**“Sick” MRI syndrome**

**M J Kushner**

At this time, MRI is a better indicator of tissue outcome rather than clinical destiny

The report by Marx et al (this issue, pp 572–575) joins the ever growing literature dealing with multiparametric MRI studies of tissue characteristics in stroke. Markers of ischaemic perturbations of cell homeostasis and perfusion, including changes in diffusion and perfusion weighted imaging (DWI and PWI), seem now to allow human stroke studies more analogous to animal models. Although debate rages, current optimism maintains that MRI delineates the three main tissue compartments in ischaemia. Conceptually these compartments are (a) irreversibly damaged tissue, (b) minimally disrupted tissue destined for eventual recovery, and (c) the “penumbra” or “borderzone” or “at-risk” region where outcome hangs in the balance between irrevocable damage and some degree of recovery, and which may be amenable to treatment.

Recent human data suggest that DWI and DWI-PWI mismatches strongly predict tissue outcome after ischaemia. Despite this progress in neuroimaging, clinical and clinicopathological MRI correlation has lagged behind. Reconciling recent MRI studies of stroke with each other is not straightforward, and it is also daunting to relate them to previous investigations of stroke using other reliable structural and haemometabolic techniques. To date, human MRI studies have used many different paradigms, and this situation stands in contrast with standardised models found in the animal stroke literature. Variations in clinical criteria, imaging protocols, and myriad other factors introduce obvious confounding influences in these studies, and it should be assumed that lack of standardised approaches can only increase exposure to not so obvious bias.

Marx et al report a relatively high incidence of DWI changes following posterior circulation ischaemia where the clinical syndrome actually was transient or reversible. DWI alterations tended to be more prominent with longer lasting ischaemic symptoms yet were detected despite ultimate clinical recovery. This bears mention, indeed, for one would expect the observed DWI changes to bode ill when in actuality full recovery was the case. This represents a potential quandary: how should one reconcile a “sick” MRI with a healthy patient? A simple explanation may be that, at this time, MRI is a better indicator of tissue outcome rather than clinical destiny. If future reports invoke MRI as an indicator of clinical outcome, or as a marker of the impact of therapeutic intervention, then data will be needed across the full spectrum of stroke severity from fixed deficits to transient syndromes. Standardised study designs are needed so that the true utility of functional MRI is established in both natural history and interventional studies of stroke. Consensus on this score would be useful now, for the widespread availability of MRI, in contrast with other techniques such as PET or SPECT, suggests that this literature will continue to proliferate.

While this field remains in flux, clinical neurology should take note. At a minimum, these MRI markers of ischaemia must be regarded as new harbingers in the menagerie threatening haemometabolic risk, and attention must be paid. Their appearance should spur the clinician to redouble efforts to apply available medical and surgical treatments, whether they be acute, or preemptive and prophylactic, to address treatable haemorheological and haemodynamic abnormalities.

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**Stroke risk**

**White matter change and stroke risk**

**P M Matthews**

Large cohort studies have defined an increasing range of therapeutic interventions that can be applied to reduce stroke risk. However, application of these in a specialist practice is complicated by the problem of interindividual variation. A major practical issue is to differentiate those patients who will do well on standard treatment from those for whom a specifically targeted, more aggressive approach is needed. Ideally, it would be possible to profile patients at presentation to identify those at highest risk. If such a strategy could be made reliable, it would help simultaneously to balance a commitment to provide the best care available for each patient and to reassure government and other funders that potentially expensive therapeutic interventions are being employed in the most cost-effective fashion.

MRI provides an attractive tool for stratifying risk in a stroke population. It offers a good definition of brain injury and promises to deliver increasingly quantitative markers of specific pathological changes. With growing access to MRI scanners and routine imaging of stroke cases, even this relatively expensive “screening test” would be of potential...
Dementia

Qualitative performance characteristics differentiate dementia with Lewy bodies and Alzheimer’s disease

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New approach to assessment of patients with dementia with Lewy bodies is clinically useful in differentiating them from patients with Alzheimer’s disease

Accurate and early clinical diagnosis of dementia with Lewy bodies (DLB) is essential for optimal management so that pitfalls such as severe neuroleptic sensitivity can be avoided, a trial of cholinesterase treatment can be instigated, and parkinsonian symptoms can be appropriately treated, weighing up the balance of psychiatric and motor symptoms and fall risk. In practice it has often been difficult to differentiate DLB from other forms of dementia, particularly Alzheimer’s disease. Several previous studies have suggested that DLB and Alzheimer’s disease can be differentiated on the basis of visual and visuospatial tasks. The tasks most easily applicable to initial diagnosis are drawing tests (such as the clock drawing). However, this approach ignores what is one of the more striking features of DLB: fluctuating cognition. Operationalised criteria for the clinical diagnosis of DLB (which do consider a broad range of symptoms) have achieved high levels of specificity (> 90%); in many studies sensitivity has been as low as 50%. Probably the major obstacle to accurate diagnosis has been the difficulty of identifying fluctuating cognition; for example, two inter-rater reliability studies have indicated that agreement

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between expert clinicians is barely better than chance.1 11

A body of work focusing on these issues has validated two clinical rating scales11 and indicated that fluctuation of reaction time on attentional tasks11 and variability in spectral frequency assessed by electroencephalography11 can also be useful methods for identifying and quantifying fluctuating cognition, which seems to be closely linked to impairments of consciousness.

In the paper by Doubleday et al11 (this issue, pp602–607) tackles these issues in an innovative way. The authors developed a standardised qualitative instrument based on observations of the patient during the course of the testing session. Information is obtained regarding a number of helpful domains including general inattention, distraction, and intrusions. Of particular interest, general inattention, distractibility, and environmentally cued intrusions were all significantly more common in DLB than in Alzheimer’s disease, the latter occurring only in patients with DB. In addition to the potential diagnostic value, this may be helpful in facilitating an understanding of the types of processing errors that are prominent in these patients. Perseveration and mental set shifting were also significantly more impaired in patients with DB and they were more likely to confabulate.

This assessment method has been used as part of the standard diagnostic protocol that has achieved a high level of accuracy in 200 people with dementia at postmortem examination, although it is unclear how many of these people had pathologically confirmed DLB. Further work is needed to examine the inter-rater reliability and concurrent validity of the instrument, and comparisons between DBL and other dementias would also be helpful. In particular, patients with vascular dementia often experience some degree of fluctuating cognition and may be more prone to difficulties with set shifting and perseveration. Diagnostic discrimination between DLB and vascular dementia can be difficult and it would be useful to know whether the proposed assessment method can also contribute to this diagnostic issue. Therefore, although the method needs to be developed a little more, the work presents a new and useful approach that is very clinically applicable and that may be of considerable value in the assessment of patients with DLB.

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