

Tuesday, October 19

in the untreated eye. These preliminary results suggest that the peripheral autonomic system in AD exhibits an abnormal cholinergic response and may provide a relatively noninvasive diagnostic test for AD.

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tw.  
caused by asy...  
the first twin pair to be...  
address this question, we sought ev...  
the unaffected twin using magnetic resonance imaging (MRI),  
18-F-fluorodeoxyglucose and 15-O-water positron emission  
tomography (PET), and neuropsychological tests. MRI was  
normal in the unaffected twin, but showed widespread atro-  
phy in the proband that was most severe in the right parietal  
lobe. In the unaffected twin, PET disclosed focal hypometab-  
olism in the left inferior parietal lobe, and despite normal  
language skills, neuropsychological testing showed signifi-  
cantly lower scores on verbally than nonverbally mediated  
tests. In the proband there was extensive hypometabolism of  
posterior cortices bilaterally, which was worse on the right,  
and neuropsychological scores, though lower on all tests,  
were higher on verbally than nonverbally mediated tests. We  
conclude that the clinically unaffected twin may have an early  
stage of ACD with a different laterality than his brother,  
which suggests that there are both genetic and nongenetic  
contributors to topographical patterns of ACD.

P192. Regional Brain-Behavior Correlations Using  
Magnetic Resonance Volumetric Measures in  
Alzheimer's Disease

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Volume loss in some brain tissue compartments has recently  
been found by volumetric magnetic resonance (MR) analysis  
in Alzheimer's disease, but the relationship between regional  
volume loss and neuropsychological performance remains  
unclear. Ten subjects with probable Alzheimer's disease and  
10 age- and education-matched controls underwent T2-  
weighted, high-resolution MR imaging (1.5 tesla). On bifea-  
ture segmentation of brain compartments, interrater reliabil-  
ity was 90%. Hippocampal volumes were measured on  
three-dimensional T1-weighted images. Brain measures were  
correlated with detailed neuropsychological testing. We con-  
firmed a significant increase in overall CSF volume in Alzhei-  
mer's subjects, as well as in frontal, temporal, and parietal  
regions ( $p < 0.002$ ). Multivariate models using age, years  
of education, and global measures of cognitive function  
showed significant effects of regional CSF volumes on corre-  
sponding neuropsychological tests. For example, left parietal  
atrophy accounted for 26% of the variance on semantic flu-  
ency ( $p < 0.02$ ), left hippocampal volume for 24% of the  
variance on verbal learning ( $p < 0.04$ ), and frontal atrophy  
for 33% of the variance on the Wisconsin card sort test ( $p$   
 $< 0.01$ ). A modulating effect of premorbid intelligence was  
seen in relating global atrophy to overall cognitive perfor-

mance. We conclude that this is a promising technique for  
longitudinal quantitation and for assessing biological benefits  
of drug interventions.

P193. Immunohistochemical Study of Copper-Zinc  
and Manganese Superoxide Dismutases  
in Senile Plaques

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Tokyo and Yamanashi, Japan

To investigate the role of the oxygen-free radical in senile  
plaque formation, we made an immunohistochemical study  
in brains of patients with dementia of the Alzheimer type  
utilizing antibodies to copper-zinc and manganese superoxide  
dismutases (CuZn-SOD, Mn-SOD). These enzymes scav-  
enge superoxide anion. CuZn-SOD is localized mainly in  
cytosol and Mn-SOD is localized in mitochondria matrix.  
The material consisted of autopsied brains from 6 patients  
with dementia of the Alzheimer type (average age, 77.0 yr;  
range, 62-89 yr). Buffered 10% formalin-fixed, paraffin-  
embedded 6- $\mu$ m-thick sections of hippocampus from all  
cases were immunostained with rabbit polyclonal antibodies  
to human CuZn-SOD and Mn-SOD with the ABC method.  
Most dystrophic neurites of primitive and classic plaques  
were immunoreactive for CuZn-SOD as fine positive struc-  
tures. Reactive astrocytes surrounding senile plaques had  
granular structures positive for Mn-SOD in their cell bodies  
and proximal processes. Diffuse or burned-out plaques had  
no immunoreactivity for either SOD. Results of this study  
suggest that neurons and astrocytes may have different re-  
sponses to oxidative stress during senile plaque formation.

P194. Neuropsychological Correlates of Competency  
Loss in Alzheimer-Type Dementia

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Our objective was to reveal the neuropsychological changes  
associated with competency loss in Alzheimer's disease (AD).  
Little is known about the cognitive changes that are associ-  
ated with loss of competency in AD and other dementias.  
Identification of neuropsychological criteria will assist physi-  
cian decision makers who currently lack objective measures  
for competency assessment. The investigators developed 2  
clinical vignettes that reliably and validly test subject compe-  
tency (treatment decision capacity) under 5 accepted and in-  
creasingly stringent legal standards. Thirty subjects (10 nor-  
mal elderly and 20 AD patients) were administered the  
vignettes, as well as a battery of neuropsychological measures  
theoretically linked to competency function. Neuropsycholo-  
gical test scores for the 2 patient groups were correlated  
with scores on the legal standards. AD patients performed  
significantly below normal controls on the more stringent  
legal standards and on all neuropsychological measures.  
Scores by AD patients on the 5 legal standards strongly cor-  
related ( $r > 0.60$ ,  $p < 0.02$ ) with neuropsychological mea-  
sures of verbal conceptualization, confrontation naming, au-  
ditory comprehension, short-term verbal memory, and verbal  
fluency. Identical correlational analyses using the normal  
older controls revealed virtually no significant correlations.  
The study revealed that neuropsychological measures of ver-  
bal conceptualization, auditory comprehension, verbal fluency,  
and short-term memory were strongly associated with the  
impaired performance of AD subjects on different stan-  
dards of competency.

(57.) \* Foster J., Black S., Stanchev P., Buck B.,  
Moscovich M., Winocur G., Bronskill M.,  
"Regional Brain - Behaviour Correlations Using  
Magnetic Resonance Volumetric Measures in  
Alzheimer's Disease", *Annals of Neurology*  
**1993**, No. 34, 293.