Preprint version

Published in Current Opinion in Psychiatry, February 2018:

https://journals.lww.com/co-

psychiatry/Abstract/publishahead/Global mental health and schizophrenia.9929 8.aspx

Global mental health and schizophrenia

Laura Asher^{1, 2*}, Abebaw Fekadu^{3, 4} and Charlotte Hanlon^{5, 6}

 $^{\rm 1}$ University of Nottingham, Division of Epidemiology and Public Health, Nottingham, UK

² London School of Hygiene and Tropical Medicine, Centre for Global Mental Health, Department of Population Health, London, UK

³ Addis Ababa University, College of Health Sciences, Centre for Innovative Drug Development and Therapeutic Trials for Africa (CDT-Africa), Addis Ababa, Ethiopia

⁴ Global Health & Infection Department, Brighton and Sussex Medical School, Brighton, UK

⁵ Addis Ababa University, College of Health Sciences, School of Medicine, Department of Psychiatry, Addis Ababa, Ethiopia.

⁶ King's College London, Institute of Psychiatry, Psychology and Neuroscience, Health Services and Population Research Department, Centre for Global Mental Health, London, UK

*Corresponding author

Contact details of corresponding author:

London School of Hygiene and Tropical Medicine, Centre for Global Mental

Health, Department of Population Health, Keppel Street, London, WC1E 7HT, UK

Laura.asher@lshtm.ac.uk

07800 820978

Abstract

Purpose of review

The aim was to synthesise recent evidence on schizophrenia illness experience and outcomes and models of care in low and middle-income countries (LMIC).

Recent findings

There is a plurality of explanatory models for psychosis and increasing evidence that context influences experiences of stigma. People with schizophrenia in LMIC are vulnerable to food insecurity, violence and physical health problems, in addition to unmet needs for mental healthcare. Family support may help to improve outcomes if present, but caregivers may be overwhelmed by the challenges faced. Despite efforts to increase availability, evidence-based care remains inaccessible to many people with schizophrenia. Non-randomised evaluations in South Africa and Mexico indicate that psychosocial support groups for people with schizophrenia and caregivers may be acceptable and useful. Randomised controlled trials in Pakistan and China show that culturally-adapted cognitive-behavioural therapy can reduce symptom severity. There is emerging evidence that traditional and alternative medicine, such as Tai Chi, may be beneficial, but to date most research is of low quality. The challenges of biomedical-traditional provider collaborations have been highlighted. Evaluations of integrated mental health care in primary care are underway and promise to provide vital information about how to scale-up quality care.

Summary

Acceptable and effective responses to schizophrenia in LMIC should be cognisant of both cultural context and universal concerns. Efforts to enhance the quality of family support should be central to models of care.

Key words

Schizophrenia, low and middle-income countries, global mental health, mental health Gap Action Programme

Introduction

The field of global mental health has emerged in the last decade, with the primary agenda of scaling up services for people with mental health problems, especially in low and middle-income countries (LMIC). Global mental health has led the way in setting the research and service priorities for mental disorders [1]. Schizophrenia is one of the priority disorders due to its association with high levels of disability and premature mortality, human rights violations and loss of productivity [2]. There is consensus that in both LMIC and high-income countries (HIC) optimal care for schizophrenia comprises 1) A combination of antipsychotic medication and psychosocial support and 2) Community-based care with access to inpatient care for acute crises [2]. However, a severe shortage of mental health specialists means that an estimated 69% of people with schizophrenia in LMIC do not access any evidence-based care; referred to as the 'treatment gap' [3]. Family members are typically the main source of support. Accordingly, efforts to scale up care focus on delivery by trained non-specialists, an approach known as task-sharing. The World Health Organization (WHO) mental health Gap Action Programme (mhGAP) represents the most widely disseminated model of task-sharing for the treatment of schizophrenia, and other priority mental disorders, in primary care [4]. Given their wide availability and acceptability for the treatment of mental health problems, collaboration with traditional and faith practitioners is also proposed as a promising approach [5].

A refined understanding of explanatory models, risk factors, illness course, and social impact of schizophrenia is paramount for the development of acceptable and effective interventions to address the treatment gap. The notion that people with schizophrenia in LMIC experience superior outcomes than those in betterresourced settings is perhaps the most influential and enduring finding from a series of multi-country studies conducted by the WHO [6]. However, some commentators have challenged the validity of these findings, highlighting, for example, the high mortality rates in LMIC, which could have caused differential attrition of severe cases, and the variation in outcomes between LMIC settings [7, 8]. The aim of this review was to synthesise recent evidence on schizophrenia illness explanations, impact and outcomes and the accessibility, feasibility and effectiveness of models of care in LMIC.

Explanations and attitudes

Recent research has consolidated the rich literature on the ways cultural and social settings shape understandings of psychotic symptoms [9, 10]. Qualitative work in India, Nigeria and Trinidad found that disruptive behaviours, wandering and decline in functioning are more often considered signs of psychosis than distorted perceptions and beliefs. Across settings, psychosis was attributed to multiple, sometimes contradictory, causes; whilst supernatural beliefs were most prominent, biological factors, substance misuse, and psychosocial factors were also important [9]. These findings strengthen the recognition of a plurality of explanatory models for psychosis in both HIC and LMIC settings.

The implications of context have also been explored in relation to the content of delusions [11], help-seeking behaviour [9, 12], and stigma [10, 13, 14]. Campbell et al.'s findings echo the axiom that whilst the form of psychotic symptoms is universal, the content is dependent on cultural milieu [11]. In this large qualitative study amongst South African Xhosa, the majority of persecutory delusions related to bewitchment and were perceived to be caused by the jealousy of friends or family [11]. Recent multi-country cross-sectional surveys have built on previous findings that supernatural explanatory models for schizophrenia tend to be associated with greater self-stigma [13] and public stigma [10]. More nuanced accounts have lent support to Yang's theory that 'what matters most' in any society shapes both the experiences of the stigmatised and the behaviours of stigmatisers [15]. Using rich qualitative and quantitative data, Koschorke et al. concluded that, in India, caregivers' experience of stigma is underpinned by their own and their relatives' lack of achievement in gendered role expectations relating to marriage, work and social standing [14]. They also found a complex relationship between knowledge and stigma, concluding that biomedical explanatory models may not be beneficial, and may even be harmful, in stigma-reduction efforts [14]. In a cross-sectional survey (n=4476) Angermeyer et al. found that amongst members of the public in Germany, there was a greater desire for social distance from people with schizophrenia in remote relationships, in contrast to Tunisia where social distance was desired for family relationships. The authors proposed this was due to the importance of family life in Tunisian culture, and the difficulties people with schizophrenia therefore face in carrying out the social role 'that matters most' in this setting [10].

Social and health impact

New research has broadened our understanding of the ways in which people with schizophrenia in LMIC represent a uniquely vulnerable group. In rural Ethiopia, 32.5% of people with severe mental disorders had severe household food insecurity, compared to 15.9% of comparison households (n= 576)[16]. In a small cross-sectional survey (n=77), 75% of women with schizophrenia in a Nigerian outpatient clinic had experienced intimate partner violence, with around three quarters reporting verbal abuse, half reporting physical abuse and a quarter reporting sexual violence [17]. This compares to 3.1% of women in a general population sample reporting sexual violence in the last 12 months [18]. In a qualitative study in rural Ethiopia, people with schizophrenia were found to be commonly physically restrained by family members as a strategy to manage the illness in the absence of other support. There was pressure from the wider community to restrain in order to protect the individual with schizophrenia and local residents [19].

In an analysis of 2002-2004 World Health Survey data (n= 242,952), significantly higher multi-morbidity was observed in people with self-reported psychosis (36.0% (95% CI, 32.1–40.2%)) compared to those with no psychotic symptoms (11.4% (95% CI, 11.0–11.8%)) across 48 LMICs [20]. Despite lower use of second-generation anti-psychotic medications in LMIC compared to HIC, angina pectoris was the most commonly reported complaint [20], suggesting a universal pattern of physical health problems amongst people with schizophrenia. The

pooled prevalence of metabolic syndrome in people with schizophrenia across 14 Indian studies was found to be 29.8%, which is comparable to HIC [21]. Liu et al. propose a multi-level model of risk, comprising individual factors, health systems and social determinants of health, to explain excess mortality in people with schizophrenia across LMIC and HIC [22]. Growing interest in the prevalence of underweight in people with schizophrenia [23], given the known association between underweight and excess mortality in the general population [24], may have particular relevance for LMIC.

Risk factors and illness course

The apparently superior schizophrenia outcomes in LMIC have been hypothesised to arise from better social support. Indirect support for this notion comes from three recent papers of the 14-year outcomes of the Chengdu Mental Health Project cohort study of schizophrenia set in rural Southwest China (n=510) [25-27]. Individuals who were either unmarried or had no family caregiver at baseline had higher rates of homelessness and suicide, and a lower rate of survival after 14 years of follow-up [25, 26]. Individuals with family caregivers also had significantly higher functioning scores, lower symptoms scores and lower relapse rates at 14 years [25]. However, the study recruited prevalent cases, meaning that poorer outcomes may predate baseline data collection; hence a causal relationship cannot be confirmed. Furthermore, it is established globally that it is not simply the presence but the nature of family support that influences outcomes. In common with findings across LMIC and HIC settings, a recent study in Pakistan (n=53) found the relapse rate for people with schizophrenia in high-expressed emotion households was 72% as compared with 36% in low-expressed households (p= <0.05) [28].

Two small-scale but in-depth qualitative papers from South Africa contest some assumptions around the nature of informal care for people with schizophrenia in LMIC. Whilst care was readily and flexibly provided by family and community members in most cases [29], this was often because they felt obligated to do so [30]. One study found that family members used non-confrontational care strategies, including unrestricted mobility, to avoid conflict with care recipients [29]. These findings contrast with the restrictive practices, including restraint, identified as part of informal care in other settings [19]. Gaps and difficulties were also identified: informal care was sometimes not available due to caregiver employment, sickness or death [29] and caregivers experienced substantial obstacles in supporting medication adherence, especially in the context of violence, food insecurity and substance abuse [30].

Recent exploratory research has proposed universal mechanisms for the development of psychotic symptoms and further challenged the notion of a more supportive family environment in LMIC. An online community-based survey in Chile, Columbia, Indonesia, Germany and US found that, whilst participants from LMIC were more likely to report that they had any family support, they also perceived more criticism compared to participants from HIC (n=1317). Moreover, the association between criticism and psychosis proneness was stronger in LMIC compared to HIC [31]. Jaya et al. used cross-sectional survey

data (n=2350) to demonstrate that cognitive vulnerability, particularly negative schemas, mediates the relationship between social adversity and self-reported psychotic symptoms in a similar way in the US, Germany and Indonesia [32]. These findings, and similar recent papers [33], are amongst the first to explore cognitive models of psychosis in LMIC and may have important implications for the use of cognitive therapies across cultural contexts.

Models of evidence-based care and access to care

A systematic review identified 33 papers relating to mhGAP implementation in LMIC across disorders, including training evaluations and country contexualisations [4], demonstrating the importance of mhGAP for global mental health practitioners and researchers. There are indications that care delivered by primary healthcare workers can support clinical improvements in people with schizophrenia but most studies to date, such as a 12-month cohort study in Nepal, have been small (n=85) and non-randomised [34]. The forthcoming results of more rigorous evaluations, including the TASCs non-inferiority RCT in rural Ethiopia [35] and the five-country Programme for Improving Mental healthcarE treatment cohorts and facility detection studies [36], promise to shed more light on the impact of the mhGAP model.

It is increasingly recognised that alongside improved availability, services must be of high quality to achieve improved patient outcomes [37]. Reflecting growing efforts to consult people with schizophrenia in service design, Mayston et al. developed and validated the Mental Health Service Satisfaction Scale for rural Ethiopia [37, 38]. Given the limited coverage of community-based services, psychiatric hospitals remain an important source of care for people with schizophrenia in LMIC, particularly for those who are destitute [39]. The WHO QualityRights initiative aims to improve human rights conditions in mental health facilities [40], though evaluations of the impact of such initiatives have so far been of low quality [41]. A new multi-level model for interventions to reduce excess mortality in persons with severe mental disorders, comprising individual, health system, community and policy-level interventions, can be readily applied to both LMIC and HICs [22]. However, implementation and evaluation of such interventions in LMIC is limited to date.

There is mounting recognition that even where biomedical services are available, people with schizophrenia may not access treatment. Descriptions of service utilisation have tended to assume that biomedical care is the 'final destination' in the treatment journey [42]. Labys et al. consulted 83 stakeholders (traditional healers, religious leaders and caregivers) in rural KwaZulu Natal, South Africa, to gain richer insights into patterns of help-seeking [12]. Nearly half of the people with psychosis discussed had never accessed biomedical care, whilst half had accessed a traditional healer. In some cases care was sought in parallel. In qualitative studies in Ethiopia and Tanzania, people with schizophrenia, caregivers and healthcare workers reported substantial barriers to engagement with care [43, 44]. In both settings there were erratic anti-psychotic medication supplies and difficulties paying for medication, due to poverty. In Tanzania whilst free mental health services are mandated, this is not usually the case in practice [44]. Other reasons for disengagement were the long illness course and unmet expectations of cure. Differing explanatory models were rarely noted as reasons for disengagement [43].

Psychosocial interventions

Previous research has demonstrated the benefits of family and psychosocial interventions for people with schizophrenia in LMIC, particularly in China [45]. Current developments include a greater emphasis on feasibility, such as delivery by non-specialists [46, 47] and peers [48], and a growing interest in rehabilitation-focused interventions [46, 47]. Using a pretest-posttest design, a 12-week family-to-family psychoeducation intervention in Mexico City produced small reductions in expressed emotion amongst caregivers of people with schizophrenia (n=230 family members) [48]. In Jordan, Al Hadihasan et al. found that even an extremely low-intensity intervention comprising psychoeducational leaflets with follow-up phone calls is acceptable and may impact on patient outcomes through improving knowledge, coping skills and self-confidence [49]. Participants in a pilot of group psychosocial rehabilitation in South Africa likewise reported improvements in knowledge and self-esteem, and also social support and self-care. However, a substantial acceptability issue was the absence of income-generating opportunities [46]. Delivery by auxiliary social workers, a non-specialist cadre, was found to be feasible [46]. Results from the Rehabilitation Intervention for people with Schizophrenia in Ethiopia clusterrandomised trial, will provide insights into the effectiveness of community-based rehabilitation delivered by lay health workers [47].

mHealth approaches offer exciting scope for intervention scalability. An RCT in rural China will evaluate the effectiveness of a multicomponent intervention comprising medication reminders via mobile text and voice messaging, an award system to incentivise medication adherence and support for family members [50]. A pilot study set in Cape Town assessed the feasibility of a single psychoeducation session and text message appointment reminders amongst people with severe mental illness (n=77). Substantial problems were identified; 41.2% did not receive the text messages as phones were lost, stolen or numbers were changed [51]. Although it was not explored in this paper, it is conceivable that difficulties using electronic devices may disproportionately affect the poorest. This pilot therefore highlights that an unintended negative consequence of mHealth interventions may be to widen health inequalities.

A recent RCT has demonstrated the effectiveness of brief culturally-adapted CBT delivered by community clinicians in improving outcomes in people with schizophrenia in China. Guo et al. found a clinically significant improvement in symptoms in 37.3% in the CBT plus treatment as usual (TAU) arm vs 19.1% in TAU alone arm (p=0.003)[52]. RCTs in Pakistan and China also demonstrated the effectiveness of CBT delivered by psychiatrists and psychologists in hospital settings [53, 54]. Qualitative studies in China and Pakistan have explored potential cultural adaptations to CBT for non-Western settings, including a bio-psycho-spiritual-social model for conceptualising and managing psychosis and the close involvement of family members [55, 56].

Traditional, faith and alternative medicine

Nortje et al.'s systematic review examined evidence for the effectiveness of any type of traditional healing (including holy water and herbal remedies) across HICs and LMICs. Evaluations were generally of poor quality, included mixed psychiatric diagnoses, and utilised non-randomised designs [57]. The authors concluded that whilst acute relapses of schizophrenia may improve whilst individuals are under the care of traditional healers, these improvements cannot be distinguished from the natural illness course. One included study indicated an association between religious healing and relapse, though reverse causality could not be ruled out [57]. A large effect of adjunctive Tai Chi on negative symptoms of schizophrenia (five RCTs; n=451) was found in a systematic review and meta-analysis (Standardised Mean Difference 0.87 (95% CI -1.51, -0.24), p=0.007) [58]. A Cochrane Review concluded Wendan decoction (a Chinese herb formula) may produce short-term improvements in psychotic symptoms compared to placebo, and whilst the formula is no better than anti-psychotic medication in improving mental state it has fewer side effects [59]. However, the quality of evidence was low in both reviews [58, 59].

There are few evaluations of biomedical-traditional provider collaborations. Van der Watt's qualitative study provides fascinating insights into the views of biomedical and alternative mental healthcare providers on potential collaboration in Ghana, Kenya and Nigeria [60]. Several barriers were identified, including a mutually strong sense of distrust towards, and superiority over, the other provider groups. The incompatibly of different belief systems and the perceived intention of biomedical providers to siphon expertise from traditional healers without acknowledgement or compensation, were also cited [60]. The COllaborative Shared care to Improve Psychosis Outcome RCT will compare the effectiveness of a collaborative care programme with usual care at improving symptom severity in people with psychosis in Nigeria and Ghana [61].

Conclusion

Social and cultural contexts continue, rightly, to be considered important forces shaping the illness experience of schizophrenia. Yet there is increasing appreciation of universal influences on schizophrenia outcomes and the problems associated with the illness, such as physical co-morbidities. Acceptable and effective responses to schizophrenia in LMIC should be cognisant of both sets of factors. Efforts to enhance the quality of family support should also be central to models of care, given the influence of the family in outcomes, and their typical role as primary care provider. Learning in this area may have wider relevance to HIC. Alternative treatment approaches such as Tai Chi may be a vehicle for the shift of global mental health from 'delivery science to discovery science' [62]. Some topics have received little attention in LMIC including involuntary treatment, early intervention and how to address excess mortality. The evolution of global mental health into a truly global discipline will benefit people with schizophrenia in low, middle and high-income country settings [62].

Key points

• There is a plurality of explanatory models for psychosis across settings and context influences experiences of stigma. People with schizophrenia in LMIC are vulnerable to food insecurity, violence and physical health problems.

- Being married or having close family may be associated with better outcomes in LMIC. However social support is not always available and caregiving is challenging.
- Evidence-based care remains inaccessible to many people with schizophrenia. Psychosocial support groups appear to be acceptable and culturally-adapted cognitive-behavioural therapy can reduce symptom severity.
- Most research on traditional and alternative treatments for schizophrenia is of low quality, but some modalities, such as Tai Chi, may be beneficial. The challenges of biomedical-traditional provider collaborations have been highlighted and forthcoming evaluations will evaluate their effectiveness.

Financial support and sponsorship

AF is funded by the Medical Research Council and Department for International Development through the African Research Leader Scheme

Conflicts of interest

LA and AF are co-principal investigators, and CH is co-investigator, of the RISE trial. CH is principal investigator, and AF is co-investigator, of the TASCs trial. CH is the research director of PRIME and AF is the principal investigator of PRIME Ethiopia.

References

- 1. Collins, P.Y., V. Patel, S.S. Joestl, et al., *Grand challenges in global mental health.* Nature, 2011. **475**(7354): p. 27-30.
- 2. Patel, V., *Universal health coverage for schizophrenia: A global mental health priority.* Schizophrenia Bulletin, 2016. **42**(4): p. 885-890.
- 3. Lora, A., R. Kohn, I. Levav, et al., *Service availability and utilization and treatment gap for schizophrenic disorders: a survey in 50 low- and middle-income countries.* Bull World Health Organ, 2012. **90**(1): p. 47-54, 54a-54b.
- 4. Keynejad, R.C., T. Dua, C. Barbui, et al., *WHO Mental Health Gap Action Programme (mhGAP) Intervention Guide: a systematic review of evidence from low and middle-income countries.* Evid Based Ment Health, 2017.
- 5. Gureje, O., G. Nortje, V. Makanjuola, et al., *The role of global traditional and complementary systems of medicine in the treatment of mental health disorders.* Lancet Psychiatry, 2015. **2**(2): p. 168-77.
- 6. Jablensky, A., N. Sartorius, G. Ernberg, et al., *Schizophrenia: manifestations, incidence and course in different cultures. A World Health Organization tencountry study.* Psychol Med Monogr Suppl, 1992. **20**: p. 1-97.
- Cohen, A., V. Patel, R. Thara, et al., *Questioning an Axiom: Better Prognosis for Schizophrenia in the Developing World?* Schizophrenia Bulletin, 2008.
 34(2): p. 229-244.
- 8. Kebede, D., A. Alem, T. Shibre, et al., *Short-term symptomatic and functional outcomes of schizophrenia in Butajira, Ethiopia.* Schizophr Res, 2005. **78**(2-3): p. 171-85.
- 9. Cohen, A., R. Padmavati, M. Hibben, et al., *Concepts of madness in diverse settings: A qualitative study from the INTREPID project.* BMC Psychiatry Vol 16 2016, ArtID 388, 2016. **16**.
- *10. Angermeyer, M.C., M.G. Carta, H. Matschinger, et al., *Cultural differences in stigma surrounding schizophrenia: Comparison between Central Europe and North Africa.* The British Journal of Psychiatry, 2016. **208**(4): p. 389-397.

Large cross-sectional community survey demonstrating how local cultural values may shape expression of public stigma towards people with schizophrenia in different settings

- 11. Campbell, M.M., G. Sibeko, S. Mall, et al., *The content of delusions in a sample of South African Xhosa people with schizophrenia.* BMC Psychiatry Vol 17 2017, ArtID 41, 2017. **17**.
- 12. Labys, C.A., E. Susser, and J.K. Burns, *Psychosis and help-seeking behavior in rural KwaZulu Natal: Unearthing local insights.* International Journal of Mental Health Systems Vol 10 2016, ArtID 57, 2016. **10**.
- Makanjuola, V., Y. Esan, B. Oladeji, et al., *Explanatory model of psychosis: impact on perception of self-stigma by patients in three sub-saharan African cities.* Social Psychiatry & Psychiatric Epidemiology, 2016. **51**(12): p. 1645-1654.
- *14. Koschorke, M., R. Padmavati, S. Kumar, et al., *Experiences of stigma and discrimination faced by family caregivers of people with schizophrenia in India.* Social Science & Medicine, 2017. **178**: p. 66-77.

In-depth mixed methods analysis of experiences of stigma by caregivers in India, including detailed conceptual frameworks.

- 15. Yang, L.H., A. Kleinman, B.G. Link, et al., *Culture and stigma: adding moral experience to stigma theory.* Soc Sci Med, 2007. **64**(7): p. 1524-35.
- *16. Tirfessa, K., C. Lund, G. Medhin, et al., *Food insecurity among people with severe mental disorder in a rural Ethiopian setting: a comparative, population-based study.* Epidemiol Psychiatr Sci, 2017: p. 1-11.

Cross-sectional study demonstating higher rates of food insecurity amongst households of people with severe mental disorder compared to the general population, including an exploration of the mediating role of disability.

- 17. Afe, T.O., T.C. Emedoh, O. Ogunsemi, et al., *Intimate partner violence, psychopathology and the women with schizophrenia in an outpatient clinic South-South, Nigeria.* BMC Psychiatry Vol 16 2016, ArtID 197, 2016. **16**.
- 18. International, N.P.C.N.N.a.I. Nigeria Demographic and Health Survey 2013. 2014.
- 19. Asher, L., A. Fekadu, S. Teferra, et al., *"I cry every day and night, I have my son tied in chains": physical restraint of people with schizophrenia in community settings in Ethiopia.* Global Health, 2017. **13**(1): p. 47.
- **20. Stubbs, B., A. Koyanagi, N. Veronese, et al., *Physical multimorbidity and psychosis: Comprehensive cross sectional analysis including 242,952 people across 48 low- and middle-income countries.* BMC Medicine, 2016. **14 (1) (no pagination)**(189).

Although limited by using self-reported, rather than clinician-diagnosed, psychosis, this very large scale cross-sectional survey is an important contribution to the sparse literature on physical health outcomes in people with schizophrenia in LMIC.

- 21. Ganesh, S., A.H. Ashok, C.N. Kumar, et al., *Prevalence and determinants of metabolic syndrome in patients with schizophrenia: A systematic review and meta-analysis of Indian studies.* Asian Journal of Psychiatry, 2016. **22**: p. 86-92.
- Liu, N.H., G.L. Daumit, T. Dua, et al., *Excess mortality in persons with severe mental disorders: a multilevel intervention framework and priorities for clinical practice, policy and research agendas.* World Psychiatry, 2017. 16(1): p. 30-40.
- 23. Sugawara, N., K. Maruo, T. Sugai, et al., *Prevalence of underweight in patients with schizophrenia: A meta-analysis.* Schizophr Res, 2017.
- 24. Berrington de Gonzalez, A., P. Hartge, J.R. Cerhan, et al., *Body-mass index and mortality among 1.46 million white adults.* N Engl J Med, 2010.
 363(23): p. 2211-9.
- 25. Ran, M.-S., C.H. Chui, I.Y.-L. Wong, et al., *Family caregivers and outcome of people with schizophrenia in rural China: 14-year follow-up study.* Social Psychiatry and Psychiatric Epidemiology, 2016. **51**(4): p. 513-520.
- **26. Ran, M.-S., Y.-L.I. Wong, S.-Y. Yang, et al., *Marriage and outcomes of people with schizophrenia in rural China: 14-year follow-up study.* Schizophrenia Research, 2017. **182**: p. 49-54.

One of a number of papers reporting outcomes of this cohort study set in rural China, which is one of the longest running cohorts of people with schizophrenia in any LMIC.

- 27. Ran, M.-S., L.H. Yang, Y.-J. Liu, et al., *The family economic status and outcome of people with schizophrenia in Xinjin, Chengdu, China: 14-year follow-up study.* International Journal of Social Psychiatry, 2017. **63**(3): p. 203-211.
- 28. Sadiq, S., K. Suhail, J. Gleeson, et al., *Expressed emotion and the course of schizophrenia in Pakistan.* Social Psychiatry and Psychiatric Epidemiology, 2017. **52**(5): p. 587-593.
- 29. den Hertog, T.N. and A.R. Gilmoor, *Informal care for people with chronic psychotic symptoms: Four case studies in a san community in South Africa.* Health & Social Care in the Community, 2017. **25**(2): p. 538-547.
- *30. Sibeko, G., P.D. Milligan, H. Temmingh, et al., *Caregiving for mental health service users: A study exploring the perceptions of mental health service users and their caregivers in Cape Town, South Africa.* Int J Soc Psychiatry, 2016. **62**(6): p. 512-21.

Qualitative study highlighting the substantial challenges faced by caregivers of people with schizophrenia.

- 31. Wusten, C. and T.M. Lincoln, *The association of family functioning and psychosis proneness in five countries that differ in cultural values and family structures.* Psychiatry Research, 2017. **253**: p. 158-164.
- 32. Jaya, E.S., L. Ascone, and T.M. Lincoln, *Social adversity and psychosis: The mediating role of cognitive vulnerability.* Schizophrenia Bulletin, 2017.
 43(3): p. 557-565.
- **33. Jaya, E.S., L. Ascone, and T.M. Lincoln, *A longitudinal mediation analysis of the effect of negative-self-schemas on positive symptoms via negative affect.* Psychol Med, 2017: p. 1-11.

Although limited by the use of an online survey and self-reported psychosis, this exploration of cognitive mechanisms relating to psychosis in a lower middle-income country (Indonesia), as well as high-income countries (United States and Germany), is a novel contribution to the literature. The identification of universal mechanisms provides an interesting contrast to the literature highlighting the importance of context.

34. Jordans, M., L. Aldridge, N.P. Luitel, et al., *Evaluation of outcomes for psychosis*

and epilepsy treatment delivered by primary

- *health care workers in Nepal: a cohort study.* International Journal of Mental Health Systems, 2017. **11**: p. 70.
- 35. Hanlon, C., A. Alem, G. Medhin, et al., *Task sharing for the care of severe mental disorders in a low-income country (TaSCS): study protocol for a randomised, controlled, non-inferiority trial.* Trials, 2016. **17**(1): p. 76.
- 36. De Silva, M.J., S.D. Rathod, C. Hanlon, et al., *Evaluation of district mental healthcare plans: the PRIME consortium methodology.* British Journal of Psychiatry, 2016. **208 Suppl 56**: p. s63-70.

- 37. Mayston, R., K. Habtamu, G. Medhin, et al., *Developing a measure of mental health service satisfaction for use in low income countries: a mixed methods study.* BMC Health Services Research, 2017. **17**(1): p. 183.
- 38. Mall, S., M. Hailemariam, M. Selamu, et al., '*Restoring the person's life': a qualitative study to inform development of care for people with severe mental disorders in rural Ethiopia.* Epidemiol Psychiatr Sci, 2016: p. 1-10.
- 39. Murthy, P., M. Isaac, and H. Dabholkar, *Mental Hospitals in India in the 21st century: Transformation and relevance.* Epidemiology and Psychiatric Sciences, 2017. **26**(1): p. 10-15.
- 40. Funk, M. and N. Drew, *WHO QualityRights: transforming mental health services.* Lancet Psychiatry, 2017. **4**(11): p. 826-827.
- 41. Shah, S., N. Desai, S. Shah, et al., *Impact of Quality Rights Gujarat program* on dropout rate of patients visiting outpatient psychiatry department of tertiary care hospital. Asian J Psychiatr, 2017. **28**: p. 4-8.
- 42. Ibrahim, A., S. Hor, O.S. Bahar, et al., *Pathways to psychiatric care for mental disorders: A retrospective study of patients seeking mental health services at a public psychiatric facility in Ghana.* International Journal of Mental Health Systems, 2016. **10 (1) (no pagination)**(63).
- 43. Hailemariam, M., A. Fekadu, M. Prince, et al., *Engaging and staying engaged: a phenomenological study of barriers to equitable access to mental healthcare for people with severe mental disorders in a rural African setting.* Int J Equity Health, 2017. **16**(1): p. 156.
- *44. Iseselo, M.K. and J.S. Ambikile, *Medication challenges for patients with severe mental illness: Experience and views of patients, caregivers and mental health care workers in Dar es Salaam, Tanzania.* International Journal of Mental Health Systems Vol 11 2017, ArtID 17, 2017. **11**.

Qualitative study from Tanzania exploring the critical issue of availability and affordability of psychotropic medications. The substantial barriers to equitable access identiifed demonstrate the importance of considering contact coverage and effective coverage, rather than simply availability coverage, in evaulations of the scale up of mental healthcare in LMIC.

- 45. Asher, L., V. Patel, and M.J. De Silva, *Community-based psychosocial interventions for people with schizophrenia in low and middle-income countries: systematic review and meta-analysis.* BMC Psychiatry, 2017. **17**(1): p. 355.
- 46. Brooke-Sumner, C., C. Lund, O. Selohilwe, et al., *Community-based psychosocial rehabilitation for schizophrenia service users in the north west province of South Africa: A formative study.* Social Work in Mental Health, 2017. **15**(3): p. 249-283.
- 47. Asher, L., M. De Silva, C. Hanlon, et al., *Community-based Rehabilitation Intervention for people with Schizophrenia in Ethiopia (RISE): study protocol for a cluster randomised controlled trial.* Trials, 2016. **17**(1): p. 299.
- 48. Dominguez-Martinez, T., M.L. Rascon-Gasca, H. Alcantara-Chabelas, et al., *Effects of family-to-family psychoeducation among relatives of patients with severe mental disorders in Mexico City.* Psychiatric Services, 2017. **68**(4): p. 415-418.

- 49. Al-HadiHasan, A., P. Callaghan, and J.S. Lymn, *Qualitative process evaluation of a psycho-educational intervention targeted at people diagnosed with schizophrenia and their primary caregivers in Jordan.* BMC Psychiatry Vol 17 2017, ArtID 68, 2017. **17**.
- 50. Xu, D.R., W. Gong, E.D. Caine, et al., *Lay health supporters aided by a mobile phone messaging system to improve care of villagers with schizophrenia in Liuyang, China: protocol for a randomised control trial.* BMJ Open, 2016. **6**(1): p. e010120.
- 51. Sibeko, G., H. Temmingh, S. Mall, et al., *Improving adherence in mental health service users with severe mental illness in South Africa: a pilot randomized controlled trial of a treatment partner and text message intervention vs. treatment as usual.* BMC Res Notes, 2017. **10**(1): p. 584.
- **52. Guo, Z.H., Z.J. Li, Y. Ma, et al., *Brief cognitive-behavioural therapy for patients in the community with schizophrenia: randomised controlled trial in Beijing, China.* British Journal of Psychiatry, 2017. **210**(3): p. 223-229.

This paper stands out from other recent RCTs and pilot studies of CBT for schizophrenia in LMIC, as the intervention was delivered in a community setting.

- 53. Naeem, F., S. Saeed, M. Irfan, et al., *Brief culturally adapted CBT for psychosis (CaCBTp): A randomized controlled trial from a low income country.* Schizophr Res, 2015. **164**(1-3): p. 143-8.
- 54. Li, Z.J., Z.H. Guo, N. Wang, et al., *Cognitive–behavioural therapy for patients* with schizophrenia: a multicentre randomized controlled trial in Beijing, *China.* Psychological Medicine, 2015. **45**(9): p. 1893-1905 13p.
- 55. Naeem, F., N. Habib, M. Gul, et al., *A qualitative study to explore patients', carers' and health professionals' views to culturally adapt CBT for psychosis (CBTp) in Pakistan.* Behavioural and Cognitive Psychotherapy, 2016. **44**(1): p. 43-55.
- 56. Li, W., L. Zhang, X. Luo, et al., *A qualitative study to explore views of patients', carers' and mental health professionals' to inform cultural adaptation of CBT for psychosis (CBTp) in China.* BMC Psychiatry Vol 17 2017, ArtID 131, 2017. **17**.
- *57. Nortje, G., B. Oladeji, O. Gureje, et al., *Effectiveness of traditional healers in treating mental disorders: a systematic review.* Lancet Psychiatry, 2016.
 3(2): p. 154-70.

Ambitious review covering evaluations of traditional medicine for a range of mental disorders across low, middle and high income countries. Although there are few randomised evaluations the authors draw out some interesting insights as to the potential role of traditional healers for the care of people with schizophrenia.

- 58. Zheng, W., Q. Li, J. Lin, et al., *Tai Chi for schizophrenia: A systematic review.* Shanghai Archives of Psychiatry, 2016. **28**(4): p. 185-194.
- 59. Deng, H. and J. Xu, *Wendan decoction (Traditional Chinese medicine) for schizophrenia.* Cochrane Database of Systematic Reviews, 2017. **6**: p. CD012217.

*60. van der Watt, A.S.J., G. Nortje, L. Kola, et al., *Collaboration Between Biomedical and Complementary and Alternative Care Providers: Barriers and Pathways.* Qual Health Res, 2017. **27**(14): p. 2177-2188.

This qualitative study provides a fascinating account of the potential barriers to biomedical-provider collaboration in mental healthcare in LMIC, and recommends a range of approaches to address the identified issues.

- 61. Gureje, O., V. Makanjuola, L. Kola, et al., *COllaborative Shared care to IMprove Psychosis Outcome (COSIMPO): study protocol for a randomized controlled trial.* Trials, 2017. **18**(1): p. 462.
- 62. Patel, V., *From delivery science to discovery science: Realising the full potential of global mental health.* Epidemiology and Psychiatric Sciences, 2016. **25**(6): p. 499-502.
 - •