Chronic Kidney Disease & the Role of Phosphorus Binders
Chronic kidney disease (CKD) is one of the leading causes of death in dogs and cats. In North America it is estimated that 8% of cats aged 10 years or older and approximately 2.4% of dogs aged 10 years or older suffer from CKD.1

Renal failure results when more than three quarters of nephrons in both kidneys are not functioning. Nephron damage associated with CKD is usually irreversible and the cause is often difficult to determine. Because of the interdependence of the vascular and tubular components of the nephron, the end point of irreversible glomerular or tubular damage is the same. The histologic changes are not process-specific, frequently making an etiologic diagnosis impossible. Progressive disease that destroys nephrons at a slow rate allows intact nephrons to undergo compensatory hypertrophy; but when renal failure finally occurs, hypertrophic nephrons can no longer maintain adequate renal function.

**DIAGNOSIS OF CKD**

A diagnosis of renal failure is confirmed when persistent azotemia with concurrent isosthenuria or minimally concentrated urine is documented. Early stages of kidney disease may be difficult to confirm because there are usually no overt clinical signs. Sequential monitoring of blood pressure, serum creatinine, urine specific gravity, and proteinuria may aid in the recognition of early kidney disease. Serum creatinine levels are commonly used to measure kidney dysfunction and are the basis of the IRIS staging system. Serum creatinine concentrations should always be interpreted in light of urine specific gravity. For example, azotemia concurrent with hypersthenuria (unusually high specific gravity and solute concentration in urine) suggests decreased renal perfusion rather than renal disease. Serial evaluation of serum creatinine concentrations on an annual or semi-annual basis may allow clinicians to detect renal disease prior to the onset of persistent azotemia. For example, an increase in serum creatinine concentration from 0.6 to 1.2 mg/dl over...
time can indicate a 50% or greater reduction in glomerular filtration rate even though a value of 1.2 mg/dl is within the normal range.

Vomiting and anorexia are also common in dogs and cats with CKD. Causes include stimulation of chemoreceptors by uremic toxins, decreased gastrin excretion and uremic vasculitis, increased gastric acid secretion, and gastrointestinal irritation secondary to uremic toxicity.

**ROLE OF PHOSPHORUS IN CKD**

Plasma phosphorus concentrations increase in CKD because of decreased renal excretion, causing hyperphosphatemia. At the same time, there is a decrease in the active form of vitamin D, that reduces intestinal absorption of calcium. This process, combined with impaired ability to reabsorb calcium in the kidney, causes ionized calcium concentrations to decrease. In response to decreased vitamin D and plasma calcium concentrations, parathyroid hormone (PTH) levels increase. This functional hyperparathyroidism...
can cause osteodystrophy, neuropathy, bone marrow suppression, and soft tissue mineralization. It has been shown that by adding enteric phosphorus binders and reducing phosphorus in the diet, hyperphosphatemia can be reduced.

**MANAGEMENT**

Even though CKD is usually irreversible from a histologic and functional viewpoint, with proper treatment, the severity of clinical signs can generally be reduced. Longevity may even be increased by decreasing phosphatemia.\(^2\) It is important that water be available ad libitum so that dehydration does not occur, that fluids be replaced parenterally if necessary, and that any infection be managed.

Therapy to slow progression of disease should be started as soon as diagnosis is conclusive. Dietary sodium levels can remain normal or if hypertension is present sodium can be gradually decreased. Dietary protein should be reduced but should never go below the patient’s protein requirement.

**Epakitin Highlights**

- Binds phosphorus
- Reduces uremia
- Improves general health of animals with CKD
- Demonstrated clinical efficacy
- Highly palatable

**Therapeutic Approaches**

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<td>Disease-specific therapy (renoliths, pyelonephritis)</td>
<td>Renoprotective therapy to slow progression (ACE-inhibitors, diet)</td>
<td>Symptomatic patient-specific therapy (anorexia, vomiting, acidosis)</td>
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**Management of CKD**

**Dogs & Cats**

- Fresh water ad libitum
- Disease-specific therapy
  - Control infection
  - Remove urolith obstructions
- Therapy to control progression
  - Sodium levels normalized or gradually decreased
  - Protein levels reduced but within protein requirements
  - Canine 2.0 to 2.2 g protein/kg/day
  - Feline 3.3 to 3.5 g protein/kg/day
  - Dietary phosphorus levels reduced
  - Phosphorus binder (Epakitin) added to diet
  - ACE inhibitors & calcium channel blockers if needed
- Patient-specific therapy
  - Control uremia
  - Control acidosis
  - Control vomiting

**Management Challenges**

- **Calorie malnutrition** may be caused by vomiting and anorexia. Controlling uremia may improve appetite and remove some of the causes of vomiting. Palatable enteric phosphorus binders will also help increase caloric intake.
- **Hypokalemia & potassium depletion** may occur in cats. Anorexia, high protein diets or acidifying diets, polyuria/polydipsia, and vomiting all contribute to potassium depletion. Oral potassium should begin early.
- **Pyelonephritis & renoliths** may occur in cats. Urinary tract infections are rare in healthy cats but may increase in older cats, especially those with CKD. Calcium oxalate uroliths are the most common type of renoliths in older cats.
- **Hyperthyroidism** is a common endocrinopathy in cats. Many hyperthyroid cats are more than 10 years old and often have concurrent renal insufficiency or CKD. Coexistence of these conditions can complicate diagnosis and treatment.

**REFERENCES**