Brief Communication Communication brève

Pure cystine and urate calculi can be clearly visible using survey digital radiography

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Abstract — Cystine and urate calculi are considered nonradiopaque to faintly radiopaque. Two canine cases in which these types of calculi are radiopaque and clearly apparent *in vivo* on survey digital radiography are described. The densities of cystine and urate calculi, as determined *in vitro* with computed tomography, are compared to other pure calculi and mixed or compound calculi to further explore the relative attenuation characteristics.

Résumé — Les calculs de cystine et d'urate purs peuvent être clairement visibles à l'aide de la radiographie numérique standard. Les calculs de cystine et d'urate sont considérés comme non radio-opaques à faiblement radio-opaques. Deux cas canins dans lesquels ces types de calculs sont radio-opaques et clairement apparents *in vivo* sur la radiographie numérique standard sont décrits. Les densités de calculs de cystine et d'urate, telles que déterminées *in vitro* par tomodensitométrie, sont comparées à d'autres calculs purs et des calculs mixtes ou composés pour explorer davantage les caractéristiques d'atténuation relatives.

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rolithiasis is common in dogs (1). Cystine and urate urinary calculi are considered to be nonradiopaque to faintly radiopaque (2). The main goal of this brief communication is to report 2 cases of *in vivo* survey digital radiographic identification of cystine and urate urinary calculi in 2 canine patients. There are few peer-reviewed reports of cystine or urate calculi detected *in vivo* on survey radiography in dogs and cats, with even fewer available radiographic images (3–6). A secondary aim is to report computed tomographic attenuation values of cystine and urate calculi compared with other calculi to explore the role of density in the radiographic findings described *in vivo*.

The first case is a 2-year-old, intact male American pit bull terrier that was presented to the University of Minnesota 2 times over a period of 6 mo for stranguria. At each episode, multiple small calculi were identified within the urinary bladder on survey digital radiography (Vet Ray; Sedecal, Arlington Heights, Illinois, USA and Canon CXDI unspecified model, Melville, New York, USA) (Figure 1A). Urethral calculi were suspected, although not definitively identified due to soft tissue superimposition. Following retropulsion and stabilization, the calculi were subsequently removed *via* cystotomy and analyzed to be 100% cystine in both episodes.

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The second case is an 8-year-old, neutered male Dalmatian dog that was presented with vomiting, anorexia, and straining to urinate. An 11×6 mm calculus was clearly identified within the plane of the urethra on survey digital radiography (Vet Ray; Sedecal) (Figure 1B). Two smaller urethral calculi, radiographically faint urinary bladder calculi, and a nephrolith were also seen. Following retropulsion and stabilization, calculi were removed via cystotomy and analyzed to be 100% ammonium urate. All calculi analyses were conducted by the Minnesota Urolith Center (University of Minnesota, St. Paul, Minnesota, USA).

To further explore the relationship between radiographic opacity and material density, several groups of calculi were evaluated *in vitro* using computed tomography (CT). The calculi were provided by the Minnesota Urolith Center. A urolith without a nidus, shell, or surface crystal layer that contained $\geq 70\%$ of one type of mineral was identified by that mineral. A urolith with < 70% of one mineral but without a nidus, shell, or surface crystals was referred to as a mixed urolith. A urolith with an identifiable nidus and/or stone with ≥ 1 surrounding layer(s) of different mineral composition was called a compound urolith (7).

Uroliths were placed into individual plastic cups that had been partially filled with a clear gelatin substrate (Gelatine; Knox, Parsippany, New Jersey, USA), allowing the uroliths to rest approximately 5 mm above the bottom of the cups. Uroliths were then covered with 0.9% saline (8). Cups were arranged in a grid and then scanned using a helical CT scanner (Toshiba Aquilion 64 CFX CT; Toshiba, Tustin, California, USA) at 120 kV and 100 mAs and reconstructed in 0.5-mm slices using a soft tissue algorithm. Freeform regions of interest (ROI) were drawn on each sample calculus or the largest calculus in a sample

CVJ / VOL 62 / JUNE 2021 605

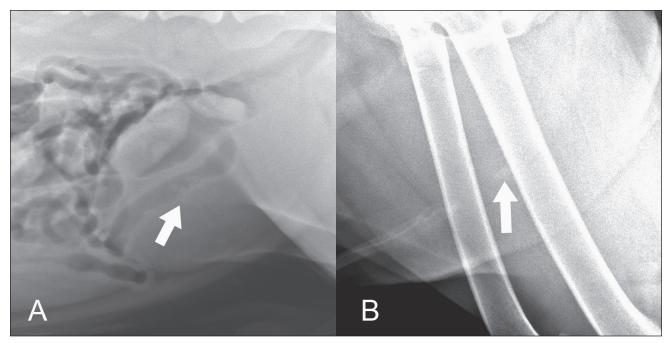


Figure 1. Survey radiographs of dogs with cystine and urate urolithiasis. A – Cystine urinary bladder calculi (arrow) (kVp 81, mAs 10.2). B – A urate urethral calculus (arrow) (kVp 92, mAs 3.2).

of multiple calculi to obtain average Hounsfield units (HU), evaluated with a window level of 25 and a window width of 350. The ROI were drawn just within the visible calculi borders to ensure that the surrounding saline was not inadvertently included in the measurements.

Computed tomographic density values (HU) were evaluated using commercially available statistical software (JMP Pro 13.2.1; SAS Institute, Cary, North Carolina, USA). Means and standard deviations of HU for all non-mixed, non-compound calculi types with at least 3 specimens were calculated. One-way analysis of variance (ANOVA) procedures were performed to compare HU values among the groups of calculi, and when the ANOVA demonstrated significant differences among calculi, differences between pairs of groups were assessed using the Tukey-Kramer HSD Experiment-wise test. Statistical significance for analyses was set at P < 0.05.

Thirty-five calculi samples were obtained from 34 dogs. Two of the cystine samples were obtained from the first case described in this report. Twenty-nine of the samples were analyzed to be 100% pure in composition, including calcium oxalate (n = 9), cystine (n = 6), urate (n = 3), struvite (n = 3), silica (n = 3), xanthine (n = 1), potassium magnesium phosphate (PMP, n = 1), and calcium phosphate carbonate (CPC, n = 1). An additional struvite sample had a nidus composed of 5% CPC and a brushite sample had a nidus of 15% calcium oxalate; these samples did not meet criteria for categorization of mixed or compound. The remaining 6 samples were mixed or compound in composition.

Hounsfield units ranged from 219 to 1848, depending on the calculus type (Table 1). Brushite and calcium oxalate were the highest attenuating calculi, similar to previous results (8,9). The HU ranges for struvite, cystine, and urate in this study were

Table 1. Hounsfield units determined for each type of calculus. For calculus types with multiple samples, results are reported as range (median).

Туре	Hounsfield units
Calcium oxalate	969 to 1496 (1141)
Struvite	487 to 643 (637)
Urate	431 to 539 (431)
Cystine	344 to 544 (427)
Silica	333 to 535 (408)
Brushite	1740
CPC	1123
PMP	908
Xanthine	219
Mixed/compound	332 to 1848

PMP — Potassium magnesium phosphate; CPC — Calcium phosphate carbonate.

similar to previous results (8,9). The HU obtained for the PMP, CPC, brushite, and xanthine samples were excluded from statistical analyses as only 1 sample for each of these calculi types was available. The HU obtained for silica, cystine, urate, and struvite were not statistically different from one another (P > 0.2 for all ordered differences). To the authors' knowledge, measured HU of silica calculi has not been reported in veterinary medicine. The HU range for calcium oxalate was statistically different from silica, cystine, urate, and struvite calculi (all P < 0.001). The range of HU for mixed and compound calculi was broad and overlapped with ranges obtained for the other calculi types. For reference; distilled water is defined as having a value of 0 HU, the reported mean HU of canine urine is 35.6 and the HU of non-contrast enhanced liver is approximately 50 to 70 (10,11).

This report provides 2 examples in which calculi types that are historically considered to be non-radiopaque were readily identified *in vivo* on survey digital radiography. When evaluating the densities of representative cystine and urate calculi

606 CVJ / VOL 62 / JUNE 2021

as determined with CT, there was no difference between the density of cystine and urate calculi and reportedly radiopaque calculi. This supports the ability to recognize cystine and urate calculi radiographically, as density is one of the main factors determining radiographic attenuation (12). A previous study of urinary calculi obtained from dogs demonstrated poor accuracy of detection of cystine and urate calculi in vitro when evaluated using screen-film radiography (13). It is speculated that the relatively radiopaque nature of the calculi of the 2 in vivo cases herein is due in part to the greater contrast resolution of digital radiography compared to screen-film systems, acknowledging that direct comparison between screen-film and digital radiography was not made in these 2 cases (14). The greater contrast resolution in digital radiography is related to the wider dynamic range associated with digital imaging receptors when compared to the silver halide crystal system of screen-film radiography. The effect of other variables such as calculus size is also acknowledged (13). Large-scale, ideally in vivo, studies evaluating the accuracy of survey digital radiography (potentially with comparison to screen-film radiography) in the diagnosis of various types and sizes of urolithiasis are warranted for further exploration of this subject, as in vivo prediction of urolith composition is used to guide case management (15).

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References

- 1. Hesse A. Canine urolithiasis: Epidemiology and analysis of urinary calculi. J Small Anim Pract 1990;31:599–604.
- Marolf A. Urinary bladder. In: Thrall D, ed. Textbook of Veterinary Diagnostic Radiology. 7th ed. Philadelphia, Pennsylvania: Saunders, 2017:851.

- 3. Grauer G. Cystine urolithiasis. Clinician's Brief 2014;12:71-73.
- 4. Butty E, Bua A, Vanstone N, Dunn M. Retained laser fiber in the nidus of a recurrent cystine urolith in an intact male English bulldog. Can Vet J 2019;60:29–32.
- 5. Grauer G. Ammonium urate urolithiasis. Clinician's Brief 2014;12: 51–55.
- Collins R, Birchard S, Chew D, Heuter KJ. Surgical treatment of urate calculi in Dalmatians: 38 cases (1980–1995). J Am Vet Med Assoc 1998;213:833–838.
- Osborne C, Clinton C, Bamman L, Moran H. Prevalence of canine uroliths: Minnesota Urolith Center. Vet Clin North Am Small Anim Pract 1986;16:27–44.
- Pressler BM, Mohammadian LA, Li E, et al. In vitro prediction of canine urolith mineral composition using computed tomographic mean beam attenuation measurements. Vet Radiol Ultrasound 2004;45:189–197.
- 9. Nykamp S. Dual-energy computed tomography of canine uroliths. Am J Vet Res 2017;18:1150–1155.
- Zwingenberger A, Carrade Holt D. Computed tomographic measurement of canine urine concentration. Can Vet J 2017;58:180–182.
- 11. Ohlerth S, Scharf G. Computed tomography in small animals Basic principles and state of the art applications. Vet J 2007;173:254–271.
- Curry T, Dowdey J, Murry R. Christensen's Physics of Diagnostic Radiology. 4th ed. Philadelphia, Pennsylvania: Lippincott Williams & Wilkins, 1990.
- 13. Weichselbaum R, Feeney D, Jessen C, Osborne C, Dreyster V, Holte J. Urocystolith detection: Comparison of survey, contrast radiographic, and ultrasonographic techniques in an in vitro bladder phantom. Vet Radiol Ultrasound 1999;40:386–400.
- Weatherburn G, Davies J. Comparison of film, hard copy computed radiography (CR) and soft copy picture archiving and communication (PACS) systems using a contrast detail test object. Br J Radiol 1999;72:856–863.
- Lulich JP, Berent AC, Adams LG, Westropp JL, Bartges JW, Osborne CA. ACVIM Small Animal Consensus Recommendations on the treatment and prevention of uroliths in dogs and cats. J Vet Intern Med 2016;30:1564–1574.

CVJ / VOL 62 / JUNE 2021 607