

**Gastroenterology****PERINUCLEAR ANTI-NEUTROPHILIC CYTOPLASMIC ANTIBODIES AND PROTEIN-LOSING ENTEROPATHY****Background**

Protein-losing enteropathy and protein-losing nephropathy are chronic and often fatal diseases in soft-coated wheaten terriers (SCWT). The diagnosis typically is made when dogs are approximately 4 to 6 years of age. The life expectancy is just a few months after diagnosis. An estimated 10% of SCWTs in the United States are believed to be affected.

Cardinal clinical findings include decreased serum albumin and globulin concentrations, an increase in urine protein:creatinine ratio (UP:C), increased fecal concentration of  $\alpha$ -proteinase inhibitor, and microalbuminuria. Advanced nephropathy is associated with increases in serum urea nitrogen and creatinine concentrations.

Renal biopsy specimens from SCWTs with protein-losing nephropathy have changes consistent with a sclerosing immune-complex glomerulonephritis. An immune-mediated process may also be involved in the pathogenesis of protein-losing enteropathy in SCWTs.

Circulating perinuclear anti-neutrophilic cytoplasmic autoantibodies (pANCAs) are associated with diet-responsive diseases in dogs of various breeds that have chronic enteropathies. Determining the presence of pANCAs has been used in humans as a reliable diagnostic test for subtypes of inflammatory bowel disease and autoimmune glomerulonephritis.

**Objectives**

To evaluate pANCA status in SCWTs and SCWT-beagle dogs and to correlate pANCA status of dogs with clinicopathologic variables of protein-losing enteropathy or protein-losing nephropathy, or both.

**Procedure**

Thirteen SCWTs and 8 SCWT-beagle dogs believed to have protein-losing diseases and a control group of 7 dogs with X-linked hereditary nephropathy and 12 healthy SCWTs more than 9 years old

were studied. Samples were obtained from affected dogs every 6 months. Collected samples included serum concentrations of albumin, globulin, creatinine, and urea nitrogen; fecal concentration of  $\alpha$ -proteainase inhibitor; and urinary protein-to-creatinine ratios. These parameters were then correlated with pANCA findings.

### **Results**

Twenty of the 21 dogs initially believed affected by protein-losing diseases had positive results for pANCAs at a minimum of 2 time points, and 18 of 21 dogs had definitive evidence of disease. None of the control dogs had positive results for pANCAs. A positive result for pANCAs was significantly associated with hypoalbuminemia but preceded the onset of hypoalbuminemia by an average of 2.4 years. Sensitivity and specificity of pANCAs to predict protein-losing enteropathy or nephropathy were 0.95 and 0.8, respectively.

### **Author Conclusion**

Most dogs affected with protein-losing enteropathy or nephropathy, or both, have positive results for pANCAs before clinicopathologic evidence of disease is detected.

### **Inclusions**

One figure, 14 references.

### **Editor Annotation**

Circulating pANCAs are associated with diet-responsive diseases in dogs of various breeds that have chronic enteropathies. The results of this study show that detection of pANCAs precede the onset of hypoalbuminemia by more than 2 years. The pANCA test may therefore be a useful early non-invasive test of disease in SCWTs. Whenever commercially available, pANCAs could help breeders gradually decrease the incidence of protein-losing nephropathy and enteropathy in SCWTs by removing these dogs from the breeding pool. (MM)

Allenspach K, Lomas B, Wieland B, et al. Evaluation of perinuclear anti-neutrophilic cytoplasmic autoantibodies as an early marker of protein-losing enteropathy and protein-losing nephropathy in Soft Coated Wheaten Terriers. *Am J Vet Res* 2008;69:1301-1304.