



University of Dundee

Post-operative AKI

Bell, Samira; Prowle, John

Published in: Journal of the American Society of Nephrology

DOI: 10.1681/ASN.2018111127

Publication date: 2019

Document Version Peer reviewed version

Link to publication in Discovery Research Portal

Citation for published version (APA): Bell, S., & Prowle, J. (2019). Post-operative AKI: prevention is better than cure? Journal of the American Society of Nephrology, 30(1), 4-6. https://doi.org/10.1681/ASN.2018111127

General rights

Copyright and moral rights for the publications made accessible in Discovery Research Portal are retained by the authors and/or other copyright owners and it is a condition of accessing publications that users recognise and abide by the legal requirements associated with these rights.

Users may download and print one copy of any publication from Discovery Research Portal for the purpose of private study or research.
You may not further distribute the material or use it for any profit-making activity or commercial gain.
You may freely distribute the URL identifying the publication in the public portal.

Take down policy If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

Post-operative AKI - prevention is better than cure?

Samira Bell^{1,2}

John Prowle^{3,4}

¹Renal Unit, Ninewells Hospital, Dundee, UK

²Division of Population Health Science and Genomics, School of Medicine, University of Dundee, Dundee, UK.

³Adult Critical Care Unit and Department of Renal Medicine and Transplantation, The Royal London Hospital, Barts Health NHS Trust, London, UK

⁴William Harvey Research Institute, Queen Mary University of London, London, UK

Corresponding Author:

Samira Bell Division of Population Health Science and Genomics, School of Medicine, University of Dundee, Dundee, UK. Samira.bell@nhs.net

Word Count: 907

Running title: Prevention is better than cure?

Acute Kidney Injury (AKI) could be considered as a sentinel post-operative complication as it is relatively common and strongly associated with increased short and long term risk of death, the development of other post-operative complications, increased length of hospital stays and potentially the development of progressive chronic kidney disease ¹. Consequently patients with AKI after surgery constitute an area of unmet clinical need and a major burden on healthcare resources. In the absence of effective interventions to treat established AKI, pre-operative identification of patients at high risk of AKI allows for peri-operative optimisation to prevent or reduce the severity of AKI. A number of large retrospective studies have examined the associations of post-operative AKI in multivariable modelling, however few of these have led to externally validated models that have been operationalised to give predictions in a clinical setting particularly in patients undergoing non-cardiac surgery². There are currently several validated AKI-risk scores for patients undergoing cardiac surgery, one in orthopedic surgery³ and most recently, a model derived through machine-learning capable of forecasting the development of a range of post-operative complications, including AKI, across a range of surgical settings⁴. In this issue of the Journal of American Society of Nephrology, Park et al address this gap in the literature through the development of the SPARK classification, an externally validated pre-operative AKI risk score for noncardiac surgery⁵. This model was developed in 51,041 patients and validated in 39,764 patients from South Korea and utilises 9 pre-operative variables: age, sex, baseline estimated glomerular filtration rate, urinary albuminuria, expected surgery duration, emergency operation, diabetes mellitus, reninangiotensin-aldosterone system blockade usage, hypoalbuminemia, anaemia and hyponatremia. In contrast to other risk scores, this risk score additionally considers AKI severity by predicting a composite 'critical AKI'; outcome of Stage 2 or greater AKI, need for renal replacement therapy within 90 days of AKI or death occurring after any AKI diagnosis. By allowing pre-operative risk stratification of patients this tool could assist clinicians in a number of ways. It can inform discussions with patients prior to

surgery, allowing for a clearer and more candid quantification of the risks of an important medical complication of surgery and the potential benefits of surgical treatment to be better weighed against risks of complications and adverse longer term outcomes. As we are increasingly facing an ageing, multimorbid population we need to carefully consider these adverse potential consequences of which can often affect organ systems remote from the site of surgical disease. In addition, identifying high risk patients can guide pre-operative planning by influencing whether patients would benefit from more intensive monitoring in the post-operative period possibly in a critical care setting.

Nevertheless, it could be argued that risk stratification is pointless since it remains unclear what the proposed intervention(s) should be for patients at high risk for AKI, and whether these interventions are in fact effective. However, recent studies have shown that risk stratification of patients at high risk of AKI diagnosis in the post-operative setting might enable effect intervention with a "bundle" of AKIdirected preventative measures. These studies examined the implementation of an AKI prevention bundle derived from recommendations in the Kidney Disease Improving Global Outcomes (KDIGO) 2012 AKI guidelines⁶ to patients deemed at high risk of AKI based on post-operative urinary AKI biomarkers (insulin-like growth factor-binding protein 7 (IGFBP7) and tissue inhibitor of metalloproteinases-2 (TIMP-2)). These interventions comprised of the avoidance of potentially nephrotoxic agents, close monitoring of serum creatinine and urine output, and optimization of volume status and haemodynamic parameters. In cardiac surgery this approach has been shown to reduce post-operative AKI diagnosis, but without evidence of benefit for longer term renal outcomes⁷, while application of a similar strategy in general surgery was overall negative, but demonstrated a signal of benefit in a subgroup of patients with moderately elevated AKI biomarkers suggestive of benefit in early but not more established AKI⁸. Whilst preliminary these results do suggest a KDIGO based AKI prevention bundle would be the obvious intervention in patients stratified as high risk for peri-operative AKI. However, AKI biomarkers are costly and not currently routinely or widely available and have thus far been only utilised in relatively small

studies with equivocal results. The application of a risk score using readily available pre-operative parameters could provide a relatively cheap and simple means of risk stratification that could be applicable to post-operative AKI prevention and management to many patients in a variety of incomesettings. Importantly in the setting of intensive care unit admissions, AKI risk-prediction algorithms based on routinely available data have been shown to have comparable diagnostic performance to AKIbiomarkers⁹, reinforcing the role of predictive models as an alternative or complement to biomarkers in risk-stratification for clinical decision making.

The SPARK risk score is thus a promising tool that could be investigated in prospective studies for targeted AKI preventative bundled intervention, either as randomised studies or perhaps, given the established nature of the components of bundled care, as a quality improvement intervention. Currently, this model is limited in its generalizability as it was both developed and validated in Korean patients and further external validation is required to ensure its applicability to other populations worldwide before consideration of widespread implementation. However, the approach of riskstratification of patients for sentinel surgical complications is likely to be an increasingly important method of improving, not just peri-operative care, but the ongoing global healthcare of surgical patients and the ability of patients and doctors to make informed decisions about their surgical treatment.

References

- 1. Meersch, M, Schmidt, C, Zarbock, A: Perioperative Acute Kidney Injury: An Under-Recognized Problem. *Anesth Analg*, 125: 1223-1232, 2017.
- 2. Hodgson, LE, Sarnowski, A, Roderick, PJ, Dimitrov, BD, Venn, RM, Forni, LG: Systematic review of prognostic prediction models for acute kidney injury (AKI) in general hospital populations. *BMJ Open*, 7: e016591, 2017.
- 3. Bell, S, Dekker, FW, Vadiveloo, T, Marwick, C, Deshmukh, H, Donnan, PT, Van Diepen, M: Risk of postoperative acute kidney injury in patients undergoing orthopaedic surgery--development and validation of a risk score and effect of acute kidney injury on survival: observational cohort study. *Bmj*, 351: h5639, 2015.
- Bihorac, A, Ozrazgat-Baslanti, T, Ebadi, A, Motaei, A, Madkour, M, Pardalos, PM, Lipori, G, Hogan, WR, Efron, PA, Moore, F, Moldawer, LL, Wang, DZ, Hobson, CE, Rashidi, P, Li, X, Momcilovic, P: MySurgeryRisk: Development and Validation of a Machine-learning Risk Algorithm for Major Complications and Death After Surgery. *Ann Surg*, 2018.
- 5. Park S, CH, Park SW, Lee S, Kim KS, Yoon HJ, Park J, Choi Y, Lee S, Kim JH, Kim S, Chin HJ, Kim DK, Joo KW, Kim YS, Lee H.: Simple postoperative AKI risk (SPARK) classification before non-cardiac surgery: a prediction index development study with external validation. *J Amer Soc Nephrology,* In press, 2018.
- 6. KDIGO Clinical Practice Guideline for Acute Kidney Injury. *Kidney Int Suppl,* 2: 1-138, 2012.
- 7. Meersch, M, Schmidt, C, Hoffmeier, A, Van Aken, H, Wempe, C, Gerss, J, Zarbock, A: Prevention of cardiac surgery-associated AKI by implementing the KDIGO guidelines in high risk patients identified by biomarkers: the PrevAKI randomized controlled trial. *Intensive Care Med*, 43: 1551-1561, 2017.
- Bocze, I, Jauch, D, Gotz, M, Kennedy, P, Jung, B, Zeman, F, Gnewuch, C, Graf, BM, Gnann, W, Banas, B, Bein, T, Schlitt, HJ, Bergler, T: Biomarker-guided Intervention to Prevent Acute Kidney Injury After Major Surgery: The Prospective Randomized BigpAK Study. *Ann Surg*, 267: 1013-1020, 2018.
- 9. Flechet, M, Guiza, F, Schetz, M, Wouters, P, Vanhorebeek, I, Derese, I, Gunst, J, Spriet, I, Casaer, M, Van den Berghe, G, Meyfroidt, G: AKIpredictor, an online prognostic calculator for acute kidney injury in adult critically ill patients: development, validation and comparison to serum neutrophil gelatinase-associated lipocalin. *Intensive Care Med*, 43: 764-773, 2017.