

# Cytology Challenge: Unusual Sediment Findings

Amy L. Weeden, DVM  
Heather L. Wamsley, DVM, PhD, DACVP (Clinical Pathology)  
*University of Florida*

**T**he checklist for identification of common urine sediment findings includes cells (eg, epithelial cells, erythrocytes, leukocytes), crystals (eg, struvite, calcium oxalate, bilirubin, ammonium biurate), and microorganisms (typically bacteria). Some practitioners, however, may encounter unusual findings that hold great diagnostic value. In the cases that follow, unusual urine sediment findings yielded interesting, if not diagnostically critical, information.



**CASE 1**

**Purulent Vaginal Discharge and Caudal Abdominal Pain**

A 5-year-old spayed beagle was presented with purulent vaginal discharge and caudal abdominal pain. Minimum database testing with urinalysis on a specimen obtained by cystocentesis showed an inflammatory leukogram, hyperglobulinemia, renal azotemia (elevated BUN and creatinine with concurrent isosthenuria), moderate proteinuria, and marked pyuria with gross hematuria. Cytologic examination of a urine sediment wet-mount specimen is shown (*Figure 1*).



▲ Unstained urine wet mount, 50× objective. Branching structures with parallel walls and perpendicular septa are present. *Courtesy of Diagnostic Cytology and Hematology of the Dog and Cat, Valenciano AC, Cowell RL (eds).*

**ASK YOURSELF**

- ▶ What are the branching structures?
- ▶ What additional diagnostic tests may be useful to confirm and better characterize this patient’s disease?

**DIAGNOSIS**

**Systemic Aspergillosis**

**Interpretation & Discussion**

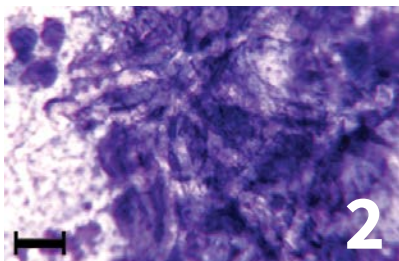
Ultrasonography showed cystic areas in the kidneys; fine-needle aspiration and cytology were pursued. In a Wright-Giemsa–stained specimen (*Figure 2*),

the branching hyphae with parallel cell walls were seen in the midst of dense nuclear debris and markedly karyolytic, degenerate neutrophils. Fungal hyphae showed variable staining cytologically. The hyphae may be nonstaining, as in this case, or they may stain bright blue with a thin, colorless cell wall.

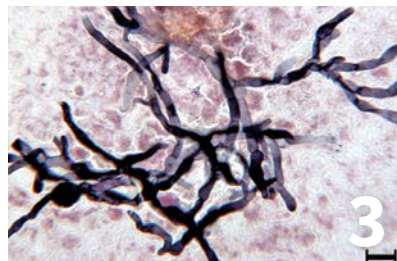
To better visualize the fungal hyphae, a

specialized silver stain was also used; the hyphae appear black with this special stain, and the septae are better visualized (*Figure 3*). This special stain can be performed on routine cytologic samples by reference laboratories.

Culture of urine showed *Aspergillus* species, a ubiquitous saprophytic fungus that may cause opportunistic infections. In 1 study, urinalysis findings in dogs with systemic aspergillosis included hematuria, pyuria, and fungal hyphae in the sediment, in decreasing order of frequency. Female dogs and German shepherd dogs appear to be overrepresented.<sup>1</sup>



▲ Wright-Giemsa–stained renal aspirate, 50× objective, bar = 10 μm. Large, non-staining, branched fungal hyphae are seen against a background of nuclear debris.



▲ Gomori methenamine silver staining of a renal aspirate, 50× objective, bar = 10 μm. The branching, septate fungal hyphae are black in the silver-stained preparation.

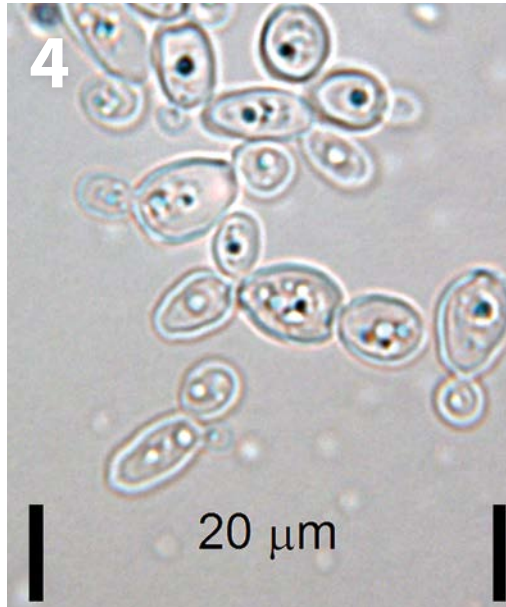
**DID YOU ANSWER?**

- ▶ Fungal hyphae. In this case, the diagnosis was *Aspergillus* spp infection.
- ▶ Fungal culture or serologic testing.

## CASE 2

### Clinically Normal Patient with Chronic Antibiotic Administration

An adult spayed female domestic shorthaired cat was undergoing routine annual examination with laboratory evaluation. The patient had received a potentiated  $\beta$ -lactam at the labeled dose intermittently for chronic upper respiratory infection and had recently finished a 7-day course. A photomicrograph of a wet-mount preparation of urine sediment obtained on cystocentesis is shown in **Figure 4**.



▲ Unstained urine wet mount, 100 $\times$  objective. A few clusters of oval organisms are shown.

## ASK YOURSELF

- ▶ What are the oval structures?
- ▶ What part of this patient's history likely contributed to this urine sediment finding?

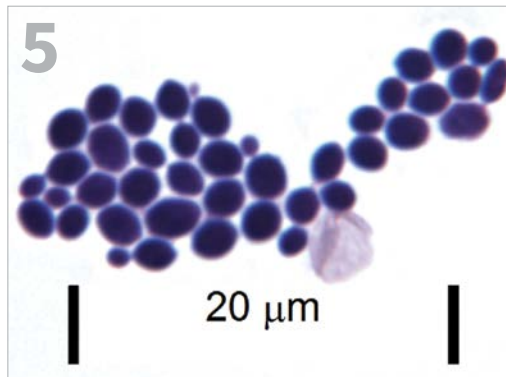
## DIAGNOSIS

### Urinary Candidiasis, Likely Secondary to Chronic Antibiotic Administration

#### Interpretation & Discussion

The patient's urine contained large numbers of yeast cells, some of which were budding (**Figure 5**); no other evidence of cystitis was present. This is an uncommon finding. False diagnosis of yeast cystitis is possible if the yeast identified microscopically is actually from wet mount stain that is contaminated by yeast or from the patient's genitals or skin. These potential sources of contamination are avoided by examination of a fresh urine sample obtained by cystocentesis that is prepared as a plain, unstained wet mount.

Yeast cystitis is typically a secondary pathology. Considerations for primary



▲ Wright-Giemsa-stained urine dry mount, 100 $\times$  objective. A cluster of 2- to 3- $\mu$ m yeast (sometimes budding) is shown along with an erythrocyte near the center of the field.

disease include bacterial cystitis, uroliths, and urinary catheterization, as well as chronic antibiotic use, diabetes mellitus, and other immune-compromising disorders.<sup>2</sup>

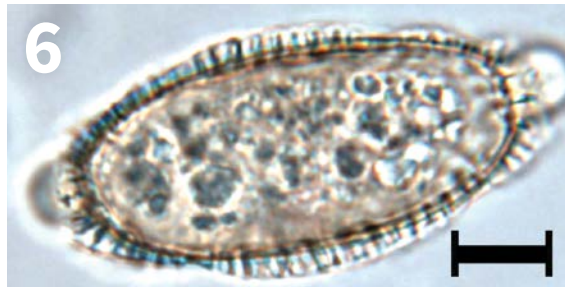
## DID YOU ANSWER?

- ▶ Budding yeast, identified as *Candida* spp on culture.
- ▶ The patient's chronic antibiotic administration likely contributed to the yeast infection.

**CASE 3**

**Partial Urethral Obstruction**

A 12-year-old indoor-outdoor castrated domestic shorthair cat was presented with stranguria, dysuria, and pollakiuria. Urinalysis with sediment evaluation was performed in-house. Based on the findings, contamination of the urine sample with fecal material was suspected, so the sediment was submitted to a reference laboratory for review (**Figure 6**).



▲ Unstimulated urine wet mount, 50× objective, bar = 10 μm. A large, oval structure is present.

**ASK YOURSELF**

- ▶ What is the structure?
- ▶ Which species may be affected with this condition?
- ▶ What is the most common presenting complaint associated with this condition?
- ▶ Why was the referring veterinarian concerned about fecal contamination of the urine sample?

**DIAGNOSIS**

**Capillariasis**

**Interpretation & Discussion**

*Capillaria plica* (*Pearsonema plica*) is a nematode found in the urinary bladder of cats, dogs, foxes, and other wild canids. The eggs may be confused with the eggs of *Trichuris vulpis* because of the bipolar opercula. However, *T vulpis* is an intestinal parasite. The opercula of *C plica* eggs are usually slightly askew, whereas those of *T vulpis* tend to be perfectly bipolar.

The shells of *C plica* eggs have a rougher, mammillated surface (**Figure 7**), whereas those of *T vulpis* are smooth. Ingestion of the earthworm intermediate host containing infective larvae is the likely mode of transmission.

Although most patients are clinically normal, lower urinary tract signs have



▲ Unstimulated wet mount, 50× objective, bar = 10 μm. A *Capillaria plica* ovum is presented to better show the mammillated shell and opercula, which are slightly askew from 180°.

**DID YOU ANSWER?**

- ▶ *Capillaria plica* ovum.
- ▶ Cats, dogs, and foxes.
- ▶ Patients are usually clinically normal.
- ▶ *Capillaria* ova resemble those of *Trichuris vulpis*, or whipworm.

been observed in dogs, while inflammation and urethral obstruction have been observed in cats.<sup>3,4</sup>

#### CASE 4

### Weight Loss and Chronic Cough

A previously stray, middle-aged, crossbreed dog was presented with inability to gain weight and chronic intermittent cough. Initial minimum database evaluation for respiratory and metabolic disease, including urinalysis on a sample obtained by cystocentesis, was performed. A photomicrograph of a wet-mount preparation of urine sediment is shown in *Figure 8*.



▲ Unstained wet mount, 40× objective. A slender larva with a tapered tail is shown. Hematuria, which was likely iatrogenic, is also observed. Squamous cells are seen at the lower left and right.

#### ASK YOURSELF

- ▶ Could the larvae be related to the patient's respiratory signs?

#### DIAGNOSIS

### *Dirofilaria immitis* Infection

#### Interpretation & Discussion

This was an unusual finding. Larvae are not usually seen in urine. Primary urinary system nematodes in small animals include *Capillaria* species and *Dioctophyme renale*, both of which may be diagnosed by finding representative ova in the urine.

Larvae of *Capillaria* species have rarely been reported in feline urine sediment,<sup>4</sup> and those of *D renale* are not usually seen. No ova were found in the urine sediment. However, this patient had a

patent *Dirofilaria immitis* infection diagnosed by occult heartworm ELISA. Microfilariae were also found on blood smear examination. The urine sediment microfilariae were likely contaminants caused by iatrogenic hemorrhage during collection.

#### DID YOU ANSWER?

- ▶ Yes. The larva was a contaminant from the iatrogenic hemorrhage in the sample and was consistent with this patient's diagnosis of *D immitis* infestation.

**Primary urinary system nematodes in small animals include *Capillaria* species and *Dioctophyme renale*, both of which may be diagnosed by finding representative ova in the urine.**

**CASE 5**

**Urethral Obstruction**

A 5-year-old intact male Australian cattle dog was presented with stranguria with minimal-to-no urine production. Physical examination showed pain on palpation of the caudal abdomen and distention of the bladder.

Imaging disclosed urinary bladder and penile urethral calculi just proximal to the os penis. Urinalysis was performed and crystals were observed in a wet-mount preparation of the urine sediment (*Figure 9*).



▲ Unstained urine wet mount, 50× objective. Flat, hexagonal, and sometimes irregular-shaped, colorless crystals are shown. They frequently overlay one another. Smaller daughter crystals are sometimes seen budding from a larger crystal.

**ASK YOURSELF**

- ▶ What type of crystal is present?
- ▶ What usually causes the formation of these crystals?
- ▶ Under what pH conditions are these crystals formed?

**DIAGNOSIS**

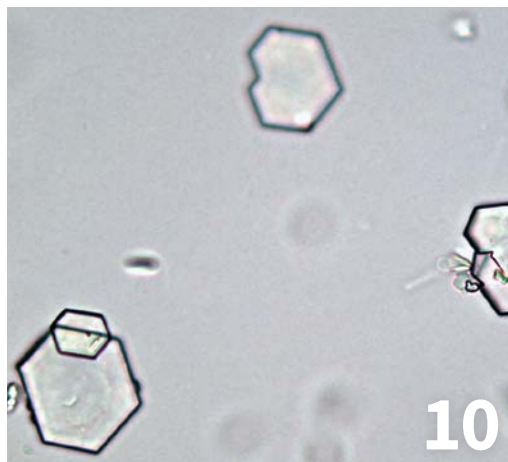
**Cystine Urolithiasis**

**Interpretation & Discussion**

Urolith analysis was consistent with the urinalysis findings of cystinuria. Cystine crystalluria and urolithiasis is an uncommon but well-documented condition in dogs. Recent studies have looked at the molecular basis and modes of inheritance of different types of canine cystinuria.<sup>5</sup> It is caused by an inherited renal tubular transport defect in which affected individuals cannot reabsorb dibasic amino acids.

Although other amino acids (eg, ornithine, lysine, arginine) are also lost in the urine, cystine is the most clinically relevant as its low solubility in acidic or neutral urine leads to the formation of crystals and uroliths along the urinary tract.<sup>6</sup>

Many breeds, including English bulldogs, dachshunds, Newfoundlands, Australian cattle dogs, Labrador retrievers, other



▲ Unstained urine wet mount, 50× objective. The cystine crystals are seen superimposed on each other with smaller crystals budding from larger ones.

purebreds, and crossbreeds, have been affected. Cystine crystalluria is a risk factor for urolithiasis and should prompt ultrasonographic evaluation for uroliths.

**DID YOU ANSWER?**

- ▶ These are cystine crystals (*Figure 10*), which are flat, hexagonal, and colorless and tend to form stacks.
- ▶ An inherited renal tubular transport defect is usually the cause.
- ▶ Acidic or neutral urine favors formation of cystine crystals.

## CASE 6

### Refractory Seizures

A 15-year-old castrated Jack Russell terrier was presented with seizures that were refractory to medical management. During hospitalization, medications included zonisamide, levetiracetam, dexamethasone, acetylcysteine, furosemide, and mannitol. As part of the diagnostic evaluation of neurologic disease, a urinalysis was performed. Photomicrographs of the urine sediment are shown in **Figure 11**.



▼ Unstained urine wet mount, 50× objective, bar = 10 μm. Photomicrographs showing crystalline structures that were numerous in the urine sediment. The crystals are dark brown, variably shaped, and exhibit a concentric layering effect. *Courtesy of Diagnostic Cytology and Hematology of the Dog and Cat, Valenciano AC, Cowell RL (eds).*

## DIAGNOSIS

### Intracranial Mass

#### Interpretation & Discussion

The patient's refractory seizures were associated with an intracranial mass. The polypharmacy used to treat this patient was likely the cause of the crystals. A number of drugs have been associated with crystalluria in human and veterinary patients. Anticonvulsant polypharmacy has specifically been studied in the human literature. Patients

receiving zonisamide, a sulfa-containing medication, and patients taking multiple medications frequently had crystalluria.<sup>7</sup>

## DID YOU ANSWER?

- ▶ No, the crystals are unusual.
- ▶ The crystals are likely formed from precipitated drugs or drug metabolites, given the patient was on multiagent therapy that included anticonvulsant drugs.

## ASK YOURSELF

- ▶ Are the crystalline structures of a common morphology seen in dogs?
- ▶ What is the likely composition of the crystal structures?

## Conclusion

Although the urine sediment findings in this series are not common, they did prove useful in the diagnosis or confirmation of many of the disease processes presented. The series highlights the diagnostic utility of urine sediment evaluation beyond the typical checklist used in most practices and the importance of pursuing unexpected or unusual laboratory findings. ■

## References

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