Children with Kidney Disease Show Blood Flow Changes in Brain

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OAK BROOK, Ill. — Blood flow changes in the brains of children, adolescents and young adults with chronic kidney disease may explain why many face a higher risk of cognitive impairment, according to a study published online in the journal Radiology.

Prior research has linked chronic kidney disease, a condition characterized by the loss of kidney function over time, with lesions in the brain’s signal-carrying white matter and deficits in cognitive performance. While chronic kidney disease in adults is frequently associated with age-related disorders such as hypertension and diabetes, the disease in childhood often occurs congenitally, yet still affects brain development and cognitive function.

Dr. Detre and colleagues assessed blood flow in the brains of 73 pediatric kidney disease patients, average age just under 16 years, and 57 similarly aged control participants. The researchers used arterial spin labeling, an MRI technique that can noninvasively quantify blood flow in the brain.

Patients with kidney disease showed higher cerebral blood flow compared with controls in certain brain regions — a surprising finding, considering that decreased cognitive performance is generally associated with decreased blood flow in the brain, such as in aging and dementia. There are a couple of possible reasons for this unusual phenomenon, Dr. Detre said.

“It may indicate compensatory hyperactivity, in which the brain regions are working extra hard to maintain performance,” he said. “Another possibility is that there’s a disturbance in the regulation of blood flow in these patients.”

White matter cerebral blood flow and blood pressure were also correlated, suggesting that kidney disease patients have problems with cerebrovascular autoregulation, the process that controls blood pressure in the brain. This type of dysfunction could potentially lead to white matter injury, according to Dr. Detre.

“Chronic kidney disease appears to affect brain physiology and function even early in the disease,” he said. “This study gives us clues about what changes in brain physiology might underlie cognitive changes.”

Among those changes were differences in blood flow between patients and controls in areas of the brain that correlated with cognitive problems in the patients. Compared with controls, kidney disease patients had cerebral blood flow differences in the default-mode network, the network of brain regions active when a person is not focused on a particular task. Patients with low executive function, or skills related to planning, organizing and paying attention, had significant differences in cerebral blood flow compared with controls.

The findings point to cerebral blood flow measurements with arterial spin labeling as a potentially valuable tool in characterizing cerebrovascular function in chronic kidney disease — an important area of research given the associations between kidney disease and neurological function, and the significantly increased risk for transient ischemic attack and stroke in even mild chronic kidney patients.

“Cerebral blood flow is a critically important physiological parameter that you can measure in just a few minutes with arterial spin labeling,” Dr. Detre said. “This technique provides a noninvasive way of quantifying cerebral blood flow that doesn’t require use of contrast agent, which is contraindicated in patients with kidney dysfunction.”

“Regional Cerebral Blood Flow in Children and Young Adults with Chronic Kidney Disease,” Collaborating with Dr. Detre were Hua-Shan Liu, Ph.D., Erum A. Hartung, M.D., Abbas F. Jawad, Ph.D., Jeffrey B. Ware, M.D., Nina Laney, B.A., Allison M. Port, B.A., Ruben C. Gur, Ph.D., Stephen R. Hooper, Ph.D., Jerilynn Radcliffe, Ph.D., and Susan L. Furth, M.D., Ph.D.

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Figure 1. Contrast shown demonstrates regions where CBF in patients with chronic kidney disease (CKD) is greater than that in the control group. There were no regions where the control group showed greater CBF than patients with CKD.

High-res (TIF) version
(Right-click and Save As)
Figure 2. Images show overlapped clusters from all individual patients with chronic kidney disease with positive extrema in cerebral blood flow (CBF) in subject-specific voxel-wise analysis. Color bar indicates total number of subjects who had subject-specific clusters with increased CBF.

High-res (TIF) version
(Right-click and Save As)
Figure 3. Scatter plot shows partial residual values of precuneus cerebral blood flow (CBF) and executive function in patients with chronic kidney disease with presence of positive extrema CBF, indicating significant correlation between precuneus CBF and executive function after controlling for hematocrit, age and sex. TNB = traditional neurocognitive battery.