The management of depression: the evidence speaks for itself

Gin S. Malhi, Erica Bell, Darryl Bassett, Philip Boyce, Richard Bryant, Malcolm Hopwood, Bill Lyndon, Roger Mulder, Richard Porter, Ajeet B. Singh and Greg Murray

Summary
Comparing the recommendations of two recently published national clinical practice guidelines for depression, this editorial highlights the concordance of advice concerning the selection and sequencing of therapies. Lifestyle and psychological interventions feature prominently and there is broad agreement regarding medication choice and optimisation strategies. The guidelines are therefore a useful resource.

Keywords
Guidelines; evidence; mood disorders; depressive disorders; major depression.

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Managing depression is the bread and butter of psychiatry. Depression is common, both on its own and comorbid with other disorders. It thus confers a significant disease burden and consequently, organisations have developed guidelines to inform management.

Further, both draw on traditional classifications (DSM-5 and ICD-11) to define the boundaries of depression. However, NG222 divides depression according to severity—coalescing the older terms ‘sub-syndromal’ and ‘mild’ to form ‘less severe’, and grouping ‘moderate’ and ‘severe’ as ‘more severe’. These relativistic terms (less and more) imply a continuum and lend the definition of depression a dimensional perspective, akin to that used in the MDcpg2020. At the same time, like its antipodean counterpart, NG222 also recognises subtypes such as psychotic depression.

For example, the MDcpg2020 views depression as a multifaceted entity that can be scaled according to severity (like NG222) but can also be subtyped depending on symptom profile, with the added sophistication of clustering symptoms into domains of activity, cognition and emotion (termed the ACE model). Both guidelines emphasise functional impairment as a critical determinant of help-seeking, interventions, long-term outcome and societal impact.

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may require ECT immediately. Thus, all treatments (psychological interventions, psychoeducation, exercise, antidepressants and ECT) can be administered from the outset, dependent on the clinical presentation and patient preference. In other words, like NG222, all these treatments can be regarded as ‘first-line’, and once again there is good consensus between the two sets of guidelines as to what treatment can, and should, be prescribed when embarking on the management of depression.

NG222 recommends that if an individual has ‘not responded at all after 4 weeks of antidepressant treatment at a recognised therapeutic dose, or after 4–6 weeks for psychological therapy or combined medication and psychological therapy’, then non-response should be explored methodically. This includes a review of the diagnosis and treatment, while maintaining a positive and reassuring stance and a willingness to switch strategies. A similar versatile approach is advocated in the MDcpg2020, which also emphasises personalising treatment where possible, based on symptom profiles, treatment history and patient preference.

**Further-line treatment**

This straightforward term coined by NG222 addresses the multiple strategies that can be employed once initial attempts to obtain response are unsuccessful. Here, a detailed diagram (hot linked as ‘visual summary on further-line treatment’ in section 1.9.1) emphasises the need to address problems that may not seem to be directly pertinent to depression, such as personal, social or environmental factors, and advises that other illnesses (especially personality dysfunction) should also be considered as potential contributors to depression. This complex NG222 schema for management contains elements of the ‘medication, increase dose, augment, switch’ (MIDAS) approach described by the MDcpg2020. In both guidelines, it is emphasised that functional improvement can occur after any of the interventions and subsequent management strategies. For instance, treatments can be optimised using increases in dose where possible, augmenting and switching, and combinations can be trialled involving different kinds of intervention or for augmentation purposes.

These strategies address the needs of the majority of people with depression. However, also important are those who do not respond to treatment – described variably as having ‘difficult to treat’ or ‘treatment-resistant’ depression (TRD). This is not an uncommon outcome and is often the result of departing from best practice. This is why NG222 also emphasises the importance of accurate diagnosis and re-evaluation of the diagnosis throughout the course of the illness: an important message that is promoted by both sets of guidelines.

**Evidence**

Guidelines rely on research evidence to formulate their recommendations. In the MDcpg2020 this was assessed using National Health and Medical Research Council (NHMRC) guidelines (www.nhmrc.gov.au).
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gov.au/guidelinesforguidelines) and the merits of various treatments were determined on the strength of evidence overall. However, NG222 dug deeper still, and conducted detailed analyses of the available evidence to underpin its recommendations regarding clinical effectiveness. In addition, NG222 considered cost-effectiveness when making specific recommendations and factored both the accessibility and the ability to implement treatments into its considerations.

The extent to which available data have been interrogated, and the manner in which it has been synthesised within NG222, has to be commended. Naturally, there are many instances in which the evidence is incomplete or of poor quality. And here, rather than not making any recommendations whatsoever, both the MDCpg2020 and NG222 have opted to offer some clinical guidance. The MDCpg2020 does this more formally and distinguishes between evidence-based recommendations (EBRs) and consensus-based recommendations (CBRs). The latter were formulated when: (a) the existing intervention evidence base was absent, ambiguous or of doubtful clinical impact in the Australian and New Zealand context; and (b) the Mood Disorders Committee (based on collective clinical and research knowledge and experience) reached consensus on the clinical utility of the recommendations. CBRs acknowledge their limitations, but nevertheless provide useful advice on how to navigate less-established care options once more established options have been reasonably exhausted. At their core, the two sets of guidelines are evidence-based and pragmatically add to this evidentiary kernel, along with experience, cost and accessibility considerations, to enhance translation into practice. Consequently, the recommendations within the guidelines overlap considerably and this not only lends strength to their findings but provides greater confidence for clinicians choosing to base their treatment decisions on the advice in the guidelines.

Nevertheless, the use of CBRs highlights a key limitation of all clinical guidelines, namely, the paucity of empirical evidence to support recommendations for many key clinical questions or decisions. Thus, to produce more comprehensive guidelines that reflect the many clinical complexities of the illness, further research on real-world depression is needed. Steps towards this goal have already been taken by a recent European Brain Council initiative, which has identified treatment gaps between ‘best’ and ‘current’ practice. Examining practices in six European countries, researchers found gaps in the detection of depression and provision of treatment, especially with respect to follow-up and access to specialist care. Consequently, they have formulated a comprehensive set of recommendations to better meet patient needs.

Conclusions

Overall, the agreement between the MDCpg2020 and NG222 yields several key guiding principles with respect to how depression should be managed. These include advocating for robust diagnosis and ongoing re-evaluation of this throughout management. They prioritise the use of psychological and lifestyle interventions where possible, and emphasise the adoption of a flexible style of management within the recommended schema of treatments and therapeutic strategies to personalise care.

References


Data availability

Data availability is not applicable to this article as no new data were created or analysed in this study.

Author contributions

G.S.M. and E.B. conducted the initial research and drafting of this piece. All other authors contributed to the writing and editing and approve of the final manuscript.

Declaration of interest

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