Validity of language lateralisation by unilateral intracarotid Wada test

The case report by Loddenkemper et al. is most important as it poses the question of whether unilateral Wada testing is sufficient for language lateralisation. It is certainly possible that similar complications resulting from epilepsy surgery based on Wada tests are underreported. The implications of erroneous language lateralisation by unilateral amobarbital injection, which is increasingly used in several epilepsy centres including our own, would be far reaching.

There are, however, some caveats regarding the interpretation of the Wada test results reported. The first two refer to the assessment of cortical inactivation in this patient with pre-existing damage to the motor cortex; the third refers to the extent of brain areas inactivated by the intracarotid Wada procedure. (1) Little is known about the relative susceptibility of the effect of amobarbital on language areas compared with damaged motor cortex. Thus it may be that a low dose of amytal (which 100 mg is) does impair a partially damaged motor cortex whereas language areas still maintain sufficient function to allow the tasks tested during the Wada procedure. In our experience, higher doses (140 to 200 mg) are often needed in adults to induce a complete contralateral hemiplegia. In order to avoid a possible risk of incomplete inactivation of the middle cerebral artery (MCA) territory and of language areas in particular, it may be advantageous to choose higher injection dosages.

(2) There is a striking discrepancy between well preserved proximal arm power (4/5) after hemispherectomy as compared with complete proximal hemiplegia (0/5) reported during the Wada procedure which requires additional explanation. This may indicate that the lack of voluntary movement observed during the Wada procedure does not result from a complete inactivation of the contralateral motor cortex but may be related to contralateral motor neglect.

(3) With regard to clinical history and magnetic resonance imaging, there is some indication that this patient may have had reorganisation of the language areas following the initial brain injury at the age of five years. Whereas the Wada test was interpreted as providing evidence for an interhemispheric shift to the right hemisphere, an intrahemispheric transfer of expressive functions to posterior temporal areas may have taken place. As the vascular supply of this region often is not restricted to MCA branches, incomplete inactivation of this area may occur. The results of the procedure could thus be explained by the incomplete inactivation of posterior left hemispheric language areas. It would be interesting to know if the authors co-injected a SPECT tracer in order to define the territory perfused by amobarbital.

Authors’ reply

We appreciate the above comments and are delighted that our case has sparked a discussion about the validity of the Wada test. The intention of our report was to question the use of the (unilateral) intracarotid amobarbital test (IAT) in selected cases, and we are pleased that Dr Schulz-Bonhage and Dr Quiske recognise the potential importance of this case. We would like to address their comments in detail.

Comment 1: No clinical data are available on the effect of amobarbital on the damaged motor cortex. Thus the effectiveness of an initial high dose of amobarbital, as well as incomplete inactivation of different MCA areas, remains an interesting speculation. Although we do not know whether higher amobarbital doses are more effective in cases like ours, we know that higher doses may increase the probability of drug related side effects such as encephalopathy and impaired consciousness, particularly in patients with pre-existing left hemispheric lesions. Additionally, amobarbital may have a stronger effect in older patients, supporting the use of a lower initial dose in our 55 year old patient. Furthermore, the recent replacement of amobarbital by methohexital, owing to a shortage of amobarbital in many centres, may subject patients to a dose dependent increase in seizure frequency during Wada testing when there is a high initial dose of methohexital. Clinical examination of the patient remains the most important factor in the careful titration of the amobarbital dose, in order to prevent adverse effects during its administration.

Comment 2: We thank the authors for highlighting the potentially interesting finding of the acute worsening of unilateral motor function during the Wada test. As documented, we would like to point out that the proximal strength of 4–5 was only achieved after eight months, following vigorous rehabilitation. The discrepancy from the results during the Wada test arises because Dr Schulz-Bonhage and Dr Quiske are comparing the examination during the acute amobarbital effect with chronic findings eight months after recovery from hemispherectomy. The patient was completely hemiplegic immediately after hemispherectomy and then gradually regained the reported outcome strength of 4–5. Thus the amobarbital test mimicked the immediate postoperative outcome well. Proximal weakness has been reported during IATs, and the documented gradual recovery of strength over 145 seconds is unlikely in patients with motor neglect.

Comment 3: IATs are often undertaken in patients undergoing epilepsy surgery. However, ipsilateral language areas are only rarely selected, especially if the test is positive. Even if the test is negative, the selected area does not necessarily include typical language areas. Therefore, little information on the validity of the IAT is available. IAT validation studies in predicting aphasia may be divided into four categories: true negatives, true positives, false negatives, and false positives. A false positive is someone who passed the IAT but nonetheless became aphasic. The incongruity of the clinically inactivated motor area and the target area (language) is a potential problem within the Wada test itself, which is not highlighted by cases like ours, and which is possibly exacerbated by an underlying subcortical lesion. Intrahemispheric reorganisation or interhemispheric language reorganisation is a possible explanation for atypical language representation in our case. The possibility of residual function and intrahemispheric reorganisation in our case is supported by intact visual fields before hemispherectomy, as previously discussed.

Another issue related to the validity of the IAT is the distribution of the amobarbital. Conventional angiography, digital subtraction angiography (DSA), and SPECT have been compared in terms of perfusion patterns during IAT. With regard to posterior cerebral artery (PCA) filling in 46 patients, SPECT showed filling in 22%, DSA showed filling in 37%, and conventional angiography showed control in 50% of patients. In 24 patients with HMPAO data, a conventional angiography before the IAT (as carried out and reported in our case) can rule out perfusion of the PCA. Based on this study, no HMPAO distribution would have been observed in the PCA territory. SPECT is therefore unnecessary.

To our knowledge, we have reported the first case in which a patient continued to speak during the Wada test and then became aphasic after ipsilateral hemispherectomy. Aphasia after hemispherectomy was not anticipated. We agree with Drs Schulze-Bonhage and Quiske that it is possible that similar complications resulting from epilepsy surgery based on Wada tests are under-reported. The teaching point of our case report is that a negative unilateral routine Wada test result for language may be misleading in an older patient with a pre-existing left subcortical lesion that was acquired after the primary development of language function. We encourage other epilepsy centres to reflect on their current practice of unilateral Wada testing for presurgical language lateralisation in selecting patients and possibly replace or supplement the Wada test by additional language lateralisation techniques.
We thank Dr Schulze-Bonhage and Dr Quiske for their comments.

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Tangier disease

With reference to the article entitled “Tangier disease—a diagnostic challenge in countries endemic for leprosy.””, I should like to point out that the name “Tangier disease” originated from a small island in Chesapeake Bay, USA and not from Morocco as stated by the authors.

The first case of this uncommon disease with neurological involvement was reported by Koen et al in the Lancet in 1967 and by Engel from the USA soon after. Detailed neuropathology on this condition in our and one other case was reported by Koen et al in 1973.

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BOOK REVIEWS

Neurology for non-neurologists, 4th edition


This book is targeted at “students, residents and practicing physicians whose primary interest is not neurology”, although at 342 pages, no colour, and very few diagrams or pictures, I felt this may be a little off putting to non-neurologists and neurologists alike! However, this is an extensive review that considers basic science, pathophysiology, and management issues, which is a useful reference tool. The first three chapters (60 pages) are an introduction to neurology and include a chapter on neuroanatomy, as well as a thorough discussion about the neurological history, examination, and common investigations. There then follows chapters on specific areas such as headache, cerebrovascular disease, dementia, and deteriorating diseases. The chapters are detailed but again may be considered overwhelming for the general physician, for example in the headache chapter four pages are dedicated to the complete International Headache Society’s classification of headache.

Interestingly, throughout the book there are very few references to when a patient should be referred on or would benefit from the input of a neurologist. Considering this book deals with rare, serious, and genetic neurological disorders, I find this approach rather disappointing. It is also written predominantly for the American market, another factor that perhaps makes the lack of reference to the need for specialised neurological involvement surprising. Drug names, dosage schedules, and units are in the accepted US format.

In summary, this is a concise but detailed reference book covering a wide range of neurological conditions. It is a useful tool for students and junior doctors but it perhaps encourages the reader to manage complex and rare conditions themselves and neglects the need for specialist input.

V L Stevenson

Neuropsychopharmacology


The European Institute of Healthcare devoted its 5th Neuropsychiatry Symposium to neuropsychopharmacology. Topics were chosen by the organisers and included, Professor Fleischhacker and Brooks, to bridge the gap between basic neuroscience and the clinical fields of neurology, but mostly psychiatry. The slim volume comprises seven chapters that cover a wide range of topics from histamine receptors to developments in substance dependence research and the pharmacotherapy of schizophrenia. The chapters are uniformly up to date and written in a challenging and interesting way. Some of the chapters are extremely comprehensive, for example that on migraine contains 199 references.

Various advances in the therapeutics of neuropsychiatric conditions are reviewed. As a psychiatrist/psychopharmacologist, I must admit to finding the three last chapters the most fascinating. Professor Deakin reviews serotonin function in relation to antisocial personality disorder and depression. He adduces interesting arguments that serotonin abnormalities are related to the difference being an anatomical selection in the dorsal raphe nucleus. This hypothesis gives rise directly to therapeutic suggestions for treating depression. These are of course largely in place. However, the treatment of antisocial personality disorder opens Orwellian prospects and will need carefully monitoring.

It was particularly interesting to see a chapter devoted to future directions and substance misuse from Professor Nutt’s group in Bristol. My only criticism of this chapter was that it was too short and only whetted one’s appetite for further information concerning receptors, brain pathways, and promising avenues of drug development.

The final chapter by one of the editors deals with new developments in the pharmacotherapy of schizophrenia. The immense clinical experience of Professor Fleischhacker is shown by his acknowledgement that improved formulations, such as rapidly dissolving tablets and syrups, have an important part to play, in view of the poor adherence to treatment of patients with schizophrenia. He also reviews polypharmacy that is the norm in many practices and points the way to further developments with new classes of compounds.

Some curmudgeonly remark the English is often not carefully sub-edited and there are numerous typos (for example, see page 65). Some of the articles contain very long paragraphs that make the content difficult to follow. There is also no uniformity of reference style. One would expect Springer-Verlag to be able to afford the services of an experienced native English speaking sub-editor.

The very last page states that the paper in the book was acid free in conformance with the requirement for archival. I suspect that the paper itself will outlive the content of these chapters, but that is not bad experience.

Overall, the book provides a rapid updating of knowledge across the field in neuropsychiatry. Inevitably, it will be out of date fairly quickly, but in the meantime it is important reading even for the cognoscenti.

M Lader

The psychology of bulimia nervosa: a cognitive perspective


The first description of bulimia nervosa was published in 1979 by Russell. Since that time there has been a burgeoning literature on all aspects of the disorder. This has coincided with the growth of cognitive psychology and cognitive behaviour therapy (CBT). As might be expected, the cognitive approach has been used in the investigation of bulimia nervosa with considerable success, and CBT has become the “gold standard” for adults requiring treatment on the basis of many
randomised controlled trials. This book is a timely review of the literature, at a time when it is still just possible for a single author to achieve a clear understanding of the field.

The background chapters cover the diagnostic features, epidemiology, and theories of aetiology of the disorder. There is description of emerging treatments, with special emphasis on CBT. Fairburn’s established model of CBT balances the focus on changing eating patterns with cognitions and is fairly described. This is the foundation for Cooper’s original work, which places more emphasis on core beliefs, for example, regarding the self and how they relate to beliefs about eating and dieting. Although Cooper’s model has some evidence in support of its rationale, there is no evidence from treatment trials that it achieves greater benefits than Fairburn’s approach, and so CBT practitioners might wonder why they should change to the new approach.

Overall this is a well written and well organised book. Each chapter has numerous useful summary tables of the main points covered in the text. Weaknesses of the book are its rather poor coverage of developmental issues, and negative bias with regard to drug treatments. Despite these points, the book achieves very well its main aim of providing a useful resource for trainees and more experienced clinicians, and should have a place in any well stocked library.

The clinical practice of critical care neurology


It was once said of the English cricket team “they played well but the only trouble is that they can’t bat, can’t bowl and can’t field”. In the United Kingdom we make similar comments about neurological intensive care— it’s a good idea and important but we have no units, no specialists, and no training. This is, of course, an exaggeration but we continue to fall behind the United States and Europe in the provision of care for patients acutely ill with neurological disorders. It must be shown that these units make a difference to the outcome of patients but it is both intuitively obvious and reflected in evidence-based medicine that providing the best care for patients will lead to better outcome. Wijdicks, together with others, has defined the field of neurological critical care with his textbooks and original contributions. The clinical practice of critical care neurology is a landmark in the field and the rapid publication of an enlarged and improved second edition reflects the importance of this book. Wijdicks writes from an extensive experience of the care of patients with critical neurological illness. The book has all the advantages of a single author text, being coherent and logical with a thorough coverage of the subject. The layout of the second edition is a considerable improvement—the text is easier to follow and many of the practical procedures are more clearly described and illustrated. The book is divided into six sections, three deal with general aspects of neurological intensive care, discussing the practical details of setting up and running a neurological intensive care unit, including the equipment and technology used in monitoring and treating the patients; and the clinical and ethical issues at the end of life. The major part of the book deals with the management of specific disorders, post-operative complications, and the management of the systemic complications seen in these patients. It is inevitable that there will be individual quirks in such a wide ranging review of neurological management but the majority of these reflect different “fashions” between the United States and the United Kingdom rather than any fundamental differences in the interpretation of the evidence base.

Neurological intensive care is a sub-speciality where the clinician has to cover a broad spectrum of the specialty and this is superbly reflected in the second edition of Wijdicks’ textbook. A single author monograph of this sort cannot hope to be exhaustive but Wijdicks has achieved a remarkable job of ensuring details are covered whilst emphasised despite the breadth of coverage. This book provides essential reading and reference for all neurologists and intensivists charged with the management of patients on intensive treatment units with primary neurological disorders.

R Howard

Stroke genetics


This is an extremely useful book that goes well beyond its stated aim as an introduction to stroke genetics for both the interested clinician seeing stroke patients, and the basic scientist entering this field of research. The book comprises 14 chapters with contributions from 18 authors. The basic science areas are well covered, including the genetic epidemiology of stroke as well as monogenic and polygenic contributions to ischaemic stroke. What marks the book out from other “neurogenetic” texts, however, is the success with which it takes genetics out of the laboratory and into the general neurological clinic. This is exemplified by the last chapter, which presents a practical approach to investigating a patient presenting with stroke for underlying genetic disorders. The three chapters dealing with the genetics of the various causes of intracerebral haemorrhage including approaches to screening for familial intracerebral aneurysms, are of particular clinical relevance, as is the excellent introduction to genetic counselling.

Cerebral venous thrombosis as a cause of stroke was mentioned in several chapters, but unfortunately not covered in a chapter of its own. I would have welcomed more detailed guidance on the utility of screening for genetically determined prothrombotic disorders, and how to deal with the uncertain predictive values of test results. However, this is a minor criticism of an otherwise successful book that should be of interest to general neurologists as much as stroke geneticists.

M R Johnson

Practical psychiatric epidemiology


It is a pleasure for a reviewer to recommend a book wholeheartedly and this I am happy to do with the present volume. It lives up to its title of taking a practical view of the topic of psychiatric epidemiology and the editors have ensured a graded approach from basic principles and their application, into the more esoteric areas of interpretation of results and the consideration of more specialised and newly developing aspects of the subject. This is not to say that it is a particularly easy book to read and the dilettante reader might well find it discouraging in its degree of detail. On the other hand, for the aspiring psychiatric epidemiologist or, just as importantly, for any researcher in psychiatry who wishes to make valid use of epidemiological methods, this book is required reading. As only one example of the latter, anyone planning to undertake a trial of a new psychotropic drug should carefully read the descriptions of potential bias in sample selection and category assignment.

I doubt if the average psychiatrist will want to buy a book on this rather specialised and research orientated subject but it should certainly be in every psychiatric library (and not just in the UK). It would be extraordinarily useful as a class text for a formal course in psychiatric epidemiology with a knowledgeable teacher acting as guide.

The editors have ensured a high quality of presentation throughout and, despite multiple authorship, there is relatively little repetition.

If I have a cavil it is not about the book but about the topic. In their introductory chapter the editors bemoan the lack of demonstrable validity of many modern research results obtained by using contemporary diagnostic and epidemiological methodologies (p 7). They also express a yearning for the discovery of genetic markers applicable to psychiatric illnesses (p 8). One can only respond to these comments because so often at present we find ourselves chasing shadows. However, I think there is increasing cause for optimism as neuropsychiatric research becomes increasingly finer grained.

As this book demonstrates, when the objective criteria emerge, there are tried and reliable methods standing ready to be applied.

A Munro

Self-assessment colour review of clinical neurology and neurosurgery


Time was when none of us would have considered voluntarily using self-assessment as a form of learning. Exams were feared and to be avoided. This book shows that some progress has been made and modern learning methods may have some validity! It is a small slim book, which makes a welcome change. It is based on the idea of using self-assessment to aid and encourage learning. On the right hand pages are a series of questions with the
Disorders of neuronal migration

This is an excellent, concise book that is written by the leaders in this rapidly growing subject. Peter Barth is to be congratulated on producing such a helpful and timely volume.

The book is written from the perspective of a clinician who wants to understand the relevance of the latest embryological, genetic, magnetic resonance, and experimental data on migration disorders and it is of particular relevance to paediatric neuroscience.

For example, the chapter on morphogenesis of the human cerebral cortex by Cainess, Takahashi, and Nowakawaki, makes regular reference to important pathological mechanisms while describing normal development in a way that is most refreshing. The account of Lissencephaly by Dobyns and Léventer is masterly. There are also very nice summaries of the role of excitotoxic damage by Gressens, Barkovich, and Evrard, and of fetal disruption by Peter Brooks.

It is not possible for a book of this size so well referenced to be clinically comprehensive and I think that this is a reasonable recognition of the imperative of providing good information on genetics, neuroradiology, and pathology. I think, therefore, that it is essential reading for those concerned with developmental and paediatric neuroscience.

Women with epilepsy: a handbook of health and treatment issues

This book was written in recognition of our increasing knowledge of the way in which epilepsy affects women specifically, and examines the issue from a biological, psycho-social, and treatment standpoint. Its aim was to provide information both for women with epilepsy and those around them, including their clinicians, families, and friends. It is a wide ranging book, covering such diverse areas as a historical perspective of the woman with epilepsy, the epidemiology, classification, and genetics of epilepsy, hormonal issues and sexual dysfunction in women with epilepsy, family planning, pregnancy and teratogenicity, legal and safety issues, and the psychosocial impact of epilepsy.

Inevitably in a book of this nature directed at a varied audience, there is a tension between clarifying the information sufficiently for lay readers while maintaining the interest of the clinician. Generally the authors succeed in this balance, but I felt that some of the chapters, particularly those addressing hormonal issues, remained rather technical for the lay reader. Other sections, such as those on the impact of epilepsy on relationships and psychiatric complications in epilepsy, were clearly written and provided useful information rarely available in standard textbooks, although the chapter on legal issues facing women with epilepsy was unfortunately of limited relevance to those outside the US.

The need for more information about the impact of epilepsy on women has been widely acknowledged in recent years, and this book will be of great interest not only to women with epilepsy and their families, but also to their doctors and epilepsy nurse specialists.

Unipolar depression: a lifespan perspective

At first glance this is a very unimposing book, with a cover design that makes little impression and barely 200 pages for so important a subject. The prevalence of major depressive disorder is 0.4–2.5% in school aged children, 3–8% in older adolescents and adults, and up to 16% in later life. This clearly has a massive impact on all aspects of the community and on the number of people who suffer depression.

Multi-author books often show a lack of balance between chapters, giving each topic the same level of importance, and often have duplication and overlap of subject matter. This book manages to avoid these problems, in large part because of the way it is structured in taking the reader through the various stages of life: infancy and pre-school years, the school aged child, adolescence, early adult life, mid life, and later life. This chronological, or maturational, perspective gives a much better appreciation of the intimate interactions of biological/genetic factors and experiences. Reading the book with this perspective is also in some ways a moving experience, as if watching and anticipating the development and progress of the illness in an individual.

The book takes us through the literature up to and including 2002, which is as good as it can be expected for a book published last year. The review of the literature is thorough and gives a balanced perspective. This is in large part because the authors are experts in the field and manage to write in a clear and pragmatic way linking clinical management with up to date research.

It is a cliché to say you should not judge a book by its cover and this is a clear example of it. I am very happy to be able to keep this copy.

M Maier

www.jnnp.com
Tangier disease

R S Kocen

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