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## Improving Incident Fistula Rates: A Process of Care Issue

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In the last decade the proportion of hemodialysis patients dialyzing with an arteriovenous fistula (prevalent fistula rate) in Canada, Europe, and Australia/New Zealand have been relatively high (50-90%)<sup>1</sup>. In the United States, the prevalent fistula rate has improved dramatically and is now 57%<sup>2</sup>. Achievement of these high rates has been due in part to national nephrology society guidelines and vascular access initiatives, such as the Kidney Disease Outcomes Quality Initiative (K/DOQI)<sup>3</sup>, Fistula First Initiative<sup>2</sup>, European Best Practice Guidelines<sup>4</sup>, and Canadian Society of Nephrology Clinical Practice Guidelines<sup>5</sup> which have focused on improving vascular access care. These guidelines and initiatives have targeted the following processes: (1) patient preparation for permanent access placement, (2) selection of type of permanent access with an emphasis on fistula evaluation and placement, and (3) cannulation. The guidelines and initiatives recognize that a multidisciplinary team comprised of nephrologists, vascular access surgeons, nephrology nurses, and vascular access coordinators are essential to accomplishing these goals <sup>2, 3, 6</sup>. However, Canada, Australia/New Zealand, and several European countries with traditionally high prevalent fistula rates have recently reported proportions of incident hemodialysis patients with a fistula (incident fistula rates) below 50%<sup>1</sup>. Furthermore, while the U.S. has seen rapid improvements in fistula rates in the prevalent population, this has not been accompanied by improvements in fistula rates among the incident population which have remained remarkably low at 18%<sup>7</sup>.

In this issue of the *American Journal of Kidney Diseases*, Lopez-Vargas et al. report the results from a prospective multicenter cohort study within an Australian/New Zealand population which evaluated the implementation of national guidelines on timing of vascular access placement and type of vascular access placed in patients who initiated hemodialysis. The main outcome of this study was the proportion of patients with a functional fistula at dialysis initiation. Secondary outcomes included physician, patient, and organizational-barriers responsible for delays in achieving a functional fistula at dialysis initiation <sup>8</sup>. Among the 319 patients who initiated hemodialysis during the six month study period, 39% of patients initiated hemodialysis with a fistula and 57% initiated hemodialysis with a catheter. Other important findings included (1) 66% and 79% of patients were under the care of a nephrologist by 12 and 3 months prior to initiating dialysis, respectively, (2) the median time from surgical referral to surgical evaluation and surgical evaluation to access surgery in this study were approximately 14 and 28 days, respectively, and (3) median eGFR

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at the time of surgical referral, access creation, and dialysis initiation were 7, 7, and 6 ml/  $min/1.73 m^2$ , respectively.

Previous studies <sup>6, 9-11</sup> have suggested that (1) early referral to a nephrologist from a primary care provider for chronic kidney disease (CKD) management, (2) timely discussion with the patient of future renal replacement modality and pre-dialysis education by the nephrologist, (3) referral to vascular access surgeon for evaluation and permanent access placement prior to dialysis initiation, and (4) close follow-up of the maturing fistula (outside of the dialysis unit) with an aggressive intervention policy (endovascular or surgical) for fistulae that fail to mature are important processes required to increase incident fistula rates (Figure 1). Close attention to all of these processes is essential in order to achieve the "holy grail" of pre-emptive fistula placement and successful cannulation with two needles by the dialysis staff at the first dialysis session. While many of the same processes are required to achieve a functional fistula in the prevalent population, the incident population, unlike prevalent dialysis patients, is not a "captive audience" which is seen thrice weekly by a healthcare provider. Thus, early and frequent pre-dialysis education and effective patient-physician interaction, while challenging, is crucial for improving fistula rates in the incident dialysis population.

There are several important observations from this study by Lopez-Vargez et. al that shed light on the processes need for improving incident fistula rates. First, late referral to nephrologists was actually low in their study population. The results from this study are similar to Europe and Canada where 62% and 63% of patients, respectively, were under the care of a nephrologist >12 months before initiating dialysis <sup>12</sup>. In marked contrast, in the United States only 24% and 33% of patients had their initial nephrology evaluation >12 months and 0 to 12 months, respectively, prior to initiating dialysis, and 43% had no nephrology care prior to dialysis initiation <sup>7</sup>. Thus, it does not appear that late nephrology referral is the major reason for low incident fistula rates in the Australia/New Zealand population in this study. Second, median time from surgical referral to evaluation and evaluation to surgical placement of permanent access was also relatively short. Interestingly, in the Dialysis Outcomes and Practice Patterns Study (DOPPS), the United States had one of the lowest median times from surgical referral to evaluation and surgical evaluation to permanent access times at 7 and 7 days, respectively, but the worst incident fistula rate among surveyed countries<sup>1</sup>. Thus, short times to surgical evaluation and placement of permanent access may not be relevant, if early referral to a nephrologist and timely referral to a surgeon does not also occur. Finally, the most striking observation from this study was low median eGFR at the time of surgical referral and access creation<sup>8</sup>. Even with early referral to a nephrologist and relatively short wait times for surgical evaluation and vascular access placement, the eGFRs suggest that the timing of surgical referral and vascular access placement occurred in the very late stages of the CKD and very close to the time of dialysis initiation. Therefore, what are the specific barriers that lead to delayed referral for surgical access evaluation and access placement?

Table 2 in the manuscript by Lopez-Vargas et al. outlines a model of perceived barriers to initiation of hemodialysis with a permanent access based on patient, physician, and organizational-levels <sup>8</sup>. An important component of the vascular access process among those patients who have pre-dialysis nephrology care is interaction between the nephrologist and patient. It is challenging for nephrologists to accurately predict the rate of kidney disease progression and whether the patient will survive until kidney failure<sup>13, 14</sup>. Very frequently, dialysis initiation occurs in conjunction with a concurrent illness or hospitalization <sup>15, 16</sup>. Therefore, it is difficult to determine the appropriate time to initiate pre-dialysis education for patients, discuss selection of renal replacement modalities, and subsequently when to refer patients for vascular access evaluation and surgery. The authors in this study reported

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that the majority of patients, 65%, attended pre-dialysis education sessions, and very few patients referred for surgical access evaluation and access placement refused or did not attend <sup>8</sup>. What is unclear from this data is the level of eGFR at the time of initial pre-dialysis education and the interval from initial pre-dialysis education to surgical evaluation and access placement. This information may provide some further insight into whether the barriers are at the level of the physician (uncertainty by the nephrologist whether the patient's CKD will progress or patient will survive to dialysis initiation) or patient (denial of CKD and need for dialysis). Furthermore, it is quite evident from this study that early placement of permanent access enhances the likelihood that a patient initiates dialysis with a fistula or graft, and not a catheter (OR 0.22, 95% CI 0.10-0.50; per 5 ml/min/1.73 m<sup>2</sup> higher in eGFR).

From an organizational-level, there are currently no specific guidelines from nephrology societies in the United States or other countries that address implementing a specific plan or benchmarks for vascular access milestones to be achieved as GFR declines. The availability and widespread use of such benchmarks would likely address both patient-based and physician-based barriers and improve incident fistula rates. In an attempt to provide such benchmarks, Hakim et al. have proposed an algorithm for the planning and placement of permanent access based on eGFR in pre-dialysis patients 17 which includes: (1) initiation of pre-dialysis education at an eGFR 30 ml/min/1.73 m<sup>2</sup>, (2) surgical evaluation and permanent access placement at an eGFR 20ml/min/1.73 m<sup>2</sup>, and (3) a mature fistula in place and ready to be used for dialysis at an eGFR 10 ml/min/1.73<sup>2</sup>. Using eGFR-based guidelines would provide objective criteria for patient education and nephrologist decisionmaking in regards to referral for surgical evaluation and placement of permanent access, with a specific emphasis on fistulae. To support this point, in the study by Lopez-Vargas et. al., among patients who initiated dialysis with a catheter, 44% never received pre-dialysis education and 48% were never referred for surgical evaluation. Furthermore, among all centers in this study, the median time from first nephrology evaluation to dialysis initiation was rather long, ranging from 1.1 to 3.4 years <sup>8</sup>. Moreover, eGFR-based guidelines would ultimately ensure timely placement of permanent access in all advanced CKD patients and allow for ample time for interventions to promote maturation, if necessary, as fistulae require several months for maturation <sup>11, 18-20</sup>. In this study, among patients with permanent access placement, the median time from permanent access placement to initiation of dialysis was significantly longer in patients who initiated dialysis with a fistula or graft vs catheter (28 vs 8 weeks; p=0.001). However, some may argue that this approach may lead to unnecessary placements of fistulae in patients with slowly progressive kidney disease, particularly in the elderly population who may die before ever requiring dialysis<sup>21</sup>.

What new information can be taken from this study? First, a major barrier influencing incident fistula rates is the delayed referral for surgical evaluation and fistula placement which occurred at an eGFR of 7 ml/min/ $1.73^2$  for both events in this study <sup>8</sup>. Second, among patients who initiated dialysis with a catheter a large percentage of these patients never received predialysis education and were not referred for surgical evaluation <sup>8</sup>. The majority of countries participating in the DOPPS study have a high proportion of patients who have been under the care of a nephrologist for > 1 year prior to dialysis initiation <sup>12</sup>. Thus, the primary focus now should be on the failures in the processes of care that occur after the patient has been under the care of a nephrologist. It appears that the most beneficial intervention towards improving incident fistula rates would be targeting earlier referral for surgical evaluation and permanent access placement. Future research studies need to directly evaluate in more detail the patient and nephrologist perspectives about pre-dialysis education and permanent vascular access placement prior to initiation of dialysis to determine the primary barriers leading to delayed referral for surgical evaluation and permanent <sup>25</sup>.

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