 0 μL⁻¹. The atypical myelo-monocytic cells were characterized by a large size, abnormal nuclear shape, irregular clumped chromatin, rarely visible nucleoli and a large amount of cytoplasm, often containing atypical granules (Fig. 3). Blasts were characterized by medium or large size, nucleus round and convoluted, irregular granular chromat, one or more nucleoli and large amount of basophilic cytoplasm. The platelets were often very dysplastic (Fig. 4). It is important to emphasize that even though a severe platelets dysplasia was noted, no specific clinical signs such as petechiae were observed. This suggests that the morphological alterations did not have a significant impact on function. The cytologic interpretation suggested a dysmyelopoiesis syndrome.

The biochemistry panel was characterized by mild hypoalbuminemia (28 g L⁻¹ RI 32-46 g L⁻¹) and mild hyperglobulinemia (45 g L⁻¹ RI 15-35 g L⁻¹) with increase in acute phase protein fractions.

As the dog’s clinical condition continued to worsen, she was euthanized at 8 month of age.

Unfortunately, permission for cytological examination of the bone marrow could not be obtained. This prevented the definitive diagnosis of a dysmyelopoiesis.

Further information on the other litter mates was supplied by the referring practitioner.
Fig. 4. Case 2. Atypical platelet characterized by the presence of “tail” and resembling blood parasites. May Grunwald Giemsa stain (100x)

1.4. Case 3-4

Another puppy from the same litter presented with similar skin lesions and died four months after birth. A diagnosis of peritonitis and suppurative mesenteric lymphadenitis was made following necropsy. Another puppy also presented with similar skin lesions in addition to a severe diarrhea. She was euthanized at five-months of age and numerous abscesses were found in the liver and mesenteric lymph nodes during necropsy.

1.5. Case 5-6

These subjects presented with similar clinical symptoms but died before they could be referred to the hospital.

2. DISCUSSION

Dysmyelopoiesis is defined as a hematologic disorder characterized by the presence of morphologic abnormalities (i.e., dysplasia) in one or more hematologic cell lines in the blood or bone marrow (Weiss, 2006; Weiss and Wardrop, 2011). Dysmyelopoieses are divided into myelodysplastic syndromes, secondary dysmyelopoiesis and congenital dysmyelopoiesis. Causes of dysmyelopoiesis include acquired mutations in hematopoietic stem cells (i.e., myelodysplastic syndromes), congenital defects in hematopoiesis and dysmyelopoietic conditions associated with various disease processes, drug treatments, or toxin exposure (Weiss, 2005). The congenital forms, causing functional defects, have been well described in some canine breeds: e.g., Canine Leucocytes Adhesion Deficiency (CLAD) in Irish Setter and Irish setter-cross-bred, chronic granulomatous disease in Dobermann Pinschers, myeloperoxidase deficiency in Gray Collies, altered chemiluminescence in Weimaraners and Pelger-Huet anomaly (Weiss and Wardrop, 2011).

CLAD was initially described in a male Irish setter by Renshaw et al. (1975) and colleagues, they called the condition “Canine granulocitopathy syndrome”. The dogs had a clinical history of recurrent infections associated with pyrexia and neutrophilia and circulating leukocytes isolated from the dog at 5 months of age showed a bactericidal defect due to abnormal expression of the CD11b/CD18 complex (Bauer et al., 2004; Gu et al., 2004). The puppies described in this report showed severe skin lesions and septicemia, both phenomena can be attributed to a neutrophil hypofunction. Neutrophils are essential to preventing bacterial infection and dogs with naturally occurring septic and non-septic inflammatory diseases had an increased cell-expression of CD11b (Weiss and Wardrop, 2011). Hence, we might speculate that they were affected by a Canine granulocitopathy syndrome.

A single case of cabot ring cells has been reported in the Dachshund. This was characterized as a severe dyserythropoiesis in an 11-year-old female that presented with depression, diarrhea, weight loss and radiographic evidence of abdominal masses (Lukszewska and Lewandowski, 2008). Examination of the peripheral blood smear revealed erythroid dysplasia, anisocytosis with macrocytosis, Howell Jolly bodies, atypical nucleated cells, neutrophil hypersegmentation and the presence of giant forms. The final diagnosis was myelodysplasia secondary to metastatic adenocarcinoma (Lukszewska and Lewandowski, 2008).

A review on Pneumocystis carinii in the miniature Dachshund reported four cases with hyperpnoea, tachypnoea, exercise intolerance and signs suggestive of immune incompetence. It is probable that Pneumocystis carinii infection in these cases was the result of an immunodeficiency (Lobetti et al., 1996).

The hematological findings of both leukocytosis and severe leukocytic dysplasia is suggestive of an abnormal function of the bactericidal cells. Tests on neutrophil function were not performed in these cases because it was not possible to access the appropriate diagnostic tests in Italy. However, both the morphological cellular abnormalities and the clinical symptoms are suggestive of reduced cell function and consequently, a functional defect of the immune system. The death of other four puppies supports the hypothesis of a familial immunodeficiency.
3. CONCLUSION

This is the first report describing a familial immunodeficiency syndrome in Dachshund dogs characterized by significant hematological changes, particularly of the leukocyte cells.

4. ACKNOWLEDGEMENT

Researchers acknowledged to Dr E. Owen (University of Queensland, Australia) the both the scientific and English editing of the manuscript.

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