Anesthesia for Patients with Renal Disease

Marlis Rezende, DVM, PhD, DACVAA Khursheed Mama, DVM, DACVAA Colorado State University

You have asked... What should I consider when anesthetizing patients with renal disease?

The experts say...

Renal disease varies in form and severity. Patients may present with a wide range of conditions, from asymptomatic status to severely compromised. Therefore, anesthetic management strategies must be individualized according to specific patient needs.

Renal disease likely goes undiagnosed at its earliest stages because blood urea nitrogen (BUN) and creatinine elevations only occur when renal function is reduced to 50%–70% of normal.¹At the other extreme, animals suffering from uremic toxemia may have CNS and myocardial depression, pericarditis, pneumonitis, coagulopathies, and increased susceptibility to infection and sepsis. Animals with significant renal disease lose the ability to adjust urine volume and content and may present with anemia, systemic hypertension, metabolic acidosis, and hyperkalemia. Chronic kidney disease is often observed as a comorbidity with other diseases, such as in dogs with heart failure caused by mitral valve insufficiency or in cats with hyperthyroidism (which may mask the severity of renal disease).

Because of the large variability in type and severity of renal disease, a complete physical examination and laboratory testing (eg, CBC, serum chemistry panel, urinalysis) are used to better define the patient's condition and determine the best perianesthetic management plan. Preanesthetic stabilization is essential to a successful outcome. For information regarding clinical management of animals presenting with complex or multifactorial disease, refer to comprehensive anesthesia textbooks or consult with a board-certified anesthesiologist (see **ACVAA Resources**).

Perianesthetic Kidney Support

The kidneys are particularly susceptible to ischemic injury because the distribution of blood flow within the kidneys is not uniform or proportional to demand; despite its higher metabolic rate, for example, the renal medulla only gets 15% of the overall renal blood flow (RBF).^{1,2} Therefore, a decrease in cardiac output and blood pressure— as may be observed with anesthetic drug administration—has the potential to further compromise renal function.

To ensure adequate oxygen delivery to the kidneys, anesthesia should be focused on maintaining circulation and oxygen-carrying capacity. Hypovolemia, hypotension,



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ACVAA Resources

The American College of Veterinary Anesthesia and Analgesia (ACVAA) is a useful resource for information regarding clinical management of animals presenting with complex or multifactorial renal disease as well as consultation with a local board-certified anesthesiologist (acvaa.org/locator). dehydration, hypoproteinemia (low colloid oncotic pressure), and acid-base and electrolyte abnormalities should thus be corrected before anesthesia is administered. Packed red cell transfusions should be considered in anemic patients. Recommendations for an exact value (for PCV) vary, as the need depends on other factors such as chronicity of disease, cardiovascular status of the patient, concurrent disease, and others. It is broadly recommended that transfusion is indicated when the PCV is 20%–23%.

Elective procedures should be delayed until patients can be stabilized adequately. While emergency procedures may not be postponed, time should be taken to improve the patient's condition as much as possible before the procedure, with special attention to the most lifethreatening abnormalities (low circulating volume, hyperkalemia, metabolic acidosis, severe anemia). Supplemental oxygen during the perianesthetic period (ie, face mask, induction box, oxygen case) may help prevent hypoxemia and hemoglobin desaturation and improve oxygen content in severely anemic patients.

Systemic hypotension has a direct effect on renal perfusion because autoregulation is compromised when mean arterial pressure decreases below 80 mm Hg and the kidneys can no longer control their own blood flow, which then becomes directly proportional to mean arterial pressure. In addition to decreasing anesthesia depth (when possible, or using a balanced anesthetic technique) and maintaining circulating blood volume via appropriate IV fluid administration,

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inotropic drugs (dopamine 5 μ g/kg/min or dobutamine 2–5 μ g/kg/min) may be used to support myocardial function and, as a consequence, increase cardiac output, blood pressure, and RBF. Vasopressors are not typically advocated because excessive vasoconstriction can negatively affect RBF despite normal or high blood pressure.

Induction Protocols

While most anesthetic drugs do not negatively affect the kidneys directly, general anesthesia can significantly decrease cardiac output and hence blood flow to the kidneys, so the anesthetic protocol should favor drugs that best preserve cardiovascular function and minimize renal vasoconstriction. Opioids (eg, hydromorphone or oxymorphone 0.05-0.1 mg/kg SC or IM) may be used to provide sedation (euphoria in cats is possible but unlikely at the lower end of the dose range) and analgesia and facilitate a reduction in the dose of anesthetic induction and maintenance agents required. If warranted, an anticholinergic (eg, atropine 0.02-0.04 mg/kg SC) may be used to prevent opioid-induced bradycardia. Because α_2 -agonists (eg, dexmedetomidine, medetomidine, xylazine) cause significant vasoconstriction and decrease cardiac output, they should be avoided in these patients.

Anesthesia induction in severely compromised dogs is performed safely with a combination of an opioid and a benzodiazepine (eg, fentanyl 10 μ g/kg IV or hydromorphone 0.1 mg/kg IV and midazolam 0.25 mg/kg IV). Supplemental administration of an anticholinergic (eg, atropine 0.01–0.02 mg/kg IV) might be needed to maintain heart rate. If additional anesthetic drug is needed to perform intubation, a small dose of ketamine or propofol (1–2 mg/kg IV titrated to effect) may be used. For severely affected cats, a lower opioid dose (eg, fentanyl $3-5 \ \mu g/kg IV$) may be used with a benzodiazepine and an adjunct drug such as ketamine (2–3 mg/kg IV). Alternatively, etomidate (0.5–2.0 mg/kg IV), which has a wide margin of cardiovascular safety, may be administered with a benzodiazepine. Dilution of this highly osmolar drug is recommended to prevent hemolysis.

Induction with ketamine and midazolam (5-7 and 0.3 mg/kg IV, respectively) is appropriate for stable dogs and cats with mild renal disease. Cats are unable to completely metabolize ketamine and eliminate the active drug through the urine. Hence prolonged effects may be noted in oliguric patients. If oliguria or anuria is post-renal, ketamine can be used provided the urethral obstruction can be corrected readily. Propofol may be used in minimally affected dogs or cats; because it has the potential to cause vasodilation and hypotension, however, its use should be avoided in debilitated patients (particularly those with fluid deficits). Combinations of propofol with opioids, benzodiazepines, and/or ketamine help reduce the dose of propofol and are thought to decrease these negative effects.

Adjunct Medications During Anesthesia

While vascular volume and blood pressure support remain the most important aspects of protecting renal function during anesthesia management, adjunct medications are often advocated. Three of these are discussed:

- Mannitol
- Dopamine
- Fenoldopam

Mannitol is an osmotic diuretic that has been used in the perioperative period to increase circulating volume, RBF, and urine output; decrease endothelial swelling; and scavenge free radicals. It may be



Maintenance, Support, & Monitoring

Anesthesia is maintained with an inhaled anesthetic agent and supplemented with adjunct cardiovascular-sparing drugs (balanced anesthetic technique) such as opioids, which may be administered by infusion or intermittent bolus. Because of the known nephrotoxic metabolites (eg, compound A) produced by sevoflurane, it is best to avoid this drug even though any toxic effect may be minimal in a stable patient with renal disease. Therefore, the use of isoflurane is generally recommended.

Support and monitoring should be consistent with the standard of care and modulated further according to disease severity and potential comorbidities as well as expected duration of anesthesia. Typically, this includes evaluation of blood pressure (direct arterial blood pressure monitoring is recommended in severe cases), heart rate and rhythm, oxygenation, ventilation, and body temperature. In compromised patients, additional monitoring of blood gases, electrolytes, acid–base status, urine output, and PCV/TP helps to facilitate timely intervention.

Appropriate IV fluid therapy should be instituted to maintain oxygen delivery, with inotropic support added as needed to support cardiac function. Isotonic crystalloid fluids (eg, lactated Ringer's solution, Plasma-Lyte) are preferred and typically used at 5–10 mL/kg/h during anesthesia (except for anuric patients, as they are at a high risk for fluid overload). Synthetic colloid use should be minimized in these patients, as there is growing evidence of potential negative renal effects in humans (at this time, no data are available to indicate whether the same is true for animals).^{3,4} Ventilatory support may be required when high doses of opioids are used during anesthesia. Supplemental oxygen should be provided before induction and during the recovery period. Hypothermia should be prevented with active warming devices.

Table. Anesthetic Protocol Samples

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	Premedication	Induction
Stable dogs and cats (mild disease)	Opioid (hydromorphone or oxymorphone 0.05 -0.1 mg/kg SC or IM) \pm anticholinergic (atropine 0.02 -0.04 mg/kg SC or IM)	Ketamine (5–7 mg/kg IV) and midazolam (0.3 mg/kg IV); or propofol (2 mg/kg IV) and ketamine (2–3 mg/kg IV)
Severely compromised dogs	Opioid (hydromorphone or oxymorphone $0.05-0.1$ mg/kg SC or IM) \pm anticholinergic (atropine $0.02-0.04$ mg/kg SC or IM)	Fentanyl (10 μ g/kg IV) and midazolam (0.25 mg/kg IV) \pm atropine 0.01–0.02 mg/kg IV \pm propofol or ketamine (1–2 mg/kg IV, titrated to effect)
Severely compromised cats	Opioid (hydromorphone or oxymorphone 0.05 mg/ kg SC or IM) \pm anticholinergic (atropine 0.02 mg/kg SC or IM)	Fentanyl (3–5 $\mu g/kg$ IV), midazolam (0.25 mg/kg IV), and ketamine (2–3 mg/kg IV); or, etomidate (0.5–2 mg/kg) and midazolam (0.25 mg/kg IV)
Table is not all inclusive. Refer to text for additional information on anesthetic protocols.		

continues

Anesthesia for patients with renal disease should focus on maintaining optimal renal perfusion and oxygenation.

administered as a single dose but is often administered during anesthesia as an infusion at 0.1 g/kg/h. To prevent dehydration, care must be taken to ensure that the patient is well hydrated and normovolemic prior to administration and that fluid therapy is provided during anesthesia and the postoperative period. Data have demonstrated clinical benefit from perioperative use of mannitol in patients undergoing renal transplantation. Prophylactic use of mannitol remains common in human patients (and veterinary patients in some practices) undergoing surgery with a high risk for acute renal injury; aortic or vena cava cross-clamping would be classic examples, but prophylactic mannitol may be used in any surgical procedure with potential for severe hypotension. The authors also use mannitol infusions for patients with elevated BUN and creatinine undergoing less invasive or elective procedures, albeit studies in humans have not been able to show improvement in outcome with its use.5

Dopamine has dose-dependent agonist effects at dopaminergic, β -, and α receptors. At low doses (<3 μ g/kg/min), dopamine is a nonselective dopaminergic agonist (DA-1 and DA-2) and can increase RBF and glomerular filtration rate, induce sodium diuresis, and decrease tubular oxygen consumption in species with renal dopamine receptors (humans, dogs, rabbits, and rats but not cats). While the beneficial effects of lowdose dopamine on renal function remain controversial,⁶⁻⁸ inotropic doses (5 µg/kg/ min) are considered beneficial because they increase blood pressure, cardiac output, and RBF. Dopamine doses of 10 µg/ kg/min or higher are generally avoided because of vasoconstriction, which may limit RBF. The authors routinely use dopamine (titrated to effect) in patients with renal disease to maintain cardiac output and blood pressure.

Fenoldopam is a selective DA-1 receptor agonist that has been shown to increase RBF and urine output and cause natriuresis in both humans and dogs.⁹⁻¹¹ In dogs, it is typically administered at 0.1-0.2 µg/ kg/min¹⁰ because higher doses are associated with hypotension during anesthesia. Hypotension is a potential side effect of fenoldopam as a result of its peripheral vasodilatory properties. However, glomerular filtration rate and RBF seem to be maintained even in the face of decreased blood pressure. Studies in humans undergoing cardiovascular surgery have offered promising results, indicating a clinical benefit and improved outcome with the use of fenoldopam,¹² but more studies are needed to confirm these findings. Nevertheless, its current high cost might limit fenoldopam use in veterinary practice.

In Sum

Anesthesia for patients with renal disease should focus on maintaining optimal renal perfusion and oxygenation. Therefore, dehydration, low circulating volume, hypotension, anemia, and oxygen desaturation should be avoided during anesthesia. A thorough evaluation to assess patient condition and anesthetic risk, followed by adequate stabilization prior to anesthesia, are key to preventing further renal damage. **Cb**

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BUN = blood urea nitrogen, DA = dopaminergic agonist, RBF = renal blood flow