EffectivenessMatters

December 2015

Acute Kidney Injury: the 5**R**s approach



This issue of *Effectiveness Matters* has been produced by CRD in collaboration with the Yorkshire and Humber AHSN Improvement Academy. The views expressed in this bulletin are those of the authors and not necessarily those of the AHSN or the University of York.

THE UNIVERSITY of York
Centre for Reviews and Dissemination

- The 5Rs approach to managing AKI includes: Risk, Recognition, Response, Renal support and Rehabilitation
- Patients at Risk of AKI should avoid episodes of dehydration and nephrotoxins
- Recognition of AKI is dependent upon good clinical judgement, careful monitoring of urine output and measurement of serum creatinine
- Rapid Response to AKI includes screening for sepsis, avoidance of toxins, optimisation of blood pressure and preventing harm (STOP AKI)
- Patients with severe AKI should be referred and receive Renal support
- Patients with or at risk of AKI, and their carers, should receive appropriate Rehabilitation including details about risk factors for AKI, preventative measures, treatment options and possible outcomes



Background

Acute kidney injury (AKI), previously known as acute renal failure, is a rapid reduction in kidney function over hours or days. The definition is based on rises in creatinine or reductions in urine output. AKI is associated with up to 100,000 deaths each year in UK hospitals.¹

A recent meta-analysis of the global burden of AKI found that 1 in 5 adults and 1 in 3 children worldwide experience AKI during a hospital stay. The pooled AKI-associated all-cause mortality rate was 22.4% (95% CI, 20.5 to 24.4) and increased with severity. The onset of AKI is insidious and often goes unrecognised, allowing deterioration that can result in uraemia, acidaemia, hyperkalaemia and ultimately death. AKI is a risk factor for chronic kidney disease and end-stage renal disease. However, with the correct care and treatment, around 30% of cases of AKI could be avoided.

There is an urgent need to raise awareness and improve the care of people at risk of, or with AKI.⁵ This edition of *Effectiveness Matters* summarises the national guidance and introduces the conceptual approach of the 5Rs to managing AKI.⁶ The evidence relates to adults and children from 1 month to 18 years of age.

Risk of AKI

Acutely ill patients or patients undergoing major surgery are at particular risk especially in the presence of additional risk factors (Table 1).⁷ It is most commonly secondary to hypotension and sepsis and more rarely due to primary kidney disease such as vasculitis or acute interstitial nephritis.^{1,8} It is rarely caused by direct trauma to the kidneys.

There is no universally accepted validated risk score for AKI for either primary or secondary care.^{7,8} Identifying relevant risk factors and undertaking appropriate blood biochemistry and urine output monitoring are key to recognising those at risk of, or with AKI in order to prevent onset or deterioration.⁷⁻⁹ This should include ensuring a baseline serum creatinine measurement for all acutely ill patients on admission.¹⁰

If an acutely ill patient has or is likely to have any of the established risk factors listed in Table 1, they should be investigated for AKI by clinical examination, urine output measurement and checking serum creatinine levels with a baseline measurement.⁷

Iodinated contrast agents used in imaging can result in or exacerbate AKI; specific risk factors should be assessed, but should not delay emergency imaging.⁷ Further information about those at risk of developing AKI is available in the NICE guidance and through the NHS England 'think Kidneys' campaign.^{7,8}

Recognising AKI

AKI can be easily recognised clinically in some cases by a low urine output (oliguria), or no urine output (anuria) but can occur without symptoms. A

comparison of serum creatinine levels with baseline will identify the presence and/or severity of AKI.^{1,9}

A national algorithm which standardises the definition of AKI and facilitates automated detection based on changes in serum creatinine levels has been introduced. A Patient Safety Alert directive requires integration of the algorithm into laboratory information management systems that communicate with patient management systems. ¹¹

A study looking specifically at the NHS England AKI algorithm found it performed well as a diagnostic adjunct in clinical practice. However, without baseline data AKI may only be identified in retrospect, so active clinical care remains essential.

There is some indication from reviews that in general early warning systems when combined with rapid response may have the potential to reduce cardiac arrests and unplanned ICU admissions. A more recent prospective observational study found that an interruptive alert as part of an electronic recognition system for AKI significantly increased the completion rate of the AKI care bundle.

New biomarkers such as neutrophil gelatinase-associated lipocalin (NGAL), kidney injury molecule-1 (KIM-1), N-acetyl- β -d-glucosaminidase (NAG), and Interleukin-18 (IL-18), for the early detection of AKI have been the subject of recent reviews. ¹⁶⁻¹⁸ Heterogeneity in populations, settings, definitions of AKI and levels of serum creatinine used for diagnosis, are consistent confounders in the evidence base. The reviews all conclude that while promising, the value of biomarkers in the early detection of AKI has yet to be demonstrated.

Response

Patients at risk

NICE guidance for recognition of and response to acute illness in adults in hospital recommends that staff have competencies in monitoring, measurement, interpretation and prompt response to the acutely ill patient. This is essential to the effective use of early warning systems. ¹⁹ Trigger thresholds for track and trigger systems should be set locally and reviewed regularly.

Identifying patients at risk of AKI means their serum creatinine levels can be monitored for changes while preventative measures are taken to avoid injury or deterioration. Evidence shows poor outcomes once AKI has progressed, making prevention even more important.^{1,7,20}

Sick day medication guidance

Prescribed medicines are implicated in around 20% of cases of AKI.²¹ When vulnerable patients develop dehydrating illnesses such as diarrhoea, vomiting or infection, they are at risk of developing hypotension. If they are taking antihypertensive medications e.g. ACE inhibitors, angiotensin-II receptor antagonists

or diuretics it may be advisable to temporarily withhold the medication.⁷

Think Kidneys' have issued an interim statement on sick day medication guidance for patients in the community when they are acutely ill. The guidance urges caution on advising patients managing their own medication and advises this should be done in discussion with their GP.²² The statement does not cover clinical management of acutely ill in-patients.

Fluid Management

Maintaining adequate hydration and optimal mean arterial pressure, avoiding hypotension, and minimizing exposure to nephrotoxic agents are general strategies that help reduce the risk of AKI in hospital in-patients.⁷ NICE guidance recommends that ward reviews should include assessment and management of patients' fluid and electrolyte needs.²³ Patients should be encouraged to help maintain their own hydration by drinking fluids, with intravenous (IV) fluid therapy only used when oral or enteral intake is insufficient.^{19,23}

IV fluids should be prescribed, delivered and monitored in accordance with a protocol following NICE recommended algorithms for Resuscitation, Routine maintenance, Replacement, Redistribution and Reassessment, depending on the clinical needs of the patient.²³ The need for careful fluid management has been highlighted in a systematic review which found an association between fluid overload and increased mortality in critically ill adult patients with AKI.²⁴

A review of interventions to prevent contrast-induced AKI found little high quality evidence, and concluded that at present hydration/volume expansion is the most effective preventative measure.²⁵ Another review found no difference in effectiveness of oral vs IV hydration in the prevention of contrast-induced AKI.²⁶ A third review found that mannitol had no benefits and was actually detrimental for preventing contrast-induced AKI.²⁷

Early and aggressive fluid resuscitation for patients with sepsis is recommended by the Surviving Sepsis Campaign.²⁸ The use of crystalloids rather than hydroxyethyl starches (HES) is supported by a subsequent systematic review, which found HES was associated with an increase in AKI incidence, need for renal replacement therapy, blood transfusion, and 90-day mortality, in patients with sepsis.²⁹ NICE guidance on sepsis is to be published in July 2016.³⁰

Care bundles

The 100,000 Lives Campaign shows that evidence-based interventions implemented together as a care bundle, have significantly better outcomes than when implemented individually.³¹ A recent prospective observational study found significantly lower risk of in-hospital case fatality in patients who had AKI care bundles completed within 24 hours.¹⁵ A systematic review found bundled care for sepsis significantly

Table 1: Risk factors for AKI in acutely ill patients

- chronic kidney disease
- heart failure
- liver disease
- diabetes mellitus
- history of acute kidney injury
- neurological or cognitive impairment or disability, which may mean limited access to fluids because of reliance on a carer
- hypovolaemia e.g. diarrhoea
- nephrotoxins e.g. NSAIDs, iodinated contrast
- symptoms or history of urological obstruction
- sepsis
- hypertension
- antihypertensive medications in the setting of hypotension

Additional factors for children and young people:

- severe diarrhoea
- haematological malignancy
- hypotension

improved survival, however the eight included studies all had limitations.³² While completion of a care bundle does not require special resources or prior training, it does need all the tasks to be carried out.³¹

NICE guidance recommends the cause of AKI be established and recorded in the patient's notes. When no identified cause of AKI is found at assessment or the patient is at risk of urinary tract obstruction, urgent (within 24 hours) ultrasound of the urinary tract should be performed.⁷

The Royal College of Physicians have recently published Acute Care Toolkit No. 12 on AKI and intravenous fluids which includes the STOP AKI management plan (Table 2).³³

Table 2: STOP AKI management plan

Sepsis - screen for sepsis

Toxins - avoid/stop toxins

Optimise blood pressure - assess volume status

- IV fluids
- hold antihypertensives
- consider vasopressors

Prevent harm

- identify cause/urinalysis
- treat complications
- review medications/fluids

Renal support

The management of AKI should be discussed with a nephrologist within 24 hours of detection of any of the following: the diagnosis is one that may need specialist treatment; there is no clear cause; poor response to treatment; complications arise; stage 3 acute kidney injury; renal transplant; or chronic kidney disease stage 4 or 5.7

Rehabilitation

Information and support for patients and carers

There is no good quality evidence directly looking at what AKI patients and carers want with respect to information and support but based on general qualitative studies of patients on dialysis, NICE provide recommendations.7

Information and support should include details about immediate and long-term treatment options, the likely monitoring required, self-management options, as well as likely prognosis. Discussions should include parents, families and/or carers if appropriate and take place in collaboration with a multidisciplinary team appropriate to the person's individual needs.7

It is recognised that opportunities to provide information or choices regarding lifesaving interventions may be limited because the patients are critically ill and any delay may risk death or significant morbidity.7 The 'Think Kidney's' campaign provides information for the public (www.thinkkidneys.nhs.uk).34

Commissioning for Quality and Innovation (CQUIN)

The care of patients with AKI is a new theme in the national CQUIN scheme for 2015/16. The goal is to improve the follow up and recovery for those who have sustained AKI, reduce the risks of readmission, re-establishing medication for other long term conditions and improving follow up of AKI.35

References

- Stewart J, Findlay G, Smith N, Kelly K, Mason M. Adding Insult to Injury. A review of the care of patients who died in hospital with a primary diagnosis of acute kidney injury (acute renal failure). National Confi dential Enquiry into Patient Outcome and Death (NCEPOD).
- Susantitaphong P, Cruz DN, Cerda J, et al. World Incidence of AKI: A Meta-Analysis. CJASN. 2013;8(9):1482-1493 Susantitaphong P. Correction. CJASN. 2014;9(6):1148 2.
- Coca SG, Singanamala S, Parikh CR. Chronic Kidney Disease after Acute Kidney İnjury: A Systematic Review and Meta-analysis. Kidney Int. 2012;81(5):442-448
- 5. Ftouh S, Lewington A, on behalf of, et al. Prevention, detection and management of acute kidney injury: concise guideline. Clin Med. 2014:14(1):61-65
- Lewington A, Cerdá J, Mehta R. Raising awareness of acute 6. kidney injury: a global perspective of a silent killer. Kidney Int. 2013;84(3):457-67
- 7. National Clinical Guideline Centre. Acute Kidney Injury. Prevention, detection and management up to the point of renal replacement therapy. Commissioned by NICE. August 2013;Clinical guideline <CG 169> Methods, evidence and recommendations
- Lewington A. Communities at risk of developing acute kidney injury. 8. Think Kidneys Risk Workstream. 2015
- NHS England. Acute Kidney Injury (AKI) Algorithm. https://www. 9. england.nhs.uk/patientsafety/akiprogramme/aki-algorithm. Accessed 27 Nov. 2015
- Sawhney S, Mitchell M, Marks A, Fluck N, Black C. Long-term prognosis after acute kidney injury (AKI): what is the role of baseline kidney function and recovery? A systematic review. BMJ Open. 2015;5(1)
- Patient Safety Alert. Standardising the early identification of Acute
- raueric Sarety Aleric. Standardising the early identification of Acute Kidney Injury. Stage Three: Directive. 9 June 2014;http://www.england.nhs.uk/wp-content/uploads/2014/06/psa-aki.pdf Sawhney S, Fluck N, Marks A, et al. Acute kidney injury—how does automated detection perform? Nephrol Dial Transpl. April 28, 2015 Alam N, Hobbelink EL, van Tienhoven AJ, van de Ven PM, Jansma EP, Nanayakkara PWB. The impact of the use of the Early Warning Score (FWS) on patient outcomes: A systematic review Provinces Score (EWS) on patient outcomes: A systematic review. Resuscitation. 2014;85(5):587-594
- McNeill G, Bryden D. Do either early warning systems or emergency response teams improve hospital patient survival? A systematic review. Resuscitation. 2013;84(12):1652-1667 Kolhe NV, Staples D, Reilly T, et al. Impact of Compliance with a

- Care Bundle on Acute Kidney Injury Outcomes: A Prospective Observational Study. PLoS ONE. 2015;10(7):e0132279 16. Lin X, Yuan J, Zhao Y, Zha Y. Urine interleukin-18 in prediction of
- acute kidney injury: a systemic review and meta-analysis. J Nephrol. 2015;28(1):7-16
- Obermüller N, Geiger H, Weipert C, Urbschat A. Current developments in early diagnosis of acute kidney injury. Int Urol Nephrol. 2014;46(1):1-7
- Shao X, Tian L, Xu W, et al. Diagnostic Value of Urinary Kidney Injury Molecule 1 for Acute Kidney Injury: A Meta-Analysis. PLoS ONE. 2014;9(1):e84131
- Centre for Clinical Practice at NICE. Acutely ill patients in hospital:
- Recognition of and response to acute illness in adults in hospital. NICE clinical guideline 50. July 2007
 Greenberg JH, Coca S, Parikh CR. Long-term risk of chronic kidney disease and mortality in children after acute kidney injury: a
- brug and Therapeutics Bulletin. Sick day rules in kidney disease. 2015;53(4):37
- Griffith K, Ashley C, Blakeman T, et al. "Sick day rules" in patients at risk of Acute Kidney Injury: an Interim Position Statement from the Think Kidneys Board. 'Think Kidneys'. 8 July 2015; Version 6
- National Clinical Guideline Centre. Intravenous fluid therapy in adults in hospital. NICE guidelines [CG174] Methods, evidence and recommendations. December 2013
- Zhang L, Chen Z, Diao Y, Yang Y, Fu P. Associations of fluid overload with mortality and kidney recovery in patients with acute kidney injury: A systematic review and meta-analysis. J Crit Care. 2015;30(4):860
- Au T, Bruckner A, Mohiuddin S, Hilleman D. The Prevention of Contrast-Induced Nephropathy. Ann. Pharmacother. 2014;48(10):1332-1342
- Cheungpasitporn W, Thongprayoon C, Brabec BA, Edmonds PJ, O'Corragain OA, Erickson SB. Oral Hydration for Prevention of Contrast-Induced Acute Kidney Injury in Elective Radiological Procedures: A Systematic Review and Meta-Analysis of Randomized Controlled Trials. North Am J Med Sci. 2014;6(12):618-624
- Yang B, Xu J, Xu F, et al. Intravascular Administration of Mannitol for Acute Kidney Injury Prevention: A Systematic Review and Meta-Analysis. PLoS ONE. 2014;9(1):e85029
- Dellinger RP, Levy MM, Rhodes A, et al. Surviving sepsis campaign: international guidelines for management of severe sepsis and septic shock: 2012. Crit Care Med. 2013;41(2):580-637 Serpa Neto A, Veelo DP, Peireira VGM, et al. Fluid resuscitation
- with hydroxyethyl starches in patients with sepsis is associated with an increased incidence of acute kidney injury and use of renal replacement therapy: A systematic review and meta-analysis of the literature. J Crit Caré. 2014;29(1):185.e181-185.e187
- National Institute for Health and Care Excellence. NICE Guidance: Sepsis (In progress). https://www.nice.org.uk/guidance/ indevelopment/gid-cgwave0686. Accessed 1 Sept 2015.
- Resar R, Griffin FA, Haraden C, Nolan TW. Using Care Bundles to Improve Health Care Quality. IHI Innovation Series white paper. Cambridge, Massachusetts: Institute for Healthcare Improvement. 2012.
- Barochia AV, Cui X, Vitberg D, et al. Bundled care for septic shock: An analysis of clinical trials*. Crit Care Med. 2010;38(2).
- Royal College of Physicians. Acute care toolkits. https://www. rcplondon.ac.uk/projects/acute-care-toolkits. Accessed 27 Nov 2015.
- Think Kidneys Programme Board. Acute Kidney Injury Information for the Public Version 1. 2014.
- NHS England. Commissioning for Quality and Innovation (CQUIN) Guidance for 2015/16. March 2015.

About Effectiveness Matters

Effectiveness Matters is a summary of reliable research evidence about the effects of important interventions for practitioners and decision makers in the NHS and public health. This issue is produced by CRD in collaboration with the Yorkshire and Humber AHSN Improvement Academy. Effectiveness *Matters* is extensively peer reviewed.

