I have no such moment to offer. But brilliant folk like Lawrie and his colleagues have that tradition and they perhaps raise the chances that such scientific inspiration can help us once again.


As psychologists who have long researched and argued for a dimensional view of psychosis, we would like to comment on Lawrie et al’s editorial.1 We are surprised that the authors pay no attention – with one exception – to the psychological literature. If they had done so they would know that considerable evidence supporting the continuum view has accrued over many decades. The one psychologist they do cite – the late Paul Meehl – is an unfortunate choice. Quite apart from the fact that it is unclear to us how Meehl’s taxonomic (categorical) approach actually helps their case, the authors ought to be aware that the theory is now on the wane. A more viable alternative is what we have termed a ‘fully dimensional’ theory that is capable of encompassing more of the

Lawrie and colleagues urge us not to reject the current categorical classification system prematurely.1 We wish to add to the argument that a categorical system is more likely to be internationally useful. More than 80% of mental illness occurs in middle- and low-income countries.2 Much of the world’s mental illness is seen in overstretched clinics, by practitioners who treat up to 100 patients a day and often have had no training in psychiatry since medical or nursing school. Administering the rating scales necessary for a dimensional system may be possible in high-income countries, but is difficult or impossible elsewhere. The categorical classification system can be used quickly by someone with relatively little training. There is also the problem of translating and validating the rating scales into hundreds of languages. Most published research currently uses the same categorical system, which means that it is useful to doctors all over the world. If the research were to refer only to a dimensional system, then it would not be useful in settings where it is impossible to administer the rating scales. The categorical system gives more people access to evidence-based treatment than any dimensional system would. A classification system that is going to be used all over the world needs to be simple and robust across healthcare systems, languages and cultures, and this is just as important as how closely it resembles the truth.


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doi: 10.1192/bjp.198.4.323a

As psychologists who have long researched and argued for a dimensional view of psychosis, we would like to comment on Lawrie et al’s editorial.1 We are surprised that the authors pay no attention – with one exception – to the psychological literature. If they had done so they would know that considerable evidence supporting the continuum view has accrued over many decades. The one psychologist they do cite – the late Paul Meehl – is an unfortunate choice. Quite apart from the fact that it is unclear to us how Meehl’s taxonomic (categorical) approach actually helps their case, the authors ought to be aware that the theory is now on the wane. A more viable alternative is what we have termed a ‘fully dimensional’ theory that is capable of encompassing more of the
known facts about psychosis, including the clear dimensionality of the risk of illness and the likely form of the heritability underpinning this, coupled with the notion of discontinuity to recognise the break in behaviour and psychological state that occurs when vulnerability translates into clinical symptoms. Importantly, the model also recognises something that Lawrie et al entirely ignore – the fact that psychotic traits can have a healthy expression that takes the individual outside the domain of psychiatric judgement.

Of course, many questions remain, such as how to deal with the overlap between schizophrenic and affective expressions of psychosis, explain the underlying biological mechanisms of these disorders, and incorporate into our thinking how expressions of vulnerability can vary from sick to benign. However, answers to these questions will not make dimensionality go away, for it is part of the essence of human variability (of which psychosis is one form).

On the practical front, these ideas admittedly make for a messy picture that is inconvenient for clinicians seeking a neat solution to diagnostic issues. But psychiatry does itself no favours by ignoring them and retreating (yet again) behind the ramparts of its traditional mode of thinking. Fortunately, as Lawrie et al will be aware, their profession actually has moved forward in recent years towards an attempt to find ways of integrating both dimensional and categorical perspectives into its future diagnostic systems. Our plea is that, in doing so, it becomes an even more ‘psychologically informed’ psychiatry.

Authors’ reply: We thank Drs Gordon and Shoesmith for their interest in our editorial, their complimentary remarks and their considered responses to what we said. Dr Gordon repeats our call to avoid prematurely abandoning categories or dimensions, and highlights the lack of known diagnostic biomarkers for psychosis, either as a whole or for current subtypes. Tandon et al1 did not really consider this, quite reasonably, as their review focuses on diagnostic issues. But psychiatry does itself no favours by ignoring them and retreating (yet again) behind the ramparts of its traditional mode of thinking. Fortunately, as Lawrie et al will be aware, their profession actually has moved forward in recent years towards an attempt to find ways of integrating both dimensional and categorical perspectives into its future diagnostic systems. Our plea is that, in doing so, it becomes an even more ‘psychologically informed’ psychiatry.

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doi: 10.1192/bjp.198.4.323b

An unjust review

In his review of my book Fiction’s Madness,1 Beveridge comments on my omission of Laurence Sterne’s Tristram Shandy in discussing the history of the novel form.2 On fictional development in the 1950s, Hawthorn3 pointedly excludes Tristram

2 Lawrie SM, Olabi B, Hall J, McIntosh AM. Do we have any solid evidence of clinical utility about the pathophysiology of schizophrenia? World Psychiatry 2011; in press.
3 Beveridge comments on my omission of Laurence Sterne’s Tristram Shandy in discussing the history of the novel form.2

Stephen M. Lawrie (email: s.lawrie@ed.ac.uk), J. Hall, A. M. McIntosh, D. G. C. Owens, E. C. Johnstone, Division of Psychiatry, Royal Edinburgh Hospital, Morningside, Edinburgh EH10 5HF, UK
doi: 10.1192/bjp.198.4.324
Shandy as anticipating the novel and I made plain that the (postmodern) changes I observed 'came into common usage in Europe and the Unites States in the last three decades or so' (Hawthorn: p. 62). To negate (my) differentiating modernist fiction from the 1950s postmodernist 'shift' might make good criticism if not merely advanced as opinion.

On my text choices being idiosyncratic, I acknowledged this inevitability (p. vi) before providing choices of others as a balance, including David Goldberg. But this was ignored and readers left with assumptions of my eccentricity.

I did not identify psychoanalysis as a dominant force in the 1930s. I asserted its significance as an interest in Freudianism, in the 1920s, with 'think-tanks' involving John Rickman, Lionel Penrose, A. G. Tansley and John Bowlby, who qualified medically in the 1930s. This interest persisted into the 1950s, some medical superintendents being conversant with psychoanalysis whose emergent tensions, in psychiatry, I addressed in my chapter on Pat Barker's Regeneration.


In effect, your reviewer ignored most of my book, opting for points of little intellectual interest. As for my (perceived) disparaging remarks about psychiatry 'throughout the book', my critical take on psychiatrists Dr Yealland (Chapter 3) and Dr Weir-Mitchell (Chapter 5) stemmed from fiction. My 'disparaging comments' were exceptionally sporadic but their effect clearly outweighed the rest of my text.

It is false that I 'dismiss' Nietzsche, Socrates and Foucault. I critically quoted Foucault thus: 'Shall we try reason: to my mind nothing could be more futile' (p. 66). I attributed only to Socrates that he was Plato's mouthpiece and placed my take on Nietzsche within Hesse's Steppenwolf and Richard III.

In general, the review was ill-considered, selectively dismissive and factually inaccurate.

Theories on the evolutionary persistence of psychosis

We note that the Darwinian models of psychosis reviewed by Kelleher et al in their editorial were all variants of the 'costly by-product' evolutionary model whereby an adaptive neuro-biological system that enhances fitness in the vast majority of the population generates the risk of error in a small minority, resulting in psychosis (including schizophrenia). Burns identified the frontotemporal and frontoparietal cortical connections of the social brain, whereas Crow proposed that the dysregulation occurs in the language centres.

We wish to propose a different and entirely environmental Darwinian formulation for the non-affective psychoses based on an 'environmental mismatch' model. We have explained elsewhere that, although we agree with Burns' proposal regarding locating the dysregulation and dysconnectivity within the social brain, we contend that the aetiology of the dysregulation relates to the effects of the novel post-Neolithic social environment.

Although the susceptibility to non-affective psychosis, including schizophrenia, is likely to be ancient, the schizophrenic and the non-affective psychosis phenotype did not manifest itself until very recently in our species' history. In other words, the risk of these disorders lay dormant and did not become evident until the post-Neolithic period.

Hence, we have proposed a reformulation of the social brain theory of schizophrenia and contend that schizophrenia (and the non-affective psychoses) are novel human phenomena that arose following the establishment of large permanent human settlements that accompanied the advent of agriculture and the abandonment of the hunter–gatherer way of life. We have contended that the blurring of the demarcation between in-group and out-group membership and living in close proximity to strangers is a stressor that can lead to perturbation in the development of the social brain in vulnerable individuals, resulting in the syndrome of schizophrenia. Hence, according to our formulation, schizophrenia is the result of a mismatch between the post-Neolithic human social environment and the

Author's reply: I would like to make the following points. First, in referring to Laurence Sterne's Tristram Shandy, which is regarded by most commentators as a novel, I was challenging the author's contention that: 'From the eighteenth century through to the nineteenth, novels were realist by nature [. . . ] from the 1950s, however, novels began to move in mysterious ways. Suddenly "Multivoiced" narratives, unreliable narrators, allegories, genre dodging, satire, and allusiveness [. . . ] became the order of the day' (Clarke, pp. 11–12). Sterne's Tristram Shandy, written in the 18th century, and James Hogg's The Private Memoirs and Confessions of a Justified Sinner, written in 1824, experiment with the genre and with the notion of the unreliable narrator. Indeed, Clarke himself (p. 17) cites Ford Madox Ford's 1915 novel The Good Soldier as representing a good example of an unreliable narrator.

Second, in his letter the author states that he did not identify psychoanalysis as a dominant force in the 1930s, but in his book he writes: 'Psychoanalysis was a major force in English psychiatry during the 1930s' (p. 150).

Third, as regards disparaging remarks about psychiatry, the quote about the smugness of male psychiatrists comes directly from the author, not from a novel. Elsewhere we find other critical remarks. Commenting on psychiatric training the author states: 'three years of preparation for membership of the Royal College of Psychiatrists [. . . ] requires not a whit of training in interpersonal relations, little of self-reflection, or what it means to be human. Such diversions might inhibit the self-assuredness provided by a medical model of madness. Alternatively, of course, the hyped confidence may simply compensate for the psychiatrists' self-perceived fragility compared with the knowledge basis and status of other medical specialties' (p. 147).

Finally, with reference to a dismissive approach to major thinkers, the author discusses what he calls 'Socrates' infamous claim that no one can knowingly do wrong,' and concludes: 'Perhaps Socrates got it wrong' (p. 156). He writes that 'Although Nietzsche's Superman (Übermensch) was realised most horrifically, in our own time, by the Nazis, the impulse to stomp on others continues' (p. 136). He also observes: 'Foucault foolishly suggests abandoning rationality itself' (p. 186).
design of the social brain. We highlight the importance of the distinction between in-group and out-group membership that lies at the heart of intergroup conflict, violence and xenophobia. Our hypothesis (the out-group intolerance hypothesis) provides an explanation for the disparities in the prevalence of schizophrenia across the world and for the higher risk of this condition among immigrants and city dwellers. We propose that our hypothesis can account for a range of disparate epidemiological and other findings regarding schizophrenia that have thus far defied explanation by other theories, including the Darwinian by-product formulations reviewed by Kelleher et al.


Kelleher et al. note the significant prevalence of non-clinical psychotic symptoms in the general population and discuss some hypotheses regarding its evolutionary survival. One theory not mentioned by them or those who have so far responded is a trait known as schizotypy. While accepting that to some degree the whole topic is rich with speculation, I suggest that schizotypy may be the missing piece in the puzzle. What follows is necessarily a brief summary of some of the relevant literature.

Differing from both schizotypal and schizoid personality disorders, schizotypy is a heritable trait associated with an increased likelihood of creativity and of religious or mystical experiences. Importantly for this discussion, schizotypy also appears to be necessary but not sufficient for the development of schizophrenia, although not all those with schizotypy develop psychotic illnesses.

The four key dimensions of schizotypy are unusual experiences (which may be considered to be related to positive symptoms), cognitive disorganisation (related to thought disorder), introverted anhedonia (related to social withdrawal and depression) and impulsive non-conformity. This last is related to some of the disturbed behaviour, such as aggression and self-harm, seen in a range of psychiatric illnesses, including psychosis.

Regarding creativity, additional research by Nettle and Clegg further finds that schizotypal traits are associated with increased ‘evolutionary fitness’ due to a greater number of sexual partners (and therefore offspring) in those with the unusual experience and impulsive non-conformity dimensions of the trait. In those with the former but not the latter dimension, the relationship with mating success is mediated by creativity. Nettle & Clegg have proposed that schizotypal traits, which in this case may be a proxy for some non-clinical psychotic symptoms, have therefore persisted because their potential negative effects are offset by enhanced mating success.

Regardless of the outcome of the search to understand the persistence of psychotic symptoms in human beings and of possible future research involving those who have the non-clinical psychosis phenotype, it is important for people working in mental health services to remember that not all those they encounter with symptoms are ill. For those that are unwell, there will be other aspects of their existence that are positive and that may be life-enhancing for them and those around them. They should be encouraged to develop these aspects of themselves as part of their long-term recovery, in addition to the treatment and support they receive from health services, carers and friends.


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doi: 10.1192/bjp.198.4.326a

Don Quixote and Sancho Panza: folie à deux?

Martins de Barros & Busatto Filho date the first report in fiction of folie à deux to the Brazilian author Machado de Assis in 1879. I submit that the first fictional account of ‘shared delusions’ was by Miguel de Cervantes over 250 years before. Cervantes wrote Don Quixote de la Mancha in or around 1604, publishing the first part in 1605 and the second, a decade later.

In Don Quixote, the eponymous hero, we have a domineering and volatile fantasist driven ‘out of his wits’ by the undue influence of books of chivalry: ‘He so buried himself in his books that he spent the nights reading from twilight till day break and the days from dawn till dark; and so from little sleep and much reading, his brain dried up and he lost his wits.’ His character is steeped in rich descriptions of grandiloquent and persecutory delusions, polymorphic hallucinations and cognitive blunting. Sancho Panza, his squire, whom he enlists as his companion for his travels, is described as ‘an honest man – if a poor man can be called honest – but without much salt in his brain-pan.’

So we have a dominant Don Quixote, who has lost his reason, and a submissive, not so bright Sancho Panza, thrown together through much of their travels, creating a situation ripe for the development of folie à deux. And indeed we see a slow erosion of reason in Sancho Panza. He initially displays some resistance and skepticism to Don Quixote’s delusions about windmills being monstrous giants or St Benedict’s monks being a crew of wicked and diabolical ‘perfidious scoundrels’. But he increasingly becomes convinced of the veracity of Don Quixote’s beliefs. One example should suffice, the example of the balsam of Fierabras. This is a concoction that Don Quixote claims he can make on the cheap. He tells Sancho Panza, ‘If ever you see me cut through the middle in some battle [. . .] you have only to take the part of my body that has fallen to the ground and place it neatly and cunningly, before the blood congeals, on to the half that is still in the saddle, taking special care to make them fit exactly. Then you must give me just two drops of this balsam to drink and, you will see, I shall be as sound as an apple.’ Sancho replies, ‘If that is so, from now on I renounce the governorship of the promised isle, and all I want in payment for all my good services is for your worship to give me the recipe for that marvelous liquor.’

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We gratefully acknowledge the support of the National Institute for Health Research Mental Health Patient and Public Involvement Research Centre for Addenbrooke’s Hospital, Cambridge.

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By the end, Sancho Panza’s descent into these fantastic delusions is complete. So much so that at his death bed, when Don Quixote regains a measure of lucidity and tries to persuade Sancho to see reason, Sancho Panza is completely insightless and unamenable.

We are not witness to the effect of the separation of Sancho Panza from Don Quixote, as the story ends before it. But apart from that, the description of folie à deux is complete in this wonderfully told tale.


The benefits of an active control arm
Lesem et al highlight the importance of rapid and safe treatment of agitation, indicating the delayed onset of action associated with intramuscular injection. They make no reference to the time from oral medication administration to onset of effect. However, the combination of oral atypical antipsychotics, with or without benzodiazepines, is well described. Small trials have compared the efficacy of oral atypical antipsychotics with that of intramuscular typicals and produced mean changes in rating scale scores similar to those in Lesem et al’s paper, on similar timescales.

When alternative treatments exist, placebo-controlled trials are appropriate if the target condition is characterised by a high placebo-response rate or a high relapse, remission or spontaneous resolution rate, or if existing therapies are partially effective or have high side-effect rates. Inclusion of an active control arm to the trial would have added to the number of patients required in each arm, but would have provided valuable information on the tolerability and efficacy of the inhaled or oral medication.

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Vijay Kumar
Access the most recent version at DOI: 10.1192/bjp.198.4.326a

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