Making sense of cystine - Common Questions on Dogs Forming Cystine Stones

How will I know if my dog has cystinuria?
1. Cystine is an amino acid that is freely filtered in urine and almost completely reabsorbed by the kidney tubules (i.e. removed from the urine).
2. As urine cystine concentration increases, a positive urine nitroprusside test, cystine crystals and cystine stones are indicators of disease.
3. Although cystinuria and cystine crystalluria do not cause clinical signs, cystine stones can irritate the lining of the urinary tract resulting in urinary accidents, urgency, straining, or bloody urine. In some cases, the stones result in life-threatening urinary obstruction. Cystine stones are not always visible on x-rays and may require special contrast studies or an ultrasound to diagnosis.
4. Genetic tests for cystinuria are available for some breeds of dogs. A genetic marker test for androgen dependent cystinuria has been developed for Mastiffs, English bulldogs and French bulldogs. There are other breeds with androgen dependent cystinuria as well as other breeds in which castration will not reduce cystinuria (Type Ia, reported in Newfoundland dogs, Labradors and Landseers, Type IIa, reported in Australian Cattle dogs and Border Collies, Type IIb, reported in Miniature Pinschers).

Will castration prevent recurrence of cystine stones?
1. Surgical or medical castration can resolve/cure cystinuria in the subset of male dogs with androgen dependent cystinuria.
2. The magnitude of cystinuria associated with stone formation is wide (100 to >10,000 μmol/g creatinine) and varies between serial measurements in the same dog. This emphasizes that other factors such as diet, urine specific gravity and urine pH influence stone formation.
3. To determine if castration reduces cystinuria, measure urine cystine before and 2 months after castration. If the urine cystine is in the normal range, it indicates that castration was beneficial, and the risk for stone formation is likely eliminated. If the urine cystine remains elevated at 2 months, check again at 4 months. If the test is persistently positive, it indicates that the dog has a non-androgen dependent form of the disease. These cases are at risk for recurrence without additional therapy. Less sensitive indicators of urine cystine that can be evaluated before and after castration include urinalyses for cystine crystalluria, and recurrence of small stones.

Is a negative nitroprusside test following castration an indication that your dog will not reform stones?
1. Possibly, but not for certain.
2. The nitroprusside test is a qualitative screening test for disease (quantitative tests are better).
3. Some dogs with cystinuria are negative on this test even though they have the disease. If the nitroprusside test was only evaluated after castration, it is difficult to determine if castration reduced cystine excretion. In this situation, measure urine cystine concentration. In most stone-free dogs, cystine concentrations are extremely low.
4. Studies on human urine indicate that the nitroprusside test is influenced by urine specific gravity and creatinine, but not by thiola or D-penicillamine. Ampicillin and sulfur containing drugs have been reported to cause false positive results. This test may also be influenced by ascorbic acid, which dogs are able to synthesize within their body (unlike people). Therefore, the accuracy of this test in dogs should be interpreted with this in mind.
5. Feeding low-protein prevention foods has been associated with negative test results consistent with a treatment effect as opposed to a change in disease status.
**What diets prevent cystine stones?**

1. A 25% reduction in 24-hour urine cystine was associated with consumption of Hill’s Prescription diet u/d.
2. Avoid diets that promote formation of acidic and concentrated urine, which are risk factors for stone formation.
3. Diets rich in methionine (a precursor of cysteine and a common amino acid in animal protein sources) may contribute to cystine recurrence. Consider lower protein foods with reduced quantities of animal protein.
4. Studies in cystinuric humans suggest that dietary sodium enhances cystinuria, and therefore high sodium diets (>100-150 mg/Kcal) should be avoided.

View our Canine Cystine Recommendations

Osborne et al. Canine cystine urolithiasis: cause, detection, treatment….. VCNA. 1999;29:193