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Arterial Spin Labeling MRI for Assessment of Perfusion in Native and Transplanted Kidneys

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Abstract

Purpose—To apply a magnetic resonance (MR) arterial spin labeling (ASL) technique to evaluate kidney perfusion in native and transplanted kidneys.

Materials and Methods—This study was compliant with the Health Insurance Portability and Accountability Act (HIPAA) and approved by the institutional review board. Informed consent was obtained from all subjects. Renal perfusion exams were performed at 1.5 T in a total of 25 subjects: 10 with native and 15 with transplanted kidneys. A flow-sensitive alternating inversion recovery (FAIR) ASL sequence was performed with respiratory triggering in all subjects and under free-breathing conditions in five transplant subjects. Thirty-two control/tag pairs were acquired and processed using a single-compartment model. Perfusion in native and transplanted

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kidneys was compared above and below an estimated glomerular filtration rate (eGFR) threshold of 60 ml/min/1.73m² and correlations with eGFR were determined.

Results—In many of the transplanted kidneys, major feeding vessels in the coronal plane required a slice orientation sagittal to the kidney. Renal motion during the examination was observed in native and transplant subjects and was corrected with registration. Cortical perfusion correlated with eGFR in native ($r=0.85$, $p=0.002$) and transplant subjects ($r=0.61$, $p=0.02$). For subjects with $eGFR \geq 60$ ml/min/1.73m², native kidneys demonstrated greater cortical ($p=0.01$) and medullary ($p=0.04$) perfusion than transplanted kidneys. For subjects with $eGFR < 60$ ml/min/1.73m², native kidneys demonstrated greater medullary perfusion ($p=0.04$) compared to transplanted kidneys. Free-breathing acquisitions provided renal perfusion measurements that were slightly lower compared to the coached/triggered technique, although no statistical differences were observed.

Conclusion—In conclusion, FAIR-ASL was able to measure renal perfusion in subjects with native and transplanted kidneys, potentially providing a clinically viable technique for monitoring kidney function.

Introduction

Current diagnostic measurements used to assess severity of kidney failure, such as serum creatinine levels, are relatively insensitive to small but potentially significant functional change. They are also non-specific, requiring a biopsy to characterize the underlying cause of renal dysfunction. Early characterization of dysfunction is crucial for transplant patients because a delay in treatment can lead to irreversible nephron loss and accelerate graft failure [1,2]. Transplant patients are often subjected to multiple biopsies to assess dysfunction longitudinally and guide treatment decisions. Biopsies are not optimal for longitudinal assessment because they are painful and can result in bleeding, infection, and even graft loss [3]. Moreover, they are expensive and cannot quantify changes in renal function associated with the identified pathology. These limitations have motivated research to find non-invasive diagnostic tools that can both characterize renal disease and provide earlier detection of functional change.

Functional magnetic resonance imaging (MRI) offers a variety of methods to assess blood flow that are especially useful in the kidney. Not only is renal blood flow essential for kidney viability, but it also plays an integral role in blood filtration and regulation. Nuclear medicine and X-ray computed tomography techniques can measure perfusion, but both require intravenous injection and radiation exposure. MR imaging offers a less invasive method of measuring kidney perfusion and has demonstrated potential in assessing renal disease in both native and transplanted kidneys [4-9]. For example, MR perfusion measurements have correlated with histology in transplanted rat kidneys [7], and human studies indicate that perfusion may help differentiate cyclosporine toxicity, acute rejection, and acute tubular necrosis following transplantation [4,5,9]. While MRI can measure renal perfusion with or without exogenous contrast agents, non-contrast techniques may be preferable due to the risk of nephrogenic systemic fibrosis associated with gadolinium-based contrast agents in patients with renal insufficiency and in the setting of longitudinal monitoring. Nephrogenic systemic fibrosis is an irreversible systemic disease, which visibly affects the skin and joints, causing skin thickening and loss of joint range of motion. Although rare, this is a devastating disease with no effective treatment [10].

Arterial spin labeling (ASL) uses the blood as an endogenous contrast agent, allowing perfusion measurements without the administration of gadolinium. Inflowing blood is selectively labeled to have an opposite magnetization compared to the destination tissue. The difference between a labeled image (tag) and a non-labeled image (control) can be used

to calculate tissue perfusion. ASL has been used extensively for brain perfusion [11] and more recently has been applied to native and transplanted kidneys with both normal and altered function [6,8,9,12-16]. ASL studies using a flow sensitive alternating inversion recovery (FAIR-ASL) scheme, first demonstrated in the kidneys by Martirosian et al. [15], appear particularly promising, correlating kidney perfusion with renal artery stenosis grade [6] and renal plasma flow [6,16].

The goal of this study was to develop methodology to examine perfusion in both native and transplanted kidneys over a broad range of function. Perfusion was measured using a FAIR-ASL technique with respiratory triggering. In a subset of transplant subjects, results using respiratory triggering were compared to measurements obtained under free breathing to assess feasibility of the technique in sick patients unable to maintain a consistent respiratory rate.

Materials and Methods

Subjects

This study was compliant with the Health Insurance Portability and Accountability Act (HIPAA) and approved by our institutional human subjects review committee. Written informed consent was obtained from all subjects. Subjects with normal kidney function were recruited from a pool of healthy volunteers who expressed interest in participating in MRI research studies. Subjects with chronic kidney disease (CKD) and kidney transplant recipients were recruited by referring nephrologists, in a consecutive fashion, when they presented to their routine clinic appointments if they met the study's inclusion criteria. Subjects were included in the study if they were adults (>18 yrs old), MRI compatible, and clinically stable. In this study estimated glomerular filtration rate (eGFR) was used to stratify patients according to their renal function and to assess the viability of ASL perfusion in transplant and native kidneys over a broad range of function.

Between February 2008 and February 2009, MR renal perfusion exams were performed on a total of 25 subjects, including ten native subjects (six men, four women; mean age \pm standard deviation (SD), 55 ± 13 years; age range, 33-79 years) and fifteen transplant subjects (twelve men, three women; mean age \pm SD, 49 ± 14 years; age range, 21-71 years). Serum creatinine was measured immediately prior to the MRI examination and eGFR was calculated using the Modification of Diet in Renal Disease formula [17]. In order to minimize the effect of recent fluid consumption on renal perfusion, all subjects refrained from fluids for four hours before MR imaging.

Scan Protocol

Scans were performed on a 1.5 T MR scanner (Signa HDx, GE Healthcare, Milwaukee, WI, USA) with an eight-element phased array cardiac coil (GE Healthcare, Milwaukee, WI, USA). ASL images were acquired using a FAIR-balanced steady state free precession (b-SSFP) acquisition scheme [15] with a 20 ms hyperbolic secant adiabatic inversion pulse and the following readout parameters: TR/TE/flip = 4.6ms/2.3ms/70°, BW = 83.33 kHz, FOV = 34-36 cm, matrix = 128 \times 128, and slice thickness = 8 mm. In a few cases, larger subjects required an increase of the FOV, negating the use of zoom gradients and lengthening the TR/TE to 5.8/2.9 ms. A single imaging slice was positioned central to a 20 mm thick slice selective inversion slab. The inversion slab was carefully chosen not to include the feeding vessels. For native kidneys the oblique-coronal orientation allowed for proper placement, however in the transplant kidney, the oblique-sagittal orientation was needed in eight of the fifteen subjects to avoid major feeding arteries to the transplanted kidney. The inversion pulse was respiratory triggered by a breathing belt post-exhalation, and following an

inversion time (TI) of 1.2 s, a centric phase encoded b-SSFP image was acquired. The subjects were coached, prior to scanning, to breathe after completion of the image readout. Respiratory rates were 12 breaths/min or lower to allow sufficient magnetization recovery between inversions (≥ 5 s between breaths). Control and tag images were alternated until 64 total images (32 control-tag pairs) were acquired. Proton density images (to measure M_0) were obtained with a NEX = 4 using the b-SSFP readout with no inversion preceding it. Scanning was completed in 6-9 min, depending on the respiratory rate.

Free-Breathing Acquisition

ASL was performed without coaching or triggering for five transplant subjects, directly after the standard coached and triggered acquisition. Subjects were instructed to breathe freely for this scan, and data were acquired using a fixed 5 s delay between each inversion (6 min for the entire exam). The slice selective inversion thickness was increased in the free-breathing acquisitions from 20 mm to between 24-28 mm to accommodate greater expected slice motion.

Segmentation and Processing

Data were analyzed using custom scripts written in MATLAB (version 7.5, The MathWorks Inc., Cambridge, MA, USA). Rectangular regions of interest (“ROIs”) were drawn around each kidney and were registered independently through the image series by using automated rigid registration based on normalized mutual information (NMI). ROIs that could not be registered, due to either through-plane motion or deformation, were excluded from the averaged data set. In one native kidney, the M_0 image could not be aligned to the FAIR images because of a difference in kidney orientation. A single, average M_0 value was used in this instance. The mean of the registered tag images, M_T , was then subtracted from the mean of the registered control images, M_C , to obtain a difference image, ΔM , of the kidney. After manually segmenting out the kidney from the T_1 -weighted, M_C image, the kidney cortex, which had higher signal intensity than the medulla, was readily differentiated with thresholding. For a limited number of cases where the coil sensitivity varied across the kidney or the segmental vessels were visible, additional manual segmentation was necessary.

Perfusion was determined on a pixel-by-pixel basis using a one compartment model:

$$f = \frac{\lambda}{2\alpha \cdot TI} \cdot \frac{\Delta M}{M_0} \cdot \exp\left(\frac{TI}{T_1}\right)$$

where $f \equiv$ perfusion, $\lambda \equiv$ partition coefficient = 80 ml/100g[13], $\alpha \equiv$ inversion efficiency = 1.0 (assumed), $TI \equiv$ inversion time = 1.2 s, and $T_1 \equiv$ longitudinal relaxation time = 966 ms for cortex and 1,410 ms for medulla [18]. The measured ΔM and M_0 from each pixel were used to generate a perfusion map for the imaged slice of the kidney. All pixels in a specific tissue (e.g. cortex) were averaged together to provide mean perfusion. Cortical pixels with $f > 1000$ ml/min/100g and medullary pixels with $f > 300$ ml/min/100g were excluded from the mean perfusion calculation as non-physiologic outliers.

Statistical Analysis

A Wilcoxon signed rank test was used to determine if there were perfusion differences between left and right native kidneys. An intra-class correlation coefficient (ICC) was also calculated to determine agreement between left and right native kidneys. For each native subject, left and right kidney perfusion measurements were averaged for a specific tissue

(e.g. cortex) and used in the remaining statistical analyses. The correlations between the perfusion values and eGFR for both native and transplant subjects were examined using Pearson (r) and Spearman (r_s) correlation coefficients. Additionally, all subjects were divided into two groups based on healthy normal vs poor function, with healthy normal function denoted by an eGFR above 60 ml/min/1.73m² and poor function below 60 ml/min/1.73m². This threshold was selected since CKD is defined as kidney damage or a GFR below 60 ml/min/1.73m² for three months or more [19]. Cortical and medullary results for these groups are shown with dotplots. Statistical differences between native and transplant subjects in the same eGFR group were determined using the Wilcoxon rank sum test. Lastly, the Wilcoxon signed rank test was again used to test for statistical differences between the free-breathing and coached/triggered ASL measurements. All statistical analyses were performed using SAS 9.1 for windows (SAS Institute Inc. Cary, NC). Statistical significance was defined as a two-tailed p-value < 0.05.

Results

Sagittal slices representing the control, tag, difference (ΔM), and perfusion map images are displayed in Figure 1 for a transplant subject who exhibited negligible motion. In four of the kidneys (two native and two transplanted), banding artifacts due to B_0 inhomogeneity affected perfusion measurements in a small portion of the cortex (Figure 2a). The affected area was then excluded from the mean perfusion calculation. In many of the transplant subjects, major feeding vessels prevented a coronal acquisition (Figure 2b), providing reason to place the slice orientation sagittal to the kidney.

Although respiratory coaching and triggering were used in the acquisition, both native and transplanted kidneys frequently shifted position throughout the exams. Figure 3 demonstrates motion in two transplanted kidneys which manifests as blurring after averaging (Figure 3c,h). Registration significantly reduced the motion artifacts (Figure 3d,i). Perfusion measurement was not obtained in one of the 34 kidneys examined in this study because retrospective image registration could not sufficiently align the images.

Quantitative perfusion results obtained from a triggered and coached acquisition are listed in Table 1 for the native subjects and Table 2 for the transplant subjects. Perfusion values for healthy native subjects in this study ranged from 407-456 ml/min/100g with mean 427 (± 20) ml/min/100g in the cortex and 47-121 ml/min/100g with mean 85 (± 33) ml/min/100g in the medulla. These measurements agree with other renal perfusion studies performed using ASL [6,12-14] and other contrast techniques [20]. For native subjects with CKD (native subjects 6-10, Table 1), the underlying etiologies varied and included: lupus nephritis, glomerulonephritis, hypertension, and diabetes. Perfusion values measured for native right and left kidneys (i.e., right kidney perfusion – left kidney perfusion) were not statistically different for cortical (17 ± 49.37 ml/min/100g; mean \pm SD; $p = 0.43$) or medullary perfusion (-2.67 ± 16.74 ml/min/100g; $p = 0.76$). Intraclass correlation coefficients also indicated a high level of agreement between right and left kidney perfusions (cortical ICC = 0.93; medullary ICC = 0.87) which provided reason to average the right and left kidney measurements for each native subject in these analyses.

Perfusion values for the transplanted kidneys with good function (eGFR ≥ 60 ml/min/100g) ranged from 262-360 ml/min/100g with mean 314 (± 41) ml/min/100g in the cortex and 11-70 ml/min/100g with mean 37 (± 21) ml/min/100g in the medulla. Four of these five healthy transplant subjects were taking calcineurin inhibitors which may account for lower perfusion values compared to the healthy native subjects in this study. These cortical perfusion values agree with a recently published study by Lanzman et al. using a similar FAIR-ASL approach in the transplanted renal cortex [9].

Cortical perfusion correlated with eGFR in both native ($r = 0.85$, $p = 0.002$; $r_s = 0.76$, $p = 0.01$) and transplant kidneys ($r = 0.61$, $p = 0.015$; $r_s = 0.57$, $p = 0.03$), while medullary perfusion did not show significant correlation with eGFR in either group. Groupwise comparison of perfusion separated by renal type (native vs transplant) and function based on eGFR is displayed for cortical and medullary perfusion in tables 3 and 4 respectively. Cortical perfusion was greater in native vs transplant subjects with $eGFR \geq 60$ ml/min/1.73m² ($p < 0.01$; Figure 4a). Medullary perfusion was greater in native vs transplant subjects for both $eGFR < 60$ ml/min/1.73m² ($p = 0.04$) and $eGFR \geq 60$ ml/min/1.73m² ($p = 0.04$; Figure 4b).

Results from the substudy comparing free-breathing and the standard respiratory triggered/coached acquisition methods are displayed in Figures 5 and 6. Figure 5 demonstrates two similar perfusion maps for the same transplanted kidney. Data in 5a were acquired with the standard respiratory coaching and triggering while those in 5b were obtained while the subject breathed freely. A comparison of all available free-breathing and triggered/coached data can be seen in Figure 6 for both cortical (a) and medullary flow (b). Although no statistical differences were observed, ASL values obtained with a free-breathing acquisition were lower on average than the triggered and coached values with a mean difference of (mean \pm SD) -37 ± 51 ml/min/100g in the cortex and -17 ± 18 ml/min/100g in the medulla.

Discussion

Functional MR imaging for the native and transplanted kidney can be performed without injecting a contrast agent and may be helpful in the longitudinal assessment of kidney function in patients with renal disease. Before these methods will be clinically useful, applicable techniques must be developed. Our group has implemented a FAIR-ASL technique to measure perfusion in both native and transplanted kidneys over a broad range of function. Specifically we chose a sagittal orientation in many of the transplanted kidneys to avoid major feeding vessels, utilized a respiratory triggered/coached breathing acquisition, and performed automatic image registration using NMI. We were able to measure the cortical and medullary perfusion in all but 1 of 35 kidneys with this technique and found correlations between cortical perfusion and eGFR. Groupwise comparisons indicate cortical perfusion differences between native and transplant subjects with $eGFR > 60$ ml/min/1.73m² and medullary perfusion differences for both $eGFR > 60$ and $eGFR < 60$ ml/min/1.73m². We also successfully implemented a free breathing protocol in a subset of transplanted kidneys to assess robustness of the technique in patients who may not be able to breathe at a consistent rate or follow respiratory coaching (i.e. very ill patients).

While perfusion values in this study are similar to other renal ASL work [6,12-14], cortical perfusion values for healthy native subjects in this study are slightly higher than values reported by Fenchel et al. using a similar pulse sequence [6]. This may partly be due to variation in acquisition and processing. As for acquisition differences, the present study allowed two additional seconds (for a total of 5 s) for magnetization recovery between inversions. This is more representative of the compartment modeling, which assumes complete recovery and should provide a larger difference between tag and control, increasing the perfusion value. As for processing differences that would lead to higher perfusion values, a more inclusive threshold for cortical perfusion of 1000 ml/min/100g (only $< 2\%$ of pixels were rejected) was used compared to 600 ml/min/100g used in the Fenchel et al. study.

Prospective respiratory coaching and triggering was applied to minimize motion artifacts, but it proved necessary to use post-acquisition image registration in most cases and this was critical in certain native and transplant subjects. Registering the M_C and M_T images

eliminated blurring, allowing better tissue segmentation and perfusion measurement. Without registration, the perfusion-weighted difference image would be blurred throughout the kidney as well as at the kidney border, where blurring could lead to artificially low cortical perfusion values. Retrospective respiratory sorting was an alternative, as it has been shown to reduce motion artifacts in renal ASL imaging compared to a simple coached acquisition (without respiratory triggering) [14]. With a FAIR acquisition, the inversion slab must encompass the imaging slice so prospective respiratory triggering was used in addition to coaching for this study.

The oblique-sagittal orientation was found to be more practical for transplant subjects whose feeding vessel trajectories precluded the possibility of a coronal acquisition. For example, in subjects with native kidneys, an oblique slice could be positioned both coronal to the kidneys and posterior to the renal arteries allowing coverage of both kidneys with one FOV. However, in many transplanted kidneys, a coronal slice orientation placed the feeding vasculature in-plane. In these cases, a coronal acquisition would have led to artificially low perfusion measurements by slice-selectively inverting the inflowing blood spins which are presumed to be at equilibrium magnetization. An oblique-sagittal slice omitted major feeding vessels in these transplant subjects and allowed for sufficient coverage of the transplanted kidney (Figure 1). Slice orientation would not be restricted for other ASL techniques that label the blood upstream, such as proximal inversion with a control for off-resonance effects (PICORE) or pulsed-continuous methods. Unfortunately, labeling via these methods would lengthen the transit time of the tagged blood and increase the variability in arrival time from subject to subject. Since diseased kidneys are likely to demonstrate more variable flow, adjusting the delay time is also a source of potential error that is avoided by use of the FAIR technique.

The respiratory triggered and coached perfusion measurements in this investigation were comparable to a study by Lanzman et al. [9] which used a free-breathing FAIR-ASL approach in transplanted kidneys. The free-breathing, sagittal perfusion measurements in this study were also similar to respiratory triggered and coached measurements, although slightly lower. Thus free-breathing may be a reasonable option for patients who are unable to follow the regimented breathing pattern. The free-breathing perfusion measurements may be lower because the slice selective inversion thickness was larger than the standard 20 mm to increase the tolerance for through-plane motion. Consequently, more blood spins outside of the imaging slice were inverted when they flowed into the imaging plane.

As expected there was a positive correlation between cortical perfusion measurements and eGFR in both transplant and native kidneys supporting the regulation of glomerular filtration rate by renal blood flow. However, perfusion values were significantly reduced in transplanted kidneys compared to native kidneys for subjects with eGFR > 60 ml/min/1.73m² (Figure 4). This may result from differential regulation of blood flow in transplanted kidneys or the vasoconstrictive effects of calcineurin inhibitors that are commonly used in kidney transplantation to prevent rejection [21-25] and is an area of future study.

Several limitations apply to the current study. No gold-standard was available to compare the perfusion results, however other studies using FAIR-ASL in the kidneys have shown good correlation with standard references [6,16,26]. The results in this work were obtained from a limited number of native and transplant subjects and must be extended to a larger population before drawing firm conclusions regarding practicality and clinical significance. This study was performed on a 1.5T system which has limited SNR compared to a 3T system but should have fewer banding artifacts. Banding was visible in four of the kidneys, although only a small portion of the kidney body was affected, and unaffected measurements were obtained for the majority of the kidney. Higher order shimming or

adjusting the center frequency of the radiofrequency pulse would likely shift these artifacts outside of the kidney [27]. This FAIR ASL sequence did not apply bi-polar gradients before readout to suppress signal from tagged spins in the arterioles. This may have introduced a bias towards higher perfusion measurements as well.

Perfusion values presented in this study were determined using a one compartment model which requires many assumptions, such as rapid water exchange between the intravascular and extravascular space. A two compartment model [28] would reduce those assumptions and allow more accurate perfusion quantification, however it requires measurements at multiple delay times which is not possible to achieve in a clinically feasible scan time. Further assumptions used in this study for perfusion measurement include a constant tissue T_1 and inversion efficiency, α . In reality, the α can vary depending on rf-amplifier performance, stability and pulse design, while the T_1 is a patient-specific parameter that may vary regionally and with kidney disease and setting. The cortical T_1 assumed in this study may have introduced a bias towards higher perfusion for lower functioning kidneys because cortical T_1 can increase with renal insufficiency [29]. Incorporating a patient-specific measured T_1 and pulse-specific α into the model should increase the quantitative accuracy of these measurements with the trade-off of slightly increased scanning time.

In conclusion, FAIR-ASL was able to measure renal perfusion in subjects with native and transplanted kidneys, potentially providing a clinically viable technique for monitoring kidney function. Results from this study demonstrate that medullary perfusion was systematically lower in transplant vs native kidneys, and that cortical perfusion was lower for transplanted kidneys as well for kidneys with higher function, $eGFR > 60 \text{ ml/min/1.73m}^2$. Cortical perfusion correlated with $eGFR$ in both native and transplanted kidneys. Future work will compare FAIR-ASL with microsphere perfusion in a swine model, assess reproducibility in human subjects, and measure the ability of this technique to characterize renal disease and provide earlier detection of functional change.

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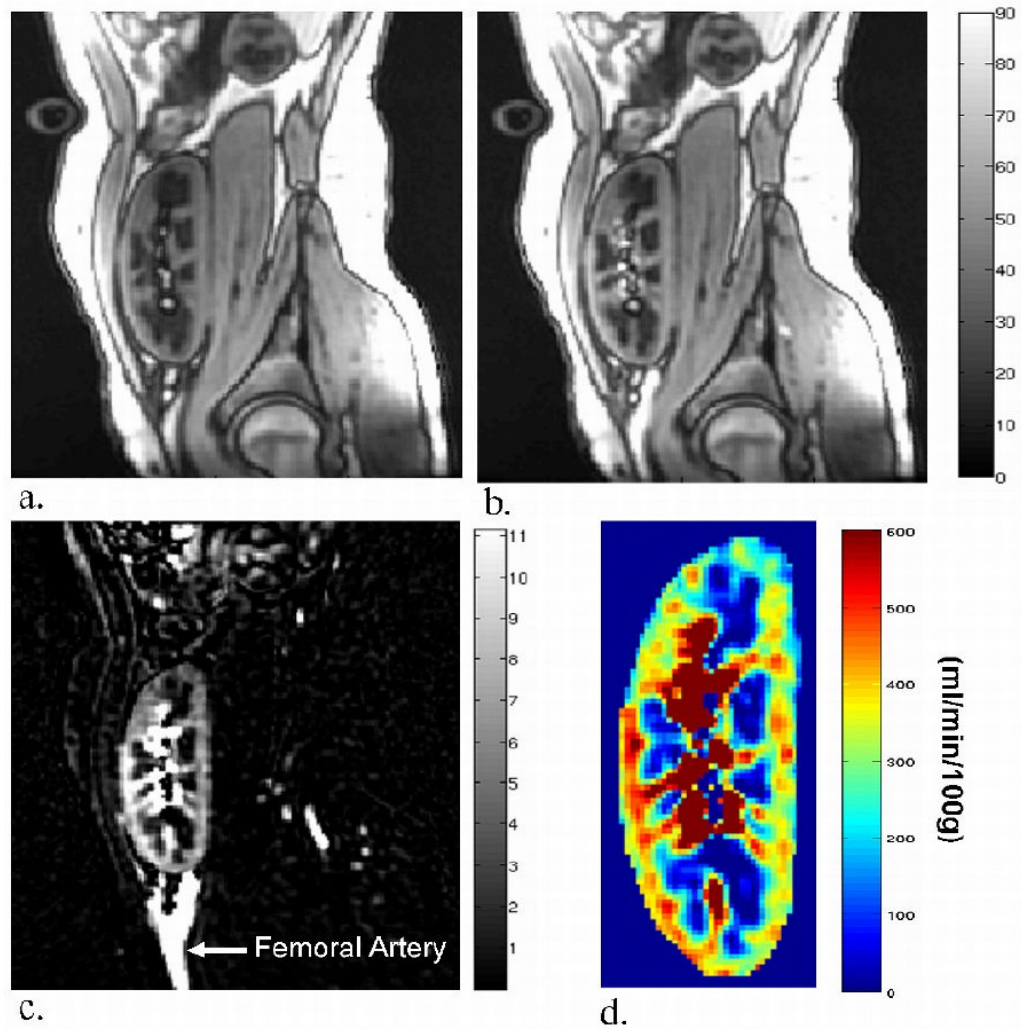


Figure 1.

Tag (a), control (b), and difference, (c) images acquired in a sagittal plane for a healthy transplant kidney subject (no. 1 in Table 2) with negligible motion (eGFR = 74 ml/min/1.73m²) along with the resulting perfusion map (d) shown in units of ml/min/100g.

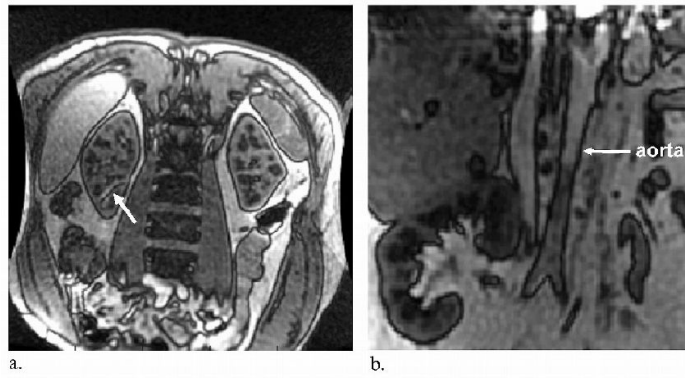


Figure 2.

(a) Worst case example of b-SSFP banding artifact inside the kidney for a healthy native kidney subject. The nulled banding area (white arrow) is excluded from mean cortical perfusion calculations. **(b)** Intersection of the imaging slice with major feeding vessels, such as the aorta in this example (white arrow), prevented a coronal acquisition in many of the transplant subjects. In these cases, a coronal acquisition would have caused the slice selective inversion to invert the inflowing blood spins which are presumed to be at equilibrium magnetization upon entrance to the kidney.

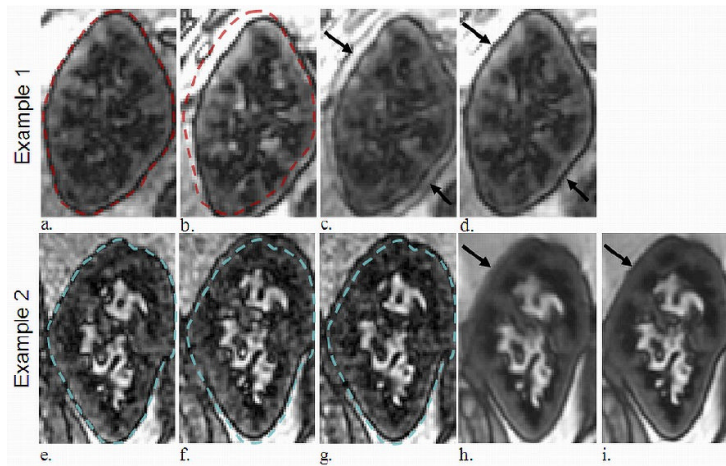


Figure 3. Two different transplanted kidneys which demonstrated significant motion during the ASL exam. Example 1 displays translation due to respiratory motion (dotted outline in **a-b**). The average of the two images pre-registration demonstrated motion artifact at the boundary of the kidney body and blurring (**arrow in c**), while averaging post-registration mitigated these (**arrow in d**). Example 2 displays another transplanted kidney in three different positions (**e-g**). The average of all 64 FAIR images pre-registration demonstrated blurring (**arrow in h**) which the post-registration average reduced (**arrow in i**).

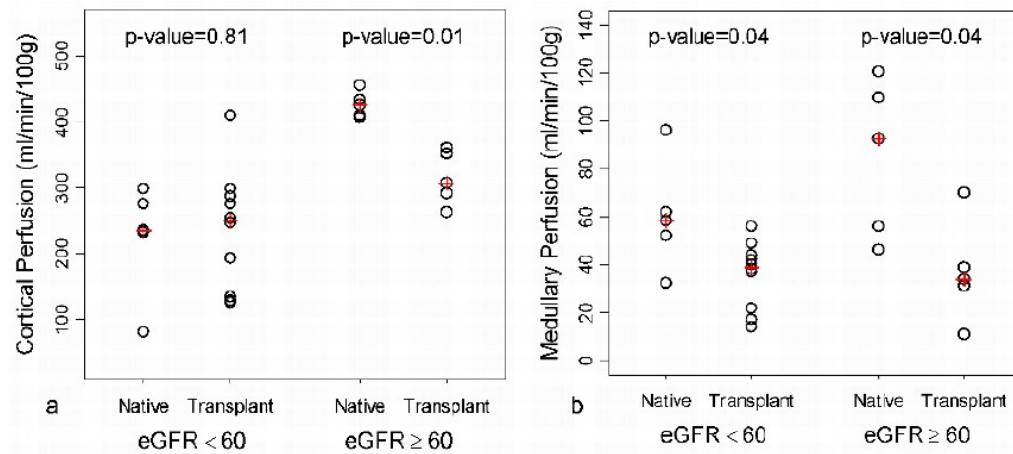


Figure 4.

Groupwise comparison of native and transplant cortical (a) and medullary (b) perfusion separated into two groups based on an eGFR threshold of 60 ml/min/1.73m². For subjects with eGFR ≥ 60 ml/min/1.73m², a statistical difference between native and transplant is observed in cortical (p = 0.01) and medullary (p = 0.04) perfusion. A statistical difference in medullary perfusion (p = 0.04) was also observed in subjects with eGFR < 60 ml/min/1.73m². (+: median)

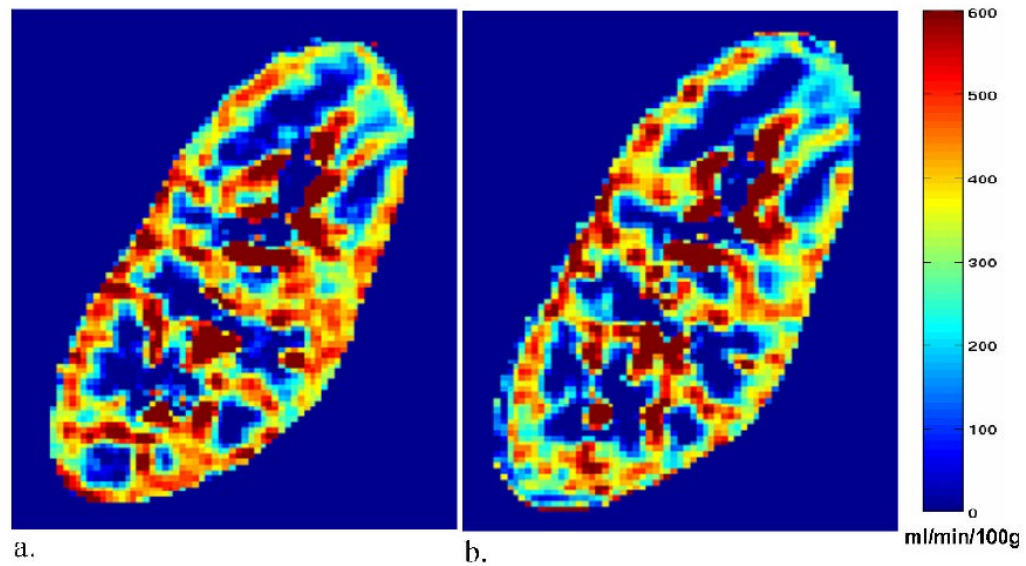


Figure 5.

Sagittal perfusion maps for a transplant subject (no. 5 in Table 2) with an eGFR of 60 ml/min/1.73m² acquired with the standard respiratory triggering and coaching (**a**) and acquired during free-breathing (no respiratory triggering or coaching) (**b**). Mean cortical perfusion under respiratory triggering/coaching and free-breathing was **a**) 382 ml/min/100g and **b**) 344 ml/min/100g, respectively. Mean medullary perfusion was **a**) 36 ml/min/100g and **b**) 13 ml/min/100g, respectively.

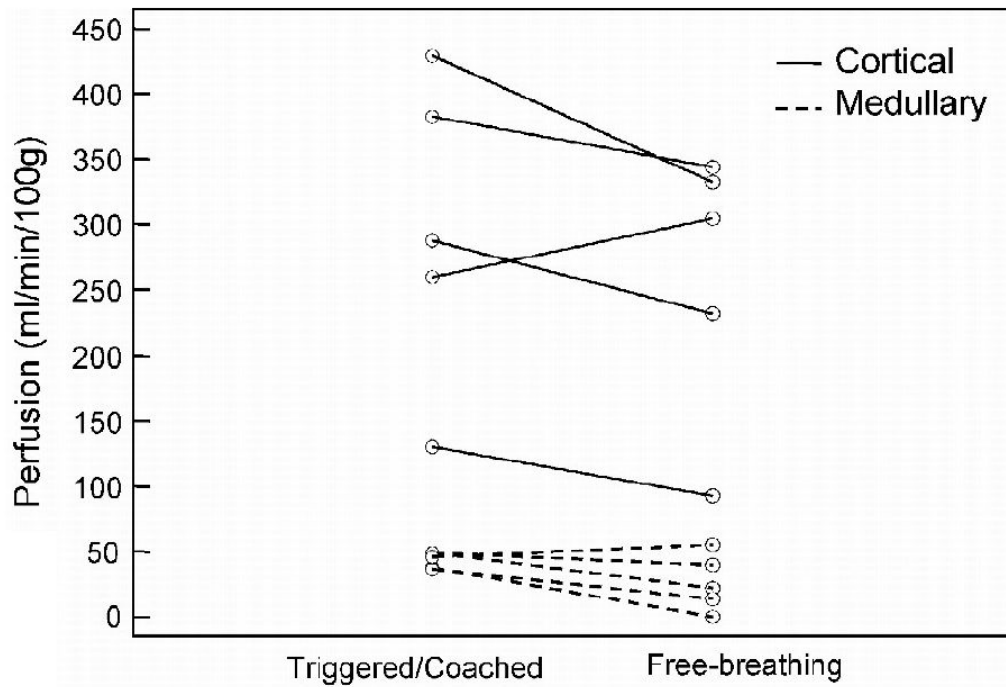


Figure 6. Comparison of respiratory triggered/coached and free-breathing perfusion measurements in five transplant subjects. All exams were performed with an oblique-sagittal orientation. Data trend toward lower perfusion values under free-breathing.

Table 1
Age, sex, eGFR and respiratory triggered/coached perfusion results for the ten subjects with native kidneys

Native Subject	Sex	Age	eGFR*	Mean Perfusion [†]	
				Cortical [‡]	Medullary [‡]
1 ^H	F	33	88	426	110
2 ^H	F	52	80	410	47
3 ^H	M	41	78	456	93
4 ^H	F	65	77	407	56
5 ^H	M	53	67	435	121
6	M	45	50	236	62
7	M	79	42	233	52
8	M	60	30	276	59
9	F	60	23	298 [§]	96 [§]
10	M	64	19	81	33

* eGFR cited in ml/min/1.73m².

[†] data represents mean of the right and left kidney.

[‡] cortical and medullary perfusion values are listed in ml/min/100g.

^H denotes a healthy subject with no renal disease per their medical history.

[§] measured only for left kidney due to a shift in the slice position of the right kidney during the exam.

Table 2
Age, sex, eGFR and respiratory triggered/coached perfusion results for the fifteen subjects with transplanted kidneys

Transplant Subject	Sex	Age	Calcineurin Inhibitors	eGFR*	Mean Perfusion [†]	
					Cortical	Medullary
1	M	21	Yes	74	360	70
2	M	34	Yes	67	262	39
3	M	38	No	66	291	31
4	M	55	Yes	61	307	34
5	F	61	Yes	60	352	11
6	M	66	No	54	276	22
7	M	64	Yes	54	194	49
8	M	56	Yes	51	247	44
9	F	54	No	50	409	42
10	M	54	Yes	48	287	56
11	M	71	No	46	134	38
12	F	41	Yes	41	299	40
13	M	30	No	22	254	37
14	M	41	Yes	21	125	14
15	M	44	No	18	128	17

* eGFR cited in ml/min/1.73m².

[†] perfusion values are listed in ml/min/100g.

Table 3

Groupwise Comparison of Cortical Perfusion

eGFR*	Group	N	Mean	Median	Std Dev	Minimum	Maximum	P-value
<60	Native	5	225	236	85	81	298	0.81
	Transplant	10	235	251	91	125	409	
≥60	Native	5	427	426	20	407	456	0.01
	Transplant	5	314	307	41	262	360	

Note – perfusion data listed in Mean, Median, Std Dev, Minimum, and Maximum are all reported in ml/min/100g.

* eGFR cited in ml/min/1.73m².

P-value < .05 indicates a significant difference.

Table 4

Groupwise Comparison for Medullary Perfusion

eGFR*	Group	N	Mean	Median	Std Dev	Minimum	Maximum	P-value
<60	Native	5	60	59	23	33	96	0.04
	Transplant	10	36	39	14	14	56	
≥60	Native	5	85	93	33	47	121	0.04
	Transplant	5	37	34	21	11	70	

Note – perfusion data listed in Mean, Median, Std Dev, Minimum, and Maximum are all reported in ml/min/100g.

* eGFR cited in ml/min/1.73m².

P-value < .05 indicates a significant difference.