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Original article

Predictors of functional status at service entry and discharge among young people with first episode psychosis

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<u>Abstract</u>

Objective: Most patients with first episode psychosis (FEP) are neither studying nor employed (have a poor functional status) when first accessing care. Knowledge of the characteristics of patients with poor functioning and the features influencing functional status over time may pave the way to better treatment.

Method: A medical file audit was used to collect data on premorbid, entry, treatment and 18-month outcome characteristics on 661 FEP patients who consecutively attended the Early Psychosis Prevention and Intervention Centre (EPPIC), Melbourne, Australia, between 1998 and 2000. Functional status was ascertained using the Modified Vocational Status Index (MVSI) and was rated at baseline (poor or good) and according to its evolution over the treatment period (stable good, stable poor, deteriorating or improved functional status).

Results: 52.0% of patients had a poor functional status at service entry. They were more likely to be male with a non-affective psychosis. They also had lower levels of premorbid global functioning and education, and were more likely to have self-reported histories of learning disability, forensic issues, traumatic experiences, and substance use. At service entry, they had more severe symptoms and poorer global functioning. 37% of these patients maintained a poor functional status at discharge, and 18% of those with a good functional status at service entry experienced a decline.

Conclusions: Although psychosocial interventions might assist a young person with FEP with working towards functional goals, for some, the impact of factors such as ongoing substance use and forensic issues on functional status need to be addressed.

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Introduction

Unemployment and/or disruption to educational goals are prevalent and debilitating issues affecting young people with first episode psychosis (FEP) [1,2]. Young people with psychosis are more likely to have disrupted and less education than their peers, which can impede future employment opportunities [3]. Rates of unemployment among FEP groups range from 13-55% [1], with rates increasing to 75-95% within the first couple of years after diagnosis [4]. This poor functioning and lack of educational and employment attainment may result in a downward spiral towards poverty and social isolation [5]. In order to effectively target and improve psychosocial interventions, it is important to first understand the characteristics of FEP patients who are either unemployed or not studying (have a poor functional status) at service entry. Only a few such studies have been conducted with FEP and most have focused on employment. Those unemployed when they first present for treatment for psychosis lack educational attainment [5-8], have poorer premorbid adjustment [9], more severe symptomatology [9-11], longer duration of untreated psychosis [DUP,8,9], dependence on disability benefits [5], poorer cognitive functioning [10,12], substance use [12], and lower guality of life [QoL, 9]. FEP patients who are students and/or employed have lower levels of depressive symptoms and higher global and social functioning [13]. Previous studies, however, have been based on small unrepresentative samples with sample sizes confined to under 200 participants [e.g., 9,11,14,13], with predominant features including non-affective [e.g., 7] or inpatient samples [e.g., 11].

Recent reports have confirmed the importance of combining early intervention programs with vocational and psychosocial support [1,2]. Findings suggest that clinical services

alone produce less favourable functional outcomes in their patients [2,15]. However, improved employment outcomes (albeit not improved educational outcomes) have been observed in studies of early intervention programs additionally providing evidence-based vocational services [2,16]. Most, however, have not considered predictors of changes in functional status after receiving treatment from a specialised early intervention service. Understanding predictors of not working/studying, particularly those, which are malleable, will enhance targeted vocational and psychosocial interventions.

This is a unique study in terms that data were collected prior to the introduction of a formal vocational intervention, namely individual placement support (IPS), to the Early Psychosis Prevention and Intervention Service [EPPIC, 3,17], Melbourne Australia. It is a naturalistic observational study with three aims: (i) to delineate the pre-morbid, diagnostic and clinical characteristics of FEP patients with a poor functional status (either unemployed or not studying) at entry to a specialist early intervention service; (ii) to examine changes in functional status over an 18-month treatment period at the service; and (iii) to identify predictors of change in functional status.

<u>Method</u>

Sample and setting

The sample consisted of 786 youth registered at the Early Psychosis Prevention and Intervention Centre (EPPIC) between 1998 and 2000. EPPIC is a specialist early intervention service for youth experiencing FEP. Participants were aged between 15 and 29 years, residing in the north-western suburbs of Melbourne, Australia and treated for up to 18 months [18]. This cohort was unique in that there were few other treatment facilities or private psychiatrists who serviced this catchment area at the time; thus this is a treated epidemiological sample from a defined catchment area.

Between 1998 and 2000, IPS had not commenced at EPPIC and there was no employment consultant within the service. Young people would have access to medical review, clinical case management and may have been referred by case managers to other services to address issues such as housing, welfare, or employment. Young people had the opportunity to access group programs at EPPIC that included some vocationally-orientated groups [19]. At the time, young people could access disability support and unemployment benefits from external agencies.

A file audit was conducted on these patients as part of the First Episode Psychosis Outcomes Study [FEPOS, 20]. Of the 786, 82 were discharged or transferred to other services and their medical files were unavailable for review, and a further 43 had a nonpsychotic diagnosis at discharge. Data were therefore available for 661 patients (see Figure 1). The local research and ethics committee approved the study.

(Insert Figure 1 about here)

Materials

To systematically assess consecutive medical files we used the Early Psychosis File Questionnaire [20]. The measures included in this questionnaire are detailed below.

Functional status

The Modified Vocational Status Index [MVSI, 21] was used to measure functional status, rating individuals on one of seven occupational levels: (1) full-time gainful employment (≥30 hours/week); (2) homemaker or student; (3) part-time gainful employment; (4) retired; (5) full- or part-time volunteer; (6) on medical or psychiatric leave of absence; and (7) unemployed, whether or not expected to work. Good functioning status was defined as full-time employment, homemaker or student, part-time gainful employment or full- or part-time volunteer. Poor functioning status was defined as either being on medical or psychiatric leave, or unemployment. Individuals could only fall into one of these 7 categories of the MVSI.

Diagnosis

Intake diagnoses were primarily performed by well-trained EPPIC clinicians working in a specialised assessment and crisis intervention team (Conus et al., 2007). Two research psychiatrists (ML and PC) reviewed all the information in the medical file pertaining to diagnosis. A consensus rating between both research psychiatrists and the case manager were performed. For a subset of 115 randomly selected patients, SCID-I/P diagnoses were available and were used to determine the validity of FEPOS discharge diag-

noses (Conus et al., 2007). There was good concordance for both psychotic (κ = 0.80) and substance use (κ = 0.74) diagnoses (Conus et al., 2007).

Pre-treatment characteristics

Characteristics of patients prior to service entry included: pre-morbid functioning in the year preceding illness onset using the Global Assessment of Functioning Scale (GAF); previous history and family history of psychiatric disorder (using DSM-IV-TR criteria); age of onset (when first experienced positive symptoms for longer than one week); DUP (age of entry into EPPIC minus age when first experienced positive psychotic symptoms for longer than one week); past substance use disorders ((SUD), using the Drug and Alcohol Assessment Schedule [DASS, 22]); suicide attempts (ICD-10 categories); exposure to trauma [physical and/or sexual; 23,24]; years in school; and forensic history [offense against property, motor traffic, drug-related offenses, offenses against persons;25].

Service entry

Characteristics assessed at service entry included: type and severity of substance use (DASS); severity of illness (using the Clinical Global Impressions – Severity of illness [CGI-S, 26]) and - Bipolar scales [CGI-BP, 27]; functioning (GAF); insight; and history of a learning disability [20].

Treatment and service discharge characteristics

Treatment factors included: illness severity (CGI-S and CGI-BP); type and severity of substance use disorders (DASS); level of functioning (GAF); involvement with the ser-

vice or disengagement, defined as the patient's refusal of treatment or the service being unable to contact the patient [28]; medication non-adherence, defined as not taking medication for one week or more [29]; and hospital admissions. At discharge, illness severity (CGI-S); global functioning (GAF), and persistent substance use were assessed.

Reliability and validity

Estimates obtained for inter-rater reliability for the CGI, CGI-BP, GAF, and insight for 40 files were good (range: $ICC_{2,1}=0.87$ for CGI-S to $ICC_{2,1}=0.89$ for insight score) [20].

Data Analyses

A series of logistic regression models determined which premorbid and service entry variables were associated with functional status at entry. From these analyses, unadjusted odds ratios (*OR*) and the 95% confidence intervals (*CI*) of the *ORs* were derived. The Wald statistic (*z*) was used to determine significance of individual predictors. From these models, variables that were predictive of good or poor functional status at service entry at the *p*<.10 level were entered in a forward stepwise fashion (based on the Wald statistic) into a multivariate logistic regression model; this model allowed determination of which variables best differentiate good or poor functional status at service entry.

We also examined predictors of change in functional status over the course of treatment. Four categories were developed based on MVSI ratings at entry and discharge: (i) stable good (studying and/or working at entry and discharge); (ii) improved (not studying, working, homemaker, or volunteer at entry but were at discharge); (iii) decline (studying, working, homemaker or volunteer and/or working at entry but not at discharge); and (iv) stable poor (not studying, working, homemaker or volunteer at entry or discharge). A series of logistic regression models were developed to delineate individual variables that discriminated between: (i) stable good versus stable poor functional status: (ii) stable good versus those who had a decline in functional status; and (iii) stable poor versus those who improved in functional status. From these analyses, adjusted odds ratios (*OR*) and the 95% confidence intervals (*CI*) of the adjusted *ORs* were derived. Again, the Wald statistic (*z*) was used to determine significance of individual predictors. Variables that were related to the outcome variables at the p<.01 level were then entered into three separate multivariate stepwise (forward based on Wald statistic) logistic regression models to determine the best predictors associated with change in functional status.

<u>Results</u>

At service entry, data on functional status were available for 659 patients (see Figure 1). Overall, 47.8% (n=316) had a good functional status at service entry according to the MVSI, meaning they were engaged in some functional activity including 20.6% (n=136) who had full-time employment and 21.5% (n=142) who were homemakers/students; 5.5% (36) with part-time employment, and 0.3% (n=2) volunteers. 52.0% (n=343) of the cohort were not studying or unemployed at service entry (poor functional status).

Comparison to population data

Unemployment rates in the cohort were compared to Australian Bureau of Statistics (ABS 2001) Census data (Table 1). In the 15-19 year age range, 49.0% of the cohort was unemployed as compared to 18.7% in this age range in the general community. Notably, as age of the cohort increased, rates of unemployment increased while the converse was true in the general population.

(Insert Table 1 about here)

Predictors of functional status at service entry

Univariate comparisons indicated that poor functional status at service entry was associated with male gender, fewer years of education, poor premorbid functioning as determined by the GAF, and histories of learning disabilities, substance use, forensic issues, sexual and/or physical trauma and a longer DUP (Table 2). Patients with poor functional status were also more likely to have an intake diagnosis of schizophrenia, a substance or poly-substance use disorder, more severe levels of psychotic symptomatology, poor insight into illness, and poorer global functioning at presentation to the service. In a multivariate logistic regression model, there were 7 predictors of poor functional status at service entry: fewer years of education, poorer premorbid functioning, past history of substance use, intake diagnosis of schizophrenia (compared to schizoaffective and bipolar disorders)¹, polysubstance use at service entry, and insight at entry (Table 3). The comparison of these predictors against the constant-only model was statistically significant, $\chi^2(11)=149.60$, p<.001, indicating that the predictors reliably delineated differences between those with poor and good functional status at service entry; however, the explained variance in functional status was modest with Nagelkerke

¹ Schizophrenia was the reference category in the logistic regression. Those with schizoaffective disorders had significantly better outcomes than those with schizophrenia. The same result was found comparing schizophrenia to bipolar disorder.

 R^2 =.30. Using these variables, the model correctly classified 70.1% of those with a good functional status and 70.7% of those with a poor functional status (overall 70.4% correct classification).

(Insert Tables 2 and 3 about here)

Discharge functional status

Data on functional status at discharge were available for 569 patients (see Figure 1). A series of comparisons were conducted to determine whether those who did and did not have 18-month data differed significantly with respect to premorbid and baseline characteristics. Those who did not have data at 18-months were more likely to have longer DUP (no data available *M*=238.4, *SD*=522.6; data available, *M*=355.5, *SD*=653.16, t(648)=-4.67, p<.001), a past history of substance use (no data available 84.8%, n=78, N=92; data available, 72.4%, n=41, N=569, $\chi^2(1)$ =6.32, p=.012), substance use at service entry (no data available 72.8%, n=67, N=92; data available, 59.6%, n=339, N=569, $\chi^2(1)$ =5.87, p=.015), more severe psychopathology (CGI-S, no data available *M*=5.8, *SD*=0.8; data available, *M*=5.5, *SD*=0.8, t(659)=2.97, p=.003), and lower GAF (no data available *M*=27.0, *SD*=9.0; data available, *M*=33.0, *SD*=9.6, t(657)=5.62, p<.001. Table 4 comprises details on changes in functional status from baseline to discharge.

Overall, 30.4% (n=173) had a stable good functional status from baseline to discharge, 4.1% (n=80) had an improved functional status, 18.3% (n=104) had a decline in functional status, and 37.3% (n=212) had stable poor functional status.

(Insert Table 4 about here)

Stable good versus poor functional status

Multivariate logistic regression indicated that as compared to those with stable good functional status, those who had stable poor functional status were more likely to have lower levels of education, a lower premorbid GAF, longer DUP, and a forensic history (Table 5). At service entry, they were older, were more likely to be polysubstance users, had poor insight, and poor functioning. Persistent substance use through treatment was also problematic. This model was significantly different from the constant-only model, $\chi^2(10)=221.51$, *p*<.001. The Nagelkerke R^2 was .69. Overall, classification success was at 83.0% (good stable functional status 82.5% and poor stable functional status 83.4%).

(Insert Table 5 about here)

Stable good versus decline in functional status

Multivariate logistic regression indicated that those who had a decline in functional status were more likely than those with a stable good functional status to have a significantly longer DUP (see Table 6). At service entry they were more likely to be using cannabis and to have a lower GAF. They had a shorter duration of time in the service, a greater number of admissions and were more likely to be persistent substance users. The model with these variables was significantly better than the constant-only model, $\chi^2(7)=48.64$, *p*<.001, and the Nagelkerke R^2 was .23. Based on these variables, correct classification of those who had a decline in functional status was 43.6% and successful classification of those with a stable good functional status was 87.6% (overall correct classification 70.6%).

(Insert Table 6 about here)

Stable poor versus improved functional status

Multivariate analyses indicated that those who had an improved functional status were more likely than those with a poor stable functional status to have a shorter DUP, be more adherent with treatment, and were less likely to be using substances during treatment (see Table 6). This model was significantly different from the constant-only model, $\chi^2(4)=55.38$, *p*<.001. The Nagelkerke R^2 was modest at .29. The classification rate for those who had improved functional status was 27.9% whereas for those who had a poor stable functional status correct classification was at 90.9% (overall success in classification 74.0%).

Discussion

Previous studies have indicated that psychosis causes significant distress for the young person by disrupting attainment of vocational and educational goals, social relationships and identity formation [30]. However, in our study, there was evidence of functional decline prior to illness onset with disruption to both education and global premorbid functioning being more likely in those who had a poor functional status at service entry. Such findings indicate that for some individuals peer concordant functioning is not established [31].

In our study, just over 50% of the patients were unemployed or not studying at service entry. Unemployment rates were markedly higher than those seen in Australian 2001 Census data. Moreover, rates of unemployment in FEP increased with age, whereas the reverse was true in the general community. This study provides more evidence of poor functional status in a representative sample of FEP patients.

Those who were not employed and/or studying at service entry had a complicated history before accessing EPPIC. Based on univariate analyses, their functional decline appeared related to numerous problems: learning disabilities; exposure to traumatic experiences; forensic issues; a longer DUP; and ongoing substance use. However, multivariate analyses indicated that education, premorbid functioning, and substance use (previous history and polysubstance use at service entry) were the strongest predictors of poor functional status. It may be the case that variables such as learning disabilities and exposure to traumatic experiences may increase the likelihood of educational difficulties and substance use. For example, cognitive problems often occur prior to the onset of clinical symptoms [32]; are observed during prodrome [33] or as early as first grade at school [34]. Poor premorbid academic functioning is associated with impairments in both verbal learning and working memory in individuals with FEP [35]. Because of learning difficulties, rates of secondary school drop-out are much higher in FEP cohorts [7]. Fewer years of formal education results may result in increased risky behaviours such as substance use [7] and criminal activity including violence towards others [36].

Exposure to traumatic experiences such as sexual abuse and/or physical abuse is common prior to the onset of psychotic disorders [37,23,24]. Such exposure has been associated with reduced psychosocial premorbid functioning including decreased educational attainment and poor work attendance [23,37]. This relationship may be con-

founded by increased risk of substance use in FEP patients with a history of trauma [38].

As a consequence of a complex history, it is unsurprising that those who had a poor functional status at service entry had more severe symptomatology, poorer insight, were more likely to be abusing multiple substances, and had poorer ratings of global functioning. 37.3% of those with a poor functional status at service entry still had a poor vocational status at discharge. Poor functional outcomes (compared with stable good outcomes) were associated with lower levels of premorbid functioning, less years of education, a forensic history, longer DUP, more severe symptoms and poorer functioning at service entry and discharge as well as more admissions and no change in substance use during treatment. Additional predictors of a stable poor functional status were a history of physical abuse, substance use or polysubstance use disorder and treatment non-adherence.

Only 30.4% of those FEP patients with a good functional status at service entry retained this status at discharge and only 14.1% achieved an improvement in their functional status at discharge. Predictors of maintenance of good functional level at discharge were virtually polar opposites of those associated with negative outcomes, highlighting potential protective factors and targets for clinical and vocational interventions.

It is worth mentioning that the predictive models pertaining to those who either declined or improved in functioning over the 18 months were less impressive in terms of classification and sensitivity. Statistical power may be one issue; however, it may be the case that factors other than those perused in this study may increase the predictive power of the models. For example, neurocognition and social cognition have been considered important predictors of functioning [39]. It may be the case that those with better neurocognition and social cognition may be more likely to improve over 18 months in terms of functioning. Further work on identifying other variables that may predict those who decline and improve in functioning over the course of treatment is needed.

While specialist early intervention programs result in better functional outcomes compared to standard care [5,40], it is clear that those studying or working at service entry can still experience a decline in functional status during treatment [2,15,16]. In our cohort 18.3% experienced decline in functional status. This may be a reflection that those who have a worsening of symptoms are more likely to decline in function [41].

In line with recent literature [2,15,16] we anticipate that those with a persistent poor functional status or who experience a decline in functional status may benefit from targeted vocational intervention in addition to SEI. Three to four years after this cohort of FEP EPPIC patients were discharged, individual placement support (IPS) was introduced at EPPIC [3,17,42]. IPS is considered suitable for anyone interested in looking for work, regardless of whether the individual is work-ready or symptom-free [17]. IPS involves an individualised approach to vocational support; employment consultants work closely with individuals to assess their interests and preferences with respect to employment. Support workers are integrated within mental health services and offer timeunlimited support, this continuing once a person has achieved the goal of competitive employment. Preliminary support for the effectiveness of IPS for FEP has been noted with respect to employment outcomes [3]. However, there are variables which may impact on the success of IPS interventions including motivation, substance use, and cognitive abilities [43]. The primary limitation of this study was the use of retrospective medical file audit methodology. Numerous problems can exist with this approach including: poor quality of documented information, rater inexperience, lack of inter-rater reliability and poor data validity. Strategies were employed to minimise the impact of these limitations, including double-rating of medical files by two consultant psychiatrists with expert knowledge of EPPIC and the treatment of FEP, determination of inter-rater reliability for clinical and functioning measures, and establishing concurrent validity of psychoses and baseline SUD for a sub-sample of patients [20]. Another limitation of medical file auditing is the lack of more specific assessments (e.g., of neurocognitive deficits). More detailed data were not available on the sociodemographic characteristics of clients and the work and educational statuses of their parents.

An additional key limitation relates to a lack of more specific data regarding concurrent functional activities. A shortcoming of the MVSI is that it only allows for one status to be recorded for each individual. In some cases this may not have accurately reflected circumstances (e.g., a person may have been engaged in both voluntary work and education). Information was also not available on utilisation of vocational group programs or access to external agencies, and with data collected at cross-sectional time-points, information was not available regarding stability of functional status from month to month over the 18-months. There also needs to be consideration that not everyone had 18 month data and those who did not have this data were more likely to have a history of substance use issues, longer DUP, more severe symptomatology and poorer functioning at presentation to the service. By not having data on this subgroup, may mean that the estimates of poor functional outcomes at 18 months are an underestimate. Further,

despite being a longitudinal study, our results do not allow for cause-effect analyses. Contrasting these limitations, the study comprised a large representative sample of patients with FEP.

In summary, a subgroup of FEP patients who are not studying and/or working at service entry appear to have a long history of functional impairment that might be related to exposure to traumatic experiences, learning disabilities, and substance use. These problems were found to be more common in males and individuals with schizophrenia. Such factors may contribute to a delay in receiving treatment for psychosis. Consequently, at service entry, such patients have more severe symptoms and poor insight. Persistent substance use, forensic issues, and lack of educational attainment all might impede future functioning and vocational opportunities. Thus, there is a need to get young people into treatment quickly, build upon engagement and relationships with young people, and offer adjunctive treatments that may address comorbid substance use. Indeed, there is some evidence from longitudinal studies that by reducing substance use in individuals with early psychosis, better functional outcomes are assured [44,45]. Apart from managing symptoms and substance use, vocational and psychosocial support during early intervention is essential [2]; highlighting the importance an integrated model of early intervention.

Importantly, further research is needed to ascertain whether these risk factors for poor outcome are also found for those young people now receiving IPS in early intervention services. Finally, the FEPOS cohort is currently being followed up; through this new study we will be able to map the trajectory of functional outcomes and to determine whether the same risk factors that apply in the short-term, predict poor outcomes over the long-term.

References

1. Rinaldi M, Killackey E, Smith J, Shepherd G, Singh SP, Craig T (2010) First episode psychosis and employment: A review. International Review of Psychiatry 22:148-162

2. Bond GR, Drake RE, Luciano A (2016) Employment and educational outcomes in early intervention programmes for early psy-chosis: a systematic review. Epidemiology and Psychiatric Sciences 24:446-457

3. Killackey E, Jackson HJ, McGorry PD (2008) Vocational intervention in first episode psychosis: individual placement and support versus treatment as usual. British Journal of Psychiatry 193:114-120

4. Marwaha S, Johnson S (2004) Schizophrenia and employment. A review. Social Psychiatry and Psychiatric Epidemiology 39:337-349

5. Dewa CS, Trojanowski L, Cheng C, Loong D (2012) Examining the factors associated with paid employment of clients enrolled in first episode of psychosis programs. Schizophr Res Treatment 2012:739616. doi:10.1155/2012/739616

6. Norman RMG, Malla AK, Manchanda R, Windell D, Harricharan R, Takhar J, Northcott S (2007) Does treatment delay predict occupational functioning in first episode psychosis. Schizophrenia Research 91:259-262

7. Goulding SM, Chien VH, Compton MT (2010) Prevalence and correlates of school drop-out prior to initial treatment of non-affective psychosis: Further evidence suggesting a need for supported education. Schizophr Research 116:228-233

8. Tapfumaneyi A, Johnson S, Joyce J, Major B, Lawrence J, Mann F, Chisholm B, Ra-haman N, Wooley J, H. F (2015) Predictors of vocational activity over the first year in inner-city early intervention in psychosis services. Early Intervention in Psychiatry 9:447-458

9. Turner N, Browne S, Clarke M, Gervin M, Larkin C, Waddington JL, O'Callaghan E (2009) Employment status amongst those with psychosis at first presentation. Social Psychiatry and Psychiatric Epidemiology 44:863-869

10. Chang WC, Man Tang JY, Ming Hui CL, Wa Chan SK, Ming Lee EH, Hai Chen EY (2014) Clinical and cognitive predictors of vocational outcome in first-episode schizophrenia: A prospective 3 year follow-up study. Psychiatry Research Accepted 19 September 2014

11. Ramsay C, Stewart T, Compton MT (2012) Unemployment among patients with newly diagnosed first-episode psychosis: prevalence and clinical correlates in a US sample. Social Psychiatry and Psychiatric Epidemiology 47:797-803

12. Dickerson FB, Stallings C, Origoni A, Boronow JJ, Sullens A, Yolken RH (2007) The association between cognitive functioning and occupational status in persons with a recent onset of psychosis. Journal of Nervous and Mental Disease 195:566-571

13. Tandberg M, Ueland T, Andreassen A, Sundet K, Melle I (2012) Factors associated with occupation and academic status in patients with first-episode psychosis with a particular focus on neurocognition. Social Psychiatry and Psychiatric Epidemiology 47:1763-1773

14. Malla AK, Norman RMG, Manchanda R, Ahmed MR, Scholten D, Harricharan R, Cortese L, Takhar J (2002) One year outcome in first episode psychosis: influence of DUP and other predictors. Schizophrenia Research 54:231-242

15. Dudley R, Nicholson M, Stott P, G S (2014) Improving vocational outcomes of service users in an Early Intervention in Psychosis service. Early Intervention in Psychiatry 8:98-102

16. Drake RE, Bond GR, Goldman HH, Hogan MF, Karakus M (2016) Individual Placement and Support services boost employment for people with serious mental illness, but funding is lacking. Health Affairs 35:1098-1105

17. Killackey E, Allott K, Cotton S, Jackson HJ, Scutella R, Tseng Y-P, Borland J, Proffitt TM, Hunt S, Kay-Lambkin F, Chinnery G, Baksheev G, Alvarez-Jimenez M, McGorry PD (2013) A randomized controlled trial of vocational intervention for young people with first-episode psychois: method. Early Intervention in Psychiatry 7:329-337

18. Schimmelmann BG, Conus P, Cotton SM, Kupferschmid S, McGorry PD, Lambert M (2012) Prevalence and impact of cannabis use disorders in adolescents with early onset first episode psychosis. European Psychiatry 27:463-469

19. Cotton SM, Luxmoore M, Woodhead G, Albiston DD, Gleeson JFM, McGor ry PD (2011) Group programmes in early intervention services. Early Intervention in Psychiatry 5:259-266

20. Conus P, Cotton SM, Schimmelmann BG, McGorry PD, Lambert M (2007) The First-Episode Psychosis Outcome Study: premorbid and baseline characteristics of an epidemiological cohort of 661 first-episode psychosis patients. Early Intervention in Psychiatry 1 (1):191-200

21. Tohen M, Hennen J, Zarate CM, Baldessarini RJ, Strakowski SM, Stoll AL, Faedda GL, Suppes T, Gebre-Medhin P, Cohen BM (2000) Two-year syndromal and functional recovery in 219 cases of first episode major affective disorder with psychotic features. Am J Psychiatry 157 (2):220-228

22. McGorry PD, Copolov DL, Singh BS (1990) Royal Park Multidiagnostic Instrument for Psychosis: Part II. Development, reliability and validity. Schizophrenia Bulletin 16 (3):517-536

23. Conus P, Cotton SM, Schimmelmann BG, McGorry PD, Lambert M (2010) Pre-treatment and outcome correlates of sexual and physical trauma in an epidemiological cohort of first episode psychosis patients. Schizophrenia Bulletin 36:1105-1114

24. Conus P, Cotton SM, Schimmelmann BG, Berk M, Daglas R, McGorry PD, Lambert M (2010) Pretreatment and outcome correlates of past sexual and physical trauma in 118 bipolar I disorder patients with a first episode of psychotic mania. Bipolar Disorders 12:244-252

25. Marion-Veyron R, Lambert M, Cotton SM, Schimmelmann BG, Gravier B, McGorry PD, Conus P (2015) History of offending behavior in first episode psychosis patients: A marker of specific clinical needs and a call for early detection strategies among young offenders. Schizophrenia Research 161:163-168

26. Guy W (1976) ECDEC Assessment manual for psychopharmacology, revised. National Institute of Mental Health, Rockville, MD

27. Spearing MK, Post RM, Leverich GS, Brandt D, Nolen W (1997) Modification of the Clinical Global Impressions (CGI) scale for use in bipolar illness (BP): the CGI-BP. Psychiatry Research 73 (3):159-171

28. Conus P, Lambert M, Cotton S, Bonsack C, McGorry PD, Schimmelmann BG (2010) Rate and predictors of service disengagement in an epidemiological first-episode psychosis cohort. Schizophrenia Research 118:256-263

29. Robinson DG, Woerner MG, Alvir JM, Bilder RM, Hinrichsen GA, Lieberman JA (2002) Predictors of medication discontinuation by patients with first-episode schizophrenia and schizoaffective disorder. Schizophrenia Research 57:209-219

30. Malla AK, Norman R, Joober R (2005) First-episode psychosis, early intervention, and outcome: What have we learned. Canadian Journal of Psychiatry 50 (14):881-891

31. Rabinowitz J, De Smedt G, Harvey PD, Davidson M (2002) Relationship between premorbid functioning and symptom severity as assessed at first episode of psychosis. Amercian Journal of Psychiatry 159:2021-2026

32. Cannon TD, Bearden CE, Hollister JM, Rosso IM, Sanchez LE, Hadley T (2000) Childhood cognitive functioning in schizophrenia patients and their unaffected siblings: a prospective cohort study. Schizophrenia Bulletin 26:379-393

33. Brewer WJ, Francey SM, Wood SJ, Jackson HJ, Pantelis C, Phillips LJ, Yung AR, Anderson VA, McGorry PD (2005) Memory impairments identified in people at ultra-high risk for psychosis who later develop first-episode psychosis. American Journal of Psychiatry 162:71-78

34. Bilder RM, Reiter G, Bates J, Lencz T, Szesko P, Goldman RS, Robinson D, Lieberman JA, Kane JM (2006) Cognitive development in schizophrenia: follow-back from the first episode. Journal of Clinical and Experimental Neuropsychology 28:270-282

35. Rund BR, Melle I, Friis S, Johannesen JO, Larsen TK, Midboe LJ, Opjordsmoen S, Simonsen E, Vaglum P, McGlashan T (2007) The course of neurocognitive functioning in first-episode psychosis and its relation to premorbid adjustment, duration of untreated psychosis, and relapse. Schizophrenia Research 91:132-140

36. Large MM, Nielssen O (2011) Violence in first-episode psychosis: a systematic review and metaanalysis. Schizophrenia Research 125:209-220

37. Ramsay C, Flanagan P, Gantt S, Broussard B, Compton MT (2011) Clinical correlates of maltreatment and traumatic experiences in childhood and adolescence among predominantly African American, socially disadvantaged, hospitalized, first-episode psychosis patients. Psychiatry Research 188:343-349

38. Compton MT, Furman AC, Kaslow NJ (2004) Preliminary evidence of an association between childhood abuse and cannabis dependence among African American first episode schizophrenia-spectrum disorder patients. Drug and Alcohol Dependence 76:311-316

39. Allott K, Cotton S, Chinnery G, Baksheev G, Massey J, Sun P, Collins Z, Barlow E, Broussard C, Wahid T, Proffit T, Jackson H, Killackey E (2013) The relative contribution of neurocognition and social cognition to 6-month vocational outcomes following Individual Placement and Support in first episode psychosis. Schizophrenia Research 150:136-143

40. Garety PA, Craig TKJ, Dunn G, Fornells-Ambrojo M, Colbert S, Rahaman N, Read J, Power P (2006) Specialised care for early psychosis: symptoms, social functioning and patient statisfaction: Randomised controlled trial. British Journal of Psychiatry 188:37-45

41. Menezes NM, Arenovich T, Zipursky RB (2006) A systematic review of longitudinal outcome studies of first-episode psychosis. Psychological Medicine 36:1349-1362

42. Killackey E, Jackson HJ, Gleeson J, Hickie I, McGorry PD (2006) Exciting career opportunity beckons! Early Intervention and vocational rehabilitation in first episode psychosis: employing cautious optimism. Australian and New Zealand Journal of Psychiatry 40:951-960

43. Lehman AF, Goldberg R, Dixon L, McNary S, Postrado L, Hackman A, McDonnell K (2002) Improving employment outcomes for persons with severe mental illnesses. Archives of General Psychiatry 59:165-172

44. Abdel-Baki A, Ouellet-Plamondon C, Salvat E, Grar K, Potvin S (2017) Symptomatic and functional outcomes of substance use disorder persistence 2 years after admission to a first-episode psychosis program. Psychiatry Research 247:113-119

45. Kerfoot KE, Rosenheck RA, Petrakis IL, Swartz S, Keefe RSE, McEvoy JP, Stroup TS (2011) Substance use and schizophrenia: Adverse correlates in the CATIE study sample. Schizophrenia Research 132:177-182



Figure 1. Participant flow

Table 1. Unemployment rates among people with first episode of treated psychosis whowere aged between 15-29 years of age at service entry compared to the data from theABS 2001 Census

Age range	ABS 2001 Census	FEPOS
	(<i>N</i> =254,139)	<i>N</i> =659
15-19 years	18.7	49.0
20-24 years	12.4	52.3
25-34 years	7.2	55.9
Across 15-34 years	9.8	52.0

*Note: ABS, Australian Bureau of Statistics; FEPOS, First Episode Psychosis Outcome

Table 2. Univariate analyses depicting the relationship between poor functional status (not studying, working, homemaker or volunteer) at service entry and pre-treatment, diagnostic and admission variables

		Functional sta	Functional status at service entry			I of OR	
Variables		Good (n=316)	Poor (n=343)	-	LCI	UCI	value
Pre-treatment variables							
Gender %Male	% (n)	60.1 (190)	70.8 (243)	1.61	1.17	2.23	.004
Years in school	% (n)	10.9 (1.4)	10.2 (1.5)	0.68	0.61	0.77	<.001
Pre-morbid GAF	M (SD)	72.6 (9.4)	66.4 (10.8)	0.94	0.93	0.96	<.001
Duration of untreated psychosis (DUP, in days) ^a	M (SD)	175.4 (384.0)	325.7 (648.8)	1.45	1.23	1.69	<.001
Age at onset (years)	M (SD)	21.2 (3.6)	21.3 (3.6)	1.01	0.96	1.05	.797
Past history of suicide attempt (%Yes)	% (n)	11.4 (36)	16.7 (56)	1.56	0.99	2.44	.055
Forensic history (%Yes)	% (n)	38.9 (129)	17.9 (56)	2.92	2.03	4.20	<.001
Past history of sexual trauma	% (n)	13.0 (41)	18.8 (64)	1.56	1.02	2.38	.042
Past history of physical trauma	% (n)	19.6 (62)	31.5 (107)	1.88	1.31	2.70	<.001
Diagnostic variables							
Past history							
Major Depressive Disorder (MDD) (%Yes)	% (n)	30.7 (97)	24.2 (83)	0.72	0.51	1.02	.062
Substance use disorder (SUD) (%Yes)	% (n)	66.5 (210)	81.0 (278)	2.16	1.51	3.09	<.001
Learning disorder (%Yes)	% (n)	0.9 (3)	3.8 (13)	4.11	1.16	14.56	.029
At service entry							
Primary diagnosis							
Schizophrenia ^b	% (n)	34.5 (109)	50.1 (172)				
Schizoaffective Disorder	% (n)	19.0 (60)	18.7 (64)	0.64	0.37	1.13	.002
Schizophreniform	% (n)	10.4 (33)	9.0 (31)	1.02	0.76	1.37	.001
Bipolar Disorder	% (n)	25.0 (79)	11.4 (39)	0.56	0.39	0.81	<.001
MDE with psychotic features	% (n)	2.8 (9)	2.6 (9)	1.06	0.51	2.21	.161
Other ^c	% (n)	8.2 (26)	8.2 (28)	1.25	0.80	1.93	.093
Substance use disorder (SUD) (%Yes)	% (n)	53.8 (170)	68.2 (234)	1.84	1.34	2.53	<.001
Polysubstance	% (n)	4.1 (13)	17.2 (59)	4.84	2.60	9.02	<.001
Cannabis	% (n)	44.6 (141)	49.0 (168)	1.19	0.88	1.62	.263

Service entry variables							
Age at service entry	M (SD)	21.7 (3.6)	22.3 (3.3)	1.05	0.99	1.09	.054
Severity of symptoms at entry							
CGI-S severity score	M (SD)	5.5 (0.8)	5.6 (0.8)	1.21	1.00	1.47	.046
CGI-BP depression	M (SD)	2.1 (1.7)	2.0 (1.6)	0.96	0.87	1.05	.357
CGI-BP mania	M (SD)	2.3 (1.9)	1.7 (1.6)	0.84	0.77	0.91	<.001
Insight at entry (%No)	% (n)	58.4 (184)	65.9 (224)	1.38	1.00	1.89	.049
Functional level at entry							
GAF	M (SD)	33.59 (9.9)	30.87 (9.5)	0.97	0.96	0.99	<.001

Note: MDE, Major Depressive Episode; CGI-S, Clinical Global Impressions - Severity of Illness; CGI-BP, Clinical Global Impressions - Bipolar Disorder; GAF, Global Assessment of Functioning; LCI, Lower Confidence Interval; HCI, Higher Confidence Interval

^a Raw data are presented, however the test statistics were based on log10 (+constant) transformed data because of extreme positive skewness

^b Schizophrenia was the reference category ^c Other diagnoses include delusional disorder (*n*=24), substance-related psychotic disorder (*n*=14), brief psychotic episode (*n*=11) and psychosis not otherwise specified (*n*=5)

		95% CI of OR		р
Variables	UK	LCI	UCI	value
Years in school	0.73	0.63	0.84	<.001
Pre-morbid GAF	0.95	0.94	0.97	<.001
Past history				
Substance use disorder (SUD) (%Yes)	1.94	1.26	3.00	.003
At service entry				
Age	1.12	1.06	1.18	<.001
Primary diagnosis				
Schizophrenia ^c				
Schizoaffective Disorder	0.36	0.18	0.72	.004
Schizophreniform	1.06	0.75	1.50	.734
Bipolar Disorder	0.54	0.34	0.85	.007
MDE with psychotic features	2.79	1.16	6.73	.022
Other	0.95	0.56	1.60	.850
Polysubstance use	4.70	2.11	10.49	<.001
Insight at entry (%No)	1.61	1.09	2.38	<.001

Table 3. Odds ratios derived from a multivariate logistic regression indicating which variables best discriminate between poor and good functional status at service entry

Note: GAF, Global Assessment of Functioning; LCI, Lower Confidence Interval; HCI, Higher Confidence Interval

 a 5 cases were removed as they were multivariate outliers based on standardized residual >3.29 (p<.001)

^b Good functional status is coded as 0 and poor functional status is coded as 1

^c Schizophrenia was the reference category

Functional status at discharge	Functional status at service entry							
C C	Full-time employment ¹	Homemaker or student	Part-time employment	Volunteer	Unemployed or no			
Full-time employment	43.4 (49)	8.7 (11)	25.0 (9)	0.0 (0)	13.0 (38			
Homemaker or student	2.7 (3)	52.4 (66)	5.6 (2)	0.0 (0)	5.1 (15)			
Part-time employment	5.3 (6)	7.9 (10)	36.1 (13)	0.0 (0)	7.2 (21)			
Volunteer	0.9 (1)	0.8 (1)	0.0 (0)	100.0 (2)	2.1 (6)			
Unemployed or not studying	47.7 (54)	30.2 (38)	33.4 (12)	0.0 (0)	72.6 (212			

Table 4. Changes in functional status over time.

 1 % (n) are calculated based on numbers within the baseline functional status

Table 5. Odds ratios derived from a multivariate logistic regression indicating which variables that best discriminate between stable poor and stable good functional status over the 18-months of treatment at EPPIC

	Univariat	e logistic reg	ression esti	mates	Final multivariate logistic regression			
Changes in functional status	OR ^a	95% C	I of OR	р	OR ^a	95% C	l of OR	р
		LCI	UCI	value		LCI	UCI	value
Stable good (n=173) vs. stable poor (n	=212)							
Gender	1.94	1.27	3.00	.002				
Years in school	0.60	0.51	0.70	<.001	0.59	0.44	0.79	<.001
Premorbid GAF	0.91	0.88	0.93	<.001	0.88	0.84	0.93	<.001
Duration of untreated psychosis								
(DUP) ³	1.83	1.48	2.26	<.001	3.99	2.54	6.26	<.001
Past history Major Depressive Disorder (MDD)								
%Yes	0.59	0.37	0.94	.027				
Learning disability %Yes	7.63	0.96	60.79	.055				
Substance use %Yes	2.69	1.69	4.29	<.001				
Forensic history %Yes	4 19	2 51	7.00	< 001	3 44	1 39	8 52	008
Physical abuse %Yes	2 25	1 40	3.63	001	0.11	1.00	0.02	.000
Service entry	2.20		0.00					
Age	1.06	1 00	1 12	061	1 18	1.06	1.32	002
Schizophropic ⁰ /Voc ^c	2.24	1.05	E 64	.001			1.02	.002
	3.31	1.95	0.04 0.40	<.001				
Cannabis use %Yes	1.41	0.94	2.12	.097	5.00	4 4 5	04 70	000
Polysubstance use % res	5.25	2.40	11.5	<.001	5.33	1.15	24.73	.033
	1.49	1.16	1.90	.001				
CGI-BP depression	0.87	0.77	0.99	.036				
CGI-BP mania	0.83	0.74	0.93	.002				
Insight % No	2.08	1.4	3.16	.001	6.85	2.88	16.25	<.001
GAF	0.95	0.93	0.98	<.001	0.94	0.90	0.98	.003
Treatment characteristics								
Time in service	1.00	0.99	1.00	.101				
Number of admissions ^d	1.35	1.15	1.58	<.001				
Non-adherent with treatment %No ^d	2.01	1.29	3.12	.002				

Substance use during treatment ^d								
No change, restart or increase	3.03	2.1	4.37	<.001	4.11	2.00	8.44	<.001
Decrease or stop	0.69	0.51	0.92	.013	0.37	0.21	0.66	.001

Note: GAF, Global Assessment of Functioning; CGI-S, Clinical Global Impressions - Severity of Illness; CGI-BP, Clinical Global Impressions - Bipolar Disorder; LCI, Lower Confidence Interval; HCI, Higher Confidence Interval

^a Stable good functional status is coded as 0 and stable poor functional status is coded as 1 ^b Because of extreme skewness, the logarithmic transformed version of DUP was

used

^c Diagnostic categories were collapsed to schizophrenia versus other ^d Time in service was used as a covariate in univariate

analyses

^e 29 cases were removed as they were multivariate outliers based on standardized residual >3.29 (p<.001)

Table 6. Odds ratios derived from two multivariate logistic regression models indicating which variables that best discriminate the stability of poor or good functional status over the 18-months of treatment at EPPIC

Changes in functional status	Univariat OR	e logistic 95% C	regression e CI of OR	stimates p	nates Final multivariate logis p OR 95% Cl c		stic regress of OR	ion ^d p
		LCI	UCI	value	_	LCI	UCI	value
Stable good (<i>n</i> =173) vs. drop (<i>n</i> =104)								
Years of education	0.84	0.70	1.00	.055				
Premorbid GAF	0.96	0.93	0.98	.001				
Duration of untreated psychosis (DUP) ^a <i>Past history</i>	1.43	1.10	1.85	.007	1.63	1.21	2.21	.002
Forensic history %Yes	2.03	1.09	3.78	.025				
Service entry								
Cannabis use %Yes	1.58	0.97	2.59	.067	3.29	1.30	8.33	.012
CGI-S	1.39	1.02	1.88	.035				
GAF	0.97	0.94	0.99	.013	0.96	0.93	0.99	.030
Insight at entry %No	1.55	0.95	2.56	.082				
Treatment characteristics								
Number of admissions	1.45	1.20	1.76	<.001	1.44	1.16	1.78	.001
Substance use during treatment								
No change, restart or increase	2.17	1.42	3.32	<.001	1.66	1.02	2.69	.042
Decrease or stop	0.66	0.46	0.95	.024	0.43	0.25	0.75	.003
Length of time in service	1.00	0.99	1.00	.230	0.99	0.98	0.99	.007
Stable poor (<i>n</i> =212) vs. improved (<i>n</i> =80) ^e							
Years in school	1.24	1.03	1.49	.026				
Premorbid GAF	1.04	1.01	1.07	.006				
Duration of untreated psychosis (DUP) ^a <i>Past history</i>	0.65	0.50	0.84	.001	0.54	0.39	0.75	<.001

Major Depressive Disorder (MDD)								
%Yes	1.74	0.97	3.09	.061				
Service entry								
Schizophrenia %Yes ^b	0.40	0.21	0.79	.007				
CGI-S at entry	0.69	0.5	0.95	.025				
CGI-BP depression	1.15	0.98	1.35	.090				
Treatment characteristics								
Length of time in service	1.01	1.00	1.02	.015				
Number of admissions	0.80	0.66	0.97	.021				
Noncompliance with treatment	0.56	0.32	0.98	.043	0.53	0.28	0.99	.047
Substance use during treatment								
No change, restart or increase	0.41	0.25	0.66	<.001	0.16	0.07	0.36	<.001
Decrease or stop	1.73	1.20	2.50	.003	2.61	1.57	4.35	<.001

Note: GAF, Global Assessment of Functioning; CGI-S, Clinical Global Impressions - Severity of Illness; CGI-BP, Clinical Global Impressions - Bipo-lar Disorder; LCI, Lower Confidence Interval; HCI, Higher Confidence Interval

^a Because of extreme skewness, the logarithmic transformed version of DUP was used

^b Diagnostic categories were collapsed to schizophrenia versus other

^c Time in service was used as a covariate in univariate analyses

^dThere were no multivariate outliers based on standardized residuals >3.29 (*p*<.001) ^e 8 cases were removed as they were multivariate outliers based on standardized residual >3.29

(*p*<.001)

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