Variability in renal dysfunction defining criteria and detection methods in intensive care units: Are the international consensus criteria used for diagnosing renal dysfunction?**


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Received 6 August 2011; accepted 15 October 2011
Available online 2 July 2012

Abstract

Objective: To evaluate variability in the detection and prevention of acute kidney injury (AKI) in the intensive care unit (ICU), and application of the international recommendations in this field (Acute Dialysis Quality Initiative [ADQI] and Acute Kidney Injury Network [AKIN]).

Setting: A prospective, observational, multicenter study.

Setting: A total of 42 ICUs in 32 hospitals (78% in third level hospitals and 70.7% general units) recruited for a study on the prevalence of AKI (COFRADE).

Interventions: Survey.

Variables: Aspects related to AKI detection and prevention and renal replacement therapy (RRT) protocols.

Results: The method used for estimating glomerular filtration rate was serum creatinine (Crs) in 36.6%, creatinine clearance in 41.5% and equations in 22%; none reported using cystatin-C. Only 39.1% ICUs acknowledged the use of stratification systems (13 RIFLE and 3 AKIN).

A total of 48.8% ICUs had no written protocols for AKI prevention, 31.7% reported using them only for contrast nephropathy, 7.3% for nephrotoxic drugs and 12.2% for both.

In contrast, 63.4% participants had written protocols for RRT, 70.7% had implemented a training program, and 53.7% had some method for adjusting doses of drugs when on RRT.

Conclusions: We observed an important variability regarding diagnostic criteria and prevention of AKI in Spanish ICUs, the application of ADQI or AKIN recommendations still being low in our units. RRT seems to generate more concern among our intensivists than AKI management.

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** This study has the scientific support of the SEMICYUC.

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VARIABILIDAD EN LOS CRITERIOS DE DEFINICIÓN Y MÉTODOS DE DETECCIÓN DE LA DISFUNCIÓN RENAL EN LAS UNIDADES DE CUIDADOS INTENSIVOS: ¿SE APlicAN LOS CONSENSOS INTERNACIONALES PARA EL DIAGNÓSTICO DE LA DISFUNCIÓN RENAL?

Resumen

Objetivo: Evaluar la variabilidad en la detección y prevención de la disfunción renal aguda (DRA) en las unidades de cuidados intensivos (UCI), así como las aplicaciones de recomendaciones internacionales en este campo (Acute Dialysis Quality Initiative [ADQI] y Acute Kidney Injury Network [AKIN]).

Diseño: Estudio multicéntrico prospectivo descritivo.

Ámbito: Un total de 42 UCI (70,7% generales) en 32 hospitales (78% de tercer nivel), participantes en un estudio de prevalencia de DRA (COFRADE).

Intervenciones: Encuesta.

Variables de interés: Métodos de detección y prevención de DRA y existencia de protocolos para depuración extrarrenal (TDE).

Resultados: Para la estimación del filtrado glomerular se usa la creatinina sérica en el 36,6%, el aclaramiento de creatinina en el 41,5%, ecuaciones basadas en una muestra de creatinina sérica en el 18% y creatinina sérica aislada en el resto; en ningún caso se utiliza la cistatina-C.

Solo el 39,1% de las UCI aplican sistemas de estratificación (13 RIFLE y 3 AKIN). El 48,8% no cuenta con protocolos de prevención de DRA, el 31,7% los tiene para nefropatía por contraste, el 7,3% para manejo de nefrotóxicos y el 12,2% para ambos.

Por contra, el 63,4% de las unidades cuenta con protocolos de manejo de TDE, el 70,7% aplica programas de formación continuada para estas y el 53,7% cuenta con protocolos de ajuste de dosis de fármacos en estos casos.

Conclusiones: Existe amplia variabilidad en cuanto al método de detección de la DRA en nuestras unidades, con escasa aplicación de sistemas de estratificación de daño. El manejo de las TDE despista más inquietud que la prevención o el diagnóstico de la DRA.

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INTRODUCCIÓN

El desarrollo de AKI en los pacientes críticos produce un aumento en la morbilidad-mortalidad y en los costos de tratamiento.1-4 Según el estudio FRAMI,1 la incidencia en nuestro medio fue del 5.7% 10 años atrás—esta figura es similar a la incidencia reportada por el estudio BEST1 que involucró al 92% de los pacientes de 12 unidades intensivas durante un año,1 lo cual sugiere que el uso de estos métodos es un requisito en el manejo de AKI. De hecho, el uso de estas técnicas permite la identificación temprana de la disfunción renal aguda, lo que puede ser un factor de pronóstico independiente.1-4,5

Hoy en día, las recomendaciones de ADQI6 y posteriormente AKIN7 parece que está cambiando, y el manejo de AKI es ahora basado en criterios estándarizados—el advenimiento de un lenguaje común que es accesible de forma uniforme y fácilmente reproducible. Sin embargo, la brecha en el manejo de AKI en nuestro entorno es aún significativa. Lo anterior obliga a considerar la necesidad de una revisión y actualización de los criterios de AKI basados en la evidencia y los estándares internacionales de tratamiento de AKI.6,7,8,9,10

En este estudio, desarrollado en el contexto de la iniciativa COFRADE,11 el objetivo es caracterizar la variabilidad en el diagnóstico y tratamiento del AKI en nuestra unidad de cuidados intensivos y comparar nuestros resultados con la literatura existente.12-17

MATERIALES Y MÉTODOS

El presente estudio representa el período inicial de una observación prospectiva que tiene como objetivo la detección de la disfunción renal aguda en el contexto de la iniciativa COFRADE,11 diseñada para analizar la incidencia de AKI y su tratamiento. Este estudio se divide en dos fases: (a) un primer período en el que se recolectaron datos de las unidades participantes y (b) un segundo período en el que se evaluó el tratamiento de AKI en nuestras unidades um participantes.

El estudio se llevó a cabo en la UCI de nuestra institución y se comparó con los resultados de la literatura existente.12-17

Nuestro principal objetivo es presentar el resultado del primer estudio y proporcionar una perspectiva del contexto actual de la disfunción renal aguda en nuestras UCI.
The study was approved by the Clinical Research Ethics Committees of each of the participating centers.

The data are presented as values (percentages), and comparisons were made using the chi-squared test, with a significance level of 95% \( (p < 0.05) \).

**Results**

A total of 32 hospitals participated in the COFRADE study: 10 in the Community of Madrid, 9 in Andalucia, three in Catalonia and the Valencian Community, and one in the Canary Islands, Castilla La Mancha, Castilla-León, the Community of Murcia, Extremadura, Galicia and the Basque Country. Twenty-three of these hospitals were third-level centers.

A total of 42 ICUs were involved, with a total of 826 beds. Of these ICUs, 30 were general Units, three medical Units, three surgical Units, and 6 specialized Units.

In relation to the estimation of glomerular filtration, 24-h urine creatinine clearance was applied in 12 Units (28.6%), and short duration urine creatinine clearance in 5 Units (11.9%). In turn, 9 Units (21.4%) made use of equations involving the determination of plasma creatinine (the Cockroft–Gault\(^6\) formula in 7 Units and the MDRD\(^7\) in two Units), while the remaining 16 Units (38.1%) only used Crs for this purpose. In no case was cystatin-C used as estimation method. There were no differences dependent upon the type of hospital or Unit surveyed (Fig. 1).

Fourteen ICUs (33.3%) adopted the RIFLE scale, while one Unit (2.4%) incorporated the AKIN system, and two Units (4.8%) adopted both scales. In contrast, 25 Units (59.5%) admitted that they applied neither system. Again in this case there was no correlation between the type of Unit and the use of the different scales, though there were differences referred to the type of hospital \( (p < 0.05) \) (Fig. 2).

Twenty Units (47.6%) had no defined protocols for the prevention of AKI, while 14 Units (33.3%) presented contrast utilization protocols, three Units (7.1%) had nephrotoxic drug use protocols, and 5 Units (11.9%) had both contrast and drug use protocols. The hospital or Unit type was not related to this variable.

Thirty Units (71.4%) claimed to have continued training programs for the management of RRT. In 26 Units (61.9%) there were specific protocols for application of the technique (without differences dependent upon the type of hospital or ICU), and of these Units, 24 had a drug adjustment protocol for patients subjected to RRT.

**Discussion**

Following the publication of the recommendations of the ADQI\(^8\) and posteriorly the AKIN\(^7\) referred to the need to stratify AKI, different studies have described a significant change in the way to approach renal dysfunction, and in fact the incidence figures\(^10\) are now considered to be far higher than the previous reference values\(^3,4\) for general critical care patients. According to our results, and despite the existence of these recommendations, there is still great variability regarding the AKI detection methods used, with
little application of the injury stratification systems. In this context, it is clear that the management of RRT (although only a part of the problem in patients with AKI) generates more concern than the prevention or diagnosis of AKI among intensivists in this country.

In recent years there have been changes in the physiopathological concepts of AKI, particularly referred to dysfunction of ischemic origin. This is important, considering that acute tubular necrosis is the main cause of AKI in critical patients, especially in relation to severe sepsis.\(^{11,12}\) This increased knowledge of the physiopathology of renal dysfunction from the moment of primary injury has led to an interest in improving the diagnosis and developing new and better methods to facilitate early diagnosis and thus minimize secondary kidney damage. In this context, the study of new kidney injury biomarkers is one of the fields of interest in AKI.\(^{13,14}\)

The recommendations of the ADQI\(^6\) referred to the diagnosis and risk stratification of AKI, as reflected in the RIFLE system, were published in 2004. This system is based on the detection of evolutive changes referred to patient basal glomerular filtration (GF) through changes in urine volume or based on the estimation of GF from Crs or creatinine clearance (CICr). One year later, in 2007, the AKIN\(^\text{©}\) group redefined this system once it became clear (initially in studies in heart surgery and posteriorly in other populations)\(^{15,16}\) that also small Crs increments imply a poorer prognosis, independently of the basal situation. The AKIN system stratifies patients into three levels according to changes in diuresis or Crs increments, without considering other GF estimators, and incorporating absolute Crs elevations (an increase of 0.3 mg/dl would classify a patient as AKIN 1, independently of the previous basal level).

On furthermore considering that when Crs rises to above 2 mg/dl, the functional kidney mass (i.e., the "kidney reserve") has dropped to 50%,\(^{17}\) it is not surprising that the RIFLE and AKIN systems, by detecting small Crs increments, are of great interest, and that the previous incidence data—mostly based on this 2 mg/dl cutoff value—possibly greatly underestimated the true incidence of AKI. It also must be considered that both systems (RIFLE and AKIN) have already been widely validated and show good correlation to mortality.\(^{18-22}\) However, despite the interest in these new defining systems, to date they have shown scant penetration in Spanish ICUs, since less than 40% of our Units use such instruments.

Another aspect to be taken into account in assessing the detection of AKI is the GF estimator used. In clinical practice, the measurement of urine output is a simple method and in most cases anticipates Crs elevation; as a result, it is widely used for the initial detection of renal dysfunction,\(^{23}\) although as a parameter it offers very poor specificity. Both systems define Crs as an alternative estimator, since it is easy to determine and is universally available. However, Crs elevation shows a delay or lag after the lowering of GF, and does not exhibit a linear relationship with decreasing GF. These aspects may pose an important problem for a system precisely aiming to secure the early detection of reductions in GF.

Taking into account that kidney function in critical patients is not stable, it seems reasonable that serial measurements of CICr (which can be made with short duration diuresis, thereby allowing for easy reproducibility)\(^{21}\) are able to detect such changes earlier than Crs—though the use of this estimator is only contemplated in the RIFLE (which in our opinion represents an advantage over AKIN). In our experience there are discrepancies in stratifying patients with the RIFLE system according to whether it is based on Crs or CICr, in the sense that utilization of the latter parameter is more closely related to the prognosis.\(^{21}\) It is notorious that in the surveyed ICUs, CICr was the parameter most commonly used to estimate GF, and it is even more surprising that 70% of the Units used determinations based on 24-h urine—these data being discordant with those obtained 10 years ago in the FRAMI study,\(^4\) where few Units made use of CICr.

Crs is the next most common estimator used in our ICUs, though in fact it constitutes the most common estimator (59%) on considering those Units that use it incorporated to equations. This is not surprising, since the method is simple and reproducible, and very familiar in the clinical setting. What is surprising is that Crs is not incorporated in either of the available and validated systems (RIFLE or AKIN) with the purpose of stratifying patients and of offering a tool for follow-up capable of contributing relevant clinical information (this in our view being the most attractive aspect of these classification systems).

On the other hand, we see that the use of formulas remains a generalized practice despite the fact that they were specifically developed for application in chronic patients. However, a significant percentage of centers (21%) apply them to estimate GF in patients with AKI, even though none of them are useful in critically ill patients.\(^{24-25}\)

None of the Units, regardless of their size, have incorporated biomarkers or cystatin-C in routine use, though we have evidenced that their application is being studied in some Units (including our own), in concrete patient groups. The incorporation of these new biomarkers may give added impulse to the early detection of AKI, though considering the current situation found in Spanish ICUs (as reflected by our data), their incorporation to the clinical setting does not appear due anytime soon.

Of note is the observation that no intervention for the diagnosis, stratification or prevention of AKI has been protocized in the majority of our Units, since less than 50% of them admit the use of protocols in this sense. Furthermore, those protocols that do exist are particularly centered on the prevention of nephropathy referred to contrast injection, and in only very sporadic cases to drug adjustment. Specifically, only 12% of the Units have protocols which include these two very important aspects. This is in stark contrast to the use of RRT, since most of the Units have written protocols for applying such techniques. This may be attributed to the fact that the diagnosis of AKI is well assimilated in our usual clinical practice, while RRT is a relatively recent practice with which we have had to become quickly familiarized in view of the advances made in this field. This has created the need to learn and reflect the practice in written protocols; indeed, almost 60% of the Spanish Units have protocols for drug adjustment associated to the use of RRT, while only 7% have protocols for drug adjustment in relation to AKI. In fact, in relation to training, over 70% of the Units reported having active training programs in RRT.
The described results of course are based on the analysis of a survey, and so must be viewed with caution. On the other hand, and as the main weakness of this study, the sample comprised a highly selected group of units—which questions the external validity of the findings. Nevertheless, this bias further emphasizes the problem, since the fact that the units in the study largely participated because of their involvement and interest in AKI leads us to suspect that the results obtained in a larger study would have proven even less adequate.

We consider it necessary to change the strategy of our approach to AKI, with incorporation of the recommendations of the ADQI and AKIN conferences to our daily clinical practice, since this would directly result in early detection. Simply acquiring the habit of detecting patients at risk and of reflecting the fact in their case history can help improve management and reduce secondary renal damage, independently of the notion that different software applications (where available) can draw our attention through specific alerts when kidney function is altered—as is the case in our center. Undoubtedly, the incorporation of written protocols for the prevention and detection of AKI, and for RRT, would have a positive effect upon patient safety—facilitating secondary prevention (limitation of damage) and avoiding classification error and iatrogenic problems (“dialytrauma”) in RRT.

Conflicts of interest

The authors declare no conflicts of interest.

Appendix 1.


References


