

Pre-morbid IQ in mental disorders: a Danish draft-board study of 7486 psychiatric patients

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Background. Longitudinal studies indicate that future schizophrenia patients exhibit lower IQ than healthy controls. Recent studies suggest that future patients with other mental illnesses obtain lower pre-morbid IQ. The aims of this study were to compare pre-morbid IQ among five diagnostic categories and normal controls, to examine the distribution of pre-morbid IQ, and to investigate the relationship between pre-morbid IQ and risk of mental illness.

Method. A total of 7486 individuals hospitalized with psychiatric disease and 20 531 controls. IQ was measured at the draft board and hospital diagnoses [schizophrenia (Sz), non-schizophrenic, non-affective psychoses (NSAP), affective (AD), personality (PD) and neurotic/stress disorders (ND)] were followed up to ages 43–54 years. Individuals hospitalized ≤ 1 year after appearing before the draft board were excluded.

Results. All future patients obtained significantly lower pre-morbid IQ than controls (3–7 IQ points), AD had the highest IQ and PD the lowest. In each diagnostic category, decreasing IQ was associated with an increasing risk of becoming a patient [odds ratios (ORs) 0.5–2.5 over the full IQ spectrum]. IQ distributions was nearly normal and uni-modal.

Conclusions. IQ deficits in each diagnostic category may reflect different functional patterns and temporal vicissitudes of the specific pathogenetic processes involved in different mental disorders.

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Introduction

Birth cohort studies, follow-back studies based on school and clinical testing, and high-risk studies all indicate that patients with schizophrenia exhibit impaired performance on tests of intelligence administered before the onset of psychosis (Jones *et al.* 1994; Ott *et al.* 1998; Bilder *et al.* 2006; Woodberry *et al.* 2008), and it is widely believed that schizophrenia is associated with a variety of cognitive deficits (Heinrichs & Zakzanis, 1998; Kremen *et al.* 1998; Mohamed *et al.* 1999). Prospective studies using draft-board tests of intelligence, conducted in Israel (Davidson *et al.* 1999; Rabinowitz *et al.* 2000; Reichenberg *et al.* 2006), Sweden (David *et al.* 1997; Zammit *et al.* 2004) and Finland (Tiihonen *et al.* 2005), show impaired pre-morbid intelligence in patients with schizophrenia.

Several studies show a linear relationship between low IQ and the risk of illness (David *et al.* 1997; Rabinowitz *et al.* 2000; Walker *et al.* 2002; Tiihonen *et al.* 2005; Reichenberg *et al.* 2006).

Comparatively few studies have examined a possible relationship between pre-morbid IQ and mental illnesses other than schizophrenia (Mortensen *et al.* 2005; David *et al.* 2008). However, from focusing exclusively on 'organic factors' in schizophrenia ('dementia in dementia praecox'), interest in cognition has gradually been extended to other psychiatric disorders. Today, an increasing number of studies suggest that pre-morbid intellectual impairment may not be unique to schizophrenia but is also detectable in other psychiatric disorders, although the results for affective illness are less clear (Reichenberg *et al.* 2002; Zammit *et al.* 2004; Mortensen *et al.* 2005; Tiihonen *et al.* 2005; Gale *et al.* 2009). These findings suggest that low pre-morbid IQ may reflect neurodevelopmental disturbances and contribute to the risk of developing mental illnesses in general (David *et al.* 2008; Koenen *et al.* 2009).

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In a previous Danish prospective investigation by Mortensen *et al.* (2005), members of the Copenhagen Perinatal Cohort with draft-board IQ were traced in the Danish Psychiatric Central Research Register. Of 3289 cohort members, 350 had been hospitalized with psychiatric illness: the diagnoses of schizophrenia, other psychosis, personality disorders, and adjustment disorder were associated with low pre-morbid IQ, whereas almost normal IQ was observed for the diagnoses of affective and 'neurotic' disorders. However, the latter patient samples were very small ($n=15$ and 21 respectively) and it is likely that the study did not have sufficient power to detect impairment of IQ scores for these diagnoses. Furthermore, the small patient samples obviously prohibited detailed analyses of the distribution of intelligence within diagnostic categories.

The present study was planned to obtain data on pre-morbid IQ in a very large sample of patients and controls (approximately 10 000 patients and 20 000 controls) with an informative, 25-year follow-up. A case-control design was used for data collection to maximize the number of patients in different diagnostic categories to facilitate generalizations on the relationship between adolescent IQ and later psychiatric diagnosis. Specifically, our aims were (1) to examine and compare pre-morbid intelligence in five diagnostic categories [schizophrenia (Sz), non-schizophrenic and non-affective psychosis (NSAP), affective disorders (AD), personality disorders (PD), neurotic/stress disorders (ND)] and the control group; (2) to examine the relationship between pre-morbid intelligence and risk of each of the five diagnoses; and (3) to examine the distribution of IQ scores within each diagnostic category.

Method

The Danish Psychiatric Central Research Register

Since 1938 this register has systematically collected information on all admissions to psychiatric hospitals and psychiatric wards in Denmark (Munk-Jørgensen & Mortensen, 1997). The register data have been manually accessible for hospital admissions prior to 1969; since then the register has been computerized. It contains the discharge diagnoses of each hospitalization. Since 1970, diagnoses were coded according to the Danish version of the ICD-8 and since 1994, according to the ICD-10. All individuals with a CPR number can be followed through the different admissions across different psychiatric hospitals and psychiatric departments of the country. The CPR number (equivalent to the US social security number) is a unique 10-digit identification number used in all Danish civil registration.

Draft-board intelligence data

With the exception of individuals with disqualifying diseases, all Danish males are required to appear before the draft board when they become liable for conscription at the age of 18 years (5–10% are exempted from appearing because of certified invalidating medical conditions such as epilepsy or diabetes). Appearing before the draft board primarily involves a medical assessment, but also intelligence testing.

Since 1957, intelligence has been assessed with Børge Prién's Prøve (BPP). This test takes 45 min to complete and comprises four subtests: letter matrices (19 items, 15 min), verbal analogies (24 items, 5 min), number series (17 items, 15 min) and geometric figures (18 items, 10 min). For a detailed description of the BPP, see Teasdale & Owen (1989). The number of correct answers is counted for each subtest and summed to a total BPP score with a range of 0–78. This total score has been found to correlate 0.8 with the Wechsler Adult Intelligence Scale (Mortensen *et al.* 1989). The draft archives only register and store the total score of the BPP and not the scores from the subtests. The archives are not computerized and necessitate a manual search.

Study population

The target population of this study comprised all Danish male citizens, born between 1950 and 1961 and drafted between 1968 and 1989, at a mean age of approximately 19.5 years. With follow-up until 2004, 26 941 individuals from this population were identified as psychiatric patients in the Danish Psychiatric Central Research Register (excluding patients with 'organic' diagnoses). The youngest men were aged 43 years and the oldest 54 years at the end of the follow-up period. The age at the first hospitalization was considered as the age of illness onset. Fig. 1 shows the sample attrition.

Collection of IQ data

The Danish draft archives are kept in five geographical locations. Because of financial constraints, our data collection was restricted to draft-board districts 1 and 2, including Copenhagen, the islands of Seeland and of Bornholm and representing more than 2 million inhabitants out of the total Danish population of 5.5 million. Using name and CPR number, draft board BPP scores were located for 8989 hospitalized index cases. By selecting the two cards flanking the index case, usually of the same or a very close birth date, intelligence scores were collected for 20 531 controls, never admitted to psychiatric hospital departments.

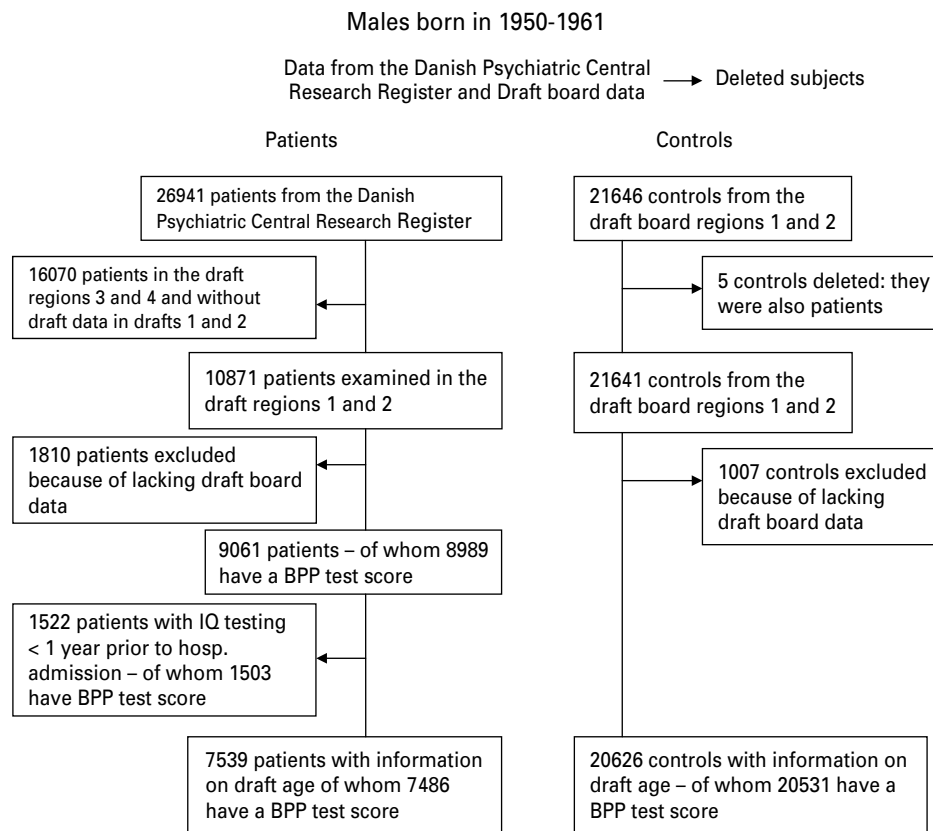


Fig. 1. The sample.

Psychiatric diagnoses

The mean number of hospitalizations per patient was 10.5 (range 1–393, *s.d.* = 20.4). Many patients had several psychiatric diagnoses and to classify such patients we used a hierarchical diagnostic approach following the rules of the ICD-10: schizophrenia (Sz), non-affective non-schizophrenic psychoses (NSAP), affective disorder (AD), neurotic disorders (ND) and personality disorder (PD). Thus, a patient with a last diagnosis of PD, but previously hospitalized for AD, would be classified as AD.

Data analyses

As the purpose of this study was to examine pre-morbid intelligence, we excluded from the analyses 1503 patients who were hospitalized ≤ 1 year following the draft-board examination, a procedure used in two similar studies (Davidson *et al.* 1999; Reichenberg *et al.* 2006). The final sample thus consisted of 7486 patients and 20 531 control subjects with pre-morbid IQ measures. The observed BPP raw score mean of the control sample was 40.11, which is close to the 39.32 mean predicted from a regression formula describing secular trends in BPP scores (Teasdale & Owen, 1989). The standard deviation was 11.24, which is also close

to the 11.38 observed for 6784 Danish conscripts born in 1954–1958 (Teasdale & Owen, 1989). Consequently, to facilitate data interpretation, the BPP intelligence scores were linearly transformed to a scale with a mean of 100 and a standard deviation of 15 in the control group, corresponding to Wechsler's frequently used intelligence scales [this transformation was also used by Mortensen *et al.* (2005)]. For the transformed BPP, age at draft-board examination and age at first admission, the means of the control group were compared with the mean of each diagnostic category and the diagnostic categories were compared two by two. The pooled *t* test or the Cochran *t* test was used depending on whether the variances could be assumed equal or not. The folded form of the *F* statistic was computed to test for equality of the two variances. Finally, for each diagnostic category, logistic regression was used to investigate the association between IQ categorized into seven intervals and the risk of becoming a patient with this particular diagnosis.

Results

The age at first hospitalization was significantly different among all diagnostic categories. The personality disorder patients were the youngest at first

Table 1. Comparison of pre-morbid IQ given by the transformed BBP between the five diagnostic categories and the control group and between each diagnostic category

Outcome variable	IQ	IQ	<i>t</i> test for equal means ^a												
			<i>n</i>	Mean	S.D.	Sz		NSAP		AD		ND		PD	
						<i>t</i> test statistic	<i>p</i> value	<i>t</i> test statistic	<i>p</i> value	<i>t</i> test statistic	<i>p</i> value	<i>t</i> test statistic	<i>p</i> value	<i>t</i> test statistic	<i>p</i> value
Schizophrenia (Sz)	1779	94.38	16.24												
Non-schizophrenic non-affective psychoses (NSAP)	1066	93.38	15.74	1.6	0.11										
Affective disorder (AD)	1845	97.09	15.41	-5.1 ^b	<0.001	-6.2	<0.001								
Neurotic or stress-related syndrome (ND)	625	94.56	15.13	-0.2 ^b	0.81	-1.5	0.13	3.6	<0.001						
Personality disorder (PD)	2171	92.88	16.00	2.9	0.004	0.8	0.40	8.5	<0.001	2.3	0.02				
Control	20 531	100.00	15.00	-14.1 ^b	<0.001	-13.4 ^b	<0.001	-8.0	<0.001	-8.9	<0.001	-19.8 ^b	<0.001		

BBP, Børge Prien's Prøve; S.D., standard deviation; Sz, schizophrenia; NSAP, non-schizophrenic non-affective psychoses; AD, affective disorder; ND, neurotic or stress-related syndrome; PD, personality disorder.

^a When the variances were assumed equal ($p \geq 0.05$), the pooled *t* test was used for the comparisons. When the variances were not assumed equal ($p < 0.05$), the Cochran *t* test was used. When the pooled *t* test was used, degrees of freedom $df = n_1 + n_2 - 2$, where n_1 is the number of subjects in the first category and n_2 is the number of subjects in the second category. When the Cochran *t* test was used, *df* is undefined.

^b The *p* values from the test for equal variances are significant ($p < 0.05$).

hospitalization (mean = 28.9 years, S.D. = 6.9) followed by Sz (mean = 30.4, S.D. = 7.8), NSAP (mean = 32.2, S.D. = 8.2) and ND (mean = 33.4, S.D. = 8.1). As expected, the patients with AD were the oldest at their first hospital contact (mean = 40.0 years, S.D. = 8.2). There were weak but significant correlations between pre-morbid IQ and age at first admission for Sz (0.053, $p = 0.02$), NSAP (0.12, $p < 0.001$), AD and PD (0.14, $p < 0.001$) but not for ND (0.05, $p = 0.16$).

The results for the draft-board BPP are summarized in Table 1. All diagnostic categories of psychiatric patients had significantly lower BPP IQ than the controls. Moreover, there were significant differences among the diagnostic categories: the AD group had a significantly higher IQ than all other diagnostic groups. Patients with Sz, NSAP and ND had a higher IQ than patients with PD, who had the lowest score of all diagnostic categories.

Table 1 also presents standard deviations and shows that all five diagnostic categories had a larger, but not dramatically larger, standard deviation than the control group. The schizophrenia category had the largest standard deviation, significantly larger than that of the control group and the AD and ND categories.

The Kolmogorov–Smirnov test showed that IQ in the control group and the five diagnostic categories was not normally distributed, but calculation of skewness and kurtosis indicated only moderate deviations from normality. In the control group skewness was -0.28 whereas it ranged from -0.01 (PD) to -0.16 (AD) in the diagnostic categories. In the control group kurtosis was -0.28 and ranged among the patients from -0.38 (AD) to -0.50 (Sz). Plots of the IQ distributions also indicated moderate deviation from normality and clearly suggested uni-modal distributions (Fig. 2).

For each of the five diagnostic categories, Table 2 presents odds ratios (ORs) for seven IQ categories. For all diagnostic categories there is increasing risk of becoming hospitalized with a given diagnosis with decreasing intelligence. The confidence intervals (CIs) indicate significantly higher risk for IQ categories <95 compared with the reference group, except for AD and ND in the 85–95 IQ category, and significantly lower risk for IQ categories ≥ 105 , except for Sz and AD disorder in the 115–125 IQ category. Intelligence differences seem to affect the risk of being diagnosed across the full IQ spectrum, but the ORs in Table 2

Table 2. IQ categories and risk of becoming a patient versus a control in each diagnostic category

IQ	Controls <i>n</i>	Schizophrenia			Non-affective non-schizophrenic psychoses			Affective disorder			Neurotic or stress-related syndrome			Personality disorder		
		<i>n</i>	OR	95% CI	<i>n</i>	OR	95% CI	<i>n</i>	OR	95% CI	<i>n</i>	OR	95% CI	<i>n</i>	OR	95% CI
[46;75]	1233	236	2.49	2.09–2.97	139	2.36	1.89–2.94	152	1.38	1.13–1.68	64	1.78	1.31–2.41	314	2.70	2.31–3.16
[75;85]	2122	264	1.62	1.37–1.92	176	1.74	1.42–2.13	261	1.38	1.17–1.62	100	1.61	1.24–2.10	378	1.89	1.63–2.19
[85;95]	3986	399	1.30	1.12–1.51	250	1.31	1.09–1.58	377	<i>1.06</i>	<i>0.92–1.22</i>	147	1.26	<i>1.00–1.60</i>	492	1.31	1.14–1.50
[95;105]	4691	360	1.00	1.00–1.00	224	1.00	1.00–1.00	419	1.00	1.00–1.00	137	1.00	1.00–1.00	442	1.00	1.00–1.00
[105;115]	5329	336	0.82	0.70–0.96	189	0.74	0.61–0.90	411	0.86	0.75–0.99	119	0.76	0.60–0.98	369	0.73	0.64–0.85
[115;125]	2397	153	0.83	<i>0.68–1.01</i>	72	0.63	0.48–0.82	180	<i>0.84</i>	<i>0.70–1.01</i>	49	0.70	0.50–0.97	129	0.57	0.47–0.70
[125;146]	773	31	0.52	0.36–0.76	16	0.43	0.26–0.72	45	0.65	0.47–0.89	9	0.40	0.20–0.79	47	0.65	0.47–0.88

OR, Odds ratio; CI, confidence interval;

Figures in italics are IQ categories not significantly different from the reference category [95;105].

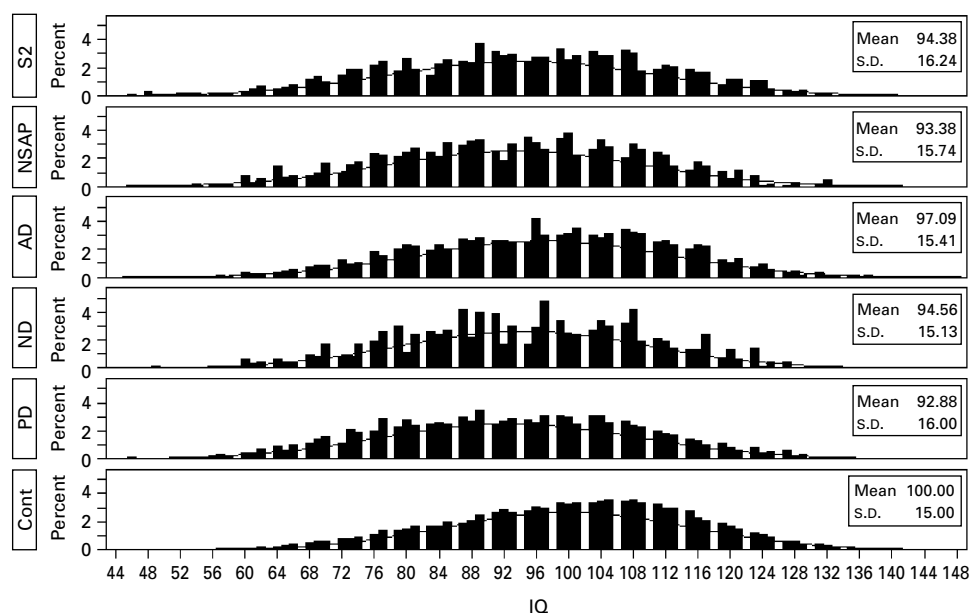


Fig. 2. IQ distribution and normal curves in diagnostic categories. Cont, Control, PD, personality disorder; ND, neurotic or stress-related syndrome; AD, affective disorder; NSAP, non-schizophrenic non-affective psychoses; Sz, schizophrenia.

suggest a non-linear relationship between IQ and risk of mental disorder, in particular for Sz, NSAP and PD. This is illustrated in Fig. 3, which for each diagnostic category shows the results of fitting logistic regression models with the continuous BPP IQ and IQ² scores as linear and quadratic predictors and the risk of being a patient *versus* a control as a binary outcome. The quadratic component was only significant for Sz, but was approaching significance for PD. The linear components were all highly significant, demonstrating a linear trend for all five diagnostic categories.

Discussion

This study focused on describing the observed relationship between draft IQ and risk of mental disorders. The study did not attempt to identify factors that may confound or modify the relationship. The main finding of this large-sample study is that psychiatric in-patients in general, that is irrespective of the diagnosis, have lower pre-morbid IQ than controls from the general population, confirming the previous Danish results by Mortensen *et al.* (2005). The range of

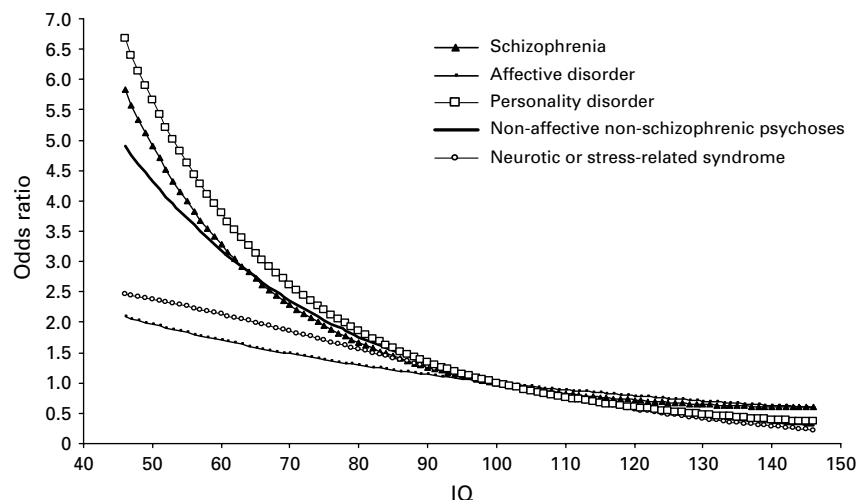


Fig. 3. Theoretical curve for the risk of being a patient *versus* being a control.

observed intelligence deficits was from about 3 to 7 IQ points below the mean of the controls (from about one-fifth to almost half of the standard deviation of 15 in the control group). PDs were associated with particularly low pre-morbid intelligence whereas ADs scored higher on pre-morbid intelligence test than all other diagnostic categories and were closest to the controls. Sz, NSAP and ND performed at similar intermediate levels, numerically closer to the scores of PD than to AD.

All diagnostic categories showed larger IQ variances than the control group, but these differences were modest. The IQ distributions of both the control group and the five diagnostic categories deviated only moderately from the normal distribution. This indicates that, apart from the difference in the mean scores, the distribution of intelligence is similar in healthy controls and in each of the five diagnostic categories. Most ORs in Table 2 are in the range from 0.5 to 2.5, and Fig. 3 shows similar risk patterns for the five diagnostic categories. For schizophrenia, NSAP and PD, $IQ < 75$ may be associated with a particularly high risk, but the general pattern suggests that intelligence differences affect the risk of being diagnosed across the full range of the IQ spectrum for all diagnostic categories. To interpret Fig. 3 it is important to note that the quadratic component was only significant for the schizophrenia category, but also that Table 2 suggests that particular high risk of Sz, NSAP and PD is associated with an $IQ < 75$.

This result is more consistent with a model assuming that the putative risk factors associated with lower intelligence operate in the majority of the patients rather than with a model in which the low mean scores in pre-morbid intelligence within each

diagnostic group stem from a specific subcategory of patients with intelligence deficits. Accordingly, the associations between pre-morbid intelligence and risk of becoming a patient may partly reflect factors common to most categories of mental disease; thus, the hypothesis may be ventured that high intelligence acts as a protective factor (Walker *et al.* 2002; Kremen *et al.* 2007; Koenen *et al.* 2009). For certain categories, such as 'neurotic' and personality disorders, high intelligence and the associated coping abilities may decrease the likelihood of hospital admission (see below), and this may have contributed to the pattern of means observed in Table 1.

Our study is based on hospital diagnoses, and it is possible that IQ is related to the probability of hospitalization given mental illness. To investigate this possibility we calculated the Spearman correlation between the draft IQ and the number of hospitalizations. This correlation was low and slightly negative for the total sample of patients (-0.06) and for all diagnostic categories (ranging from -0.12 for Sz to -0.05 for AD). Because of the large sample size these low correlations were significant, but nevertheless they indicate that the correlation between IQ and number of hospital admissions is too weak to explain the observed associations between IQ and risk of mental disorders. Thus, low IQ is not just a common risk factor for hospitalization but also has an impact on the risk of developing specific mental disorders.

Certain methodological aspects of our study deserve attention. First, it is an inherent weakness of the draft studies that IQ is obtained by late adolescence, which is rather late in the natural course of mental illness (Kessler *et al.* 2005). As described, we minimized this problem by excluding from the analysis

individuals with hospitalization ≤ 1 year after IQ assessment. One limitation of the study is the availability of the total IQ only, but this may not be a serious limitation because aggregate measures of cognitive function tend to predict schizophrenia better than tests of single, very specific cognitive functions (Fearon & Murray, 2002). Second, the study is limited to males only, and the association between intelligence and specific diagnostic categories may be sex dependent. However, in the Israeli draft studies (Reichenberg *et al.* 2005) and in the meta-analysis of Woodberry *et al.* (2008), pre-morbid IQ in schizophrenia did not differ between the sexes. Third, this study only concerns patients identified because of their contact with psychiatric hospitals. In-patients are usually more severely affected than diagnostically similar individuals, who are either untreated or only treated in out-patient clinics (Parnas & Teasdale, 1987). This bias is probably mainly operative for the AD, ND and PD categories (Parnas *et al.* 1995). Fourth, socio-economic status may be related to both intelligence and the risk of mental disorder, and information on socio-economic status was not available for the conscripts. However, Mortensen *et al.* (2005) adjusted BPP scores for parental social status and showed that the low IQ associated with the different categories of mental disorder could not be explained by family socio-economic background. A fifth methodological issue is the diagnostic classification of the patients with multiple hospitalizations and diagnoses. Not all researchers explain how they deal with this problem and those who do, use either the latest diagnosis or a hierarchical approach. We relied on a hierarchical approach following the rules of the ICD-10 as in the Israeli and Swedish studies (Reichenberg *et al.* 2002; David *et al.* 2008). Finally, the diagnoses in this study are clinical and not based on research interviews. This is an intrinsic methodological constraint in epidemiological studies based on hospital diagnoses. However, two Danish studies have demonstrated that the clinical ICD-8 schizophrenia diagnosis, as *de facto* used by the Danish clinicians, was narrow and conservative, compatible with the DSM-III-R operational criteria of schizophrenia (Munk-Jørgensen, 1985; Jørgensen *et al.* 1987). There is also evidence supporting the validity of register-derived diagnoses of affective disorders (Kessing, 1998). It is also possible that changes in diagnostic practice and hospital admission criteria during the follow-up period may have affected the differences in IQ between each diagnostic category and the control group. To investigate this possibility we divided the sample into those born 1950–1955 and those born 1956–1961 and used a 43-year follow-up for both subsamples. There was no significant interaction between the year of birth two-level factor and the

patient *versus* control factor. Thus, the IQ difference between the patients and the controls was similar in two subsamples defined by year of birth for all diagnostic categories, and it seems that that changes in diagnostic practice and hospital admission criteria during the follow-up period have not influenced the differences in IQ between patients and control substantially.

The mean pre-morbid IQ of 94.4 in our schizophrenic subsample was very close to the mean IQ of 94.7 observed in schizophrenic patients in a recent meta-analysis (Woodberry *et al.* 2008). Thus, our results based on the so far largest sample of schizophrenia patients strongly confirm previous findings showing that future schizophrenic patients obtain pre-morbid IQ scores that are about one-half standard deviation lower than that of the controls, but not significantly different from the NSAP or ND patients. Schizophrenia patients performed better than patients with PD and worse than affective patients who showed the best performance of all diagnostic groups, yet nonetheless significantly lower than the performance of the controls (David *et al.* 1997; Davidson *et al.* 1999; Zammit *et al.* 2004; Tiihonen *et al.* 2005; Reichenberg *et al.* 2006).

Both cohort and clinical studies show mixed results with respect to pre-morbid intellectual functioning in affective disorders (van Os *et al.* 1997; Reichenberg *et al.* 2002; Zammit *et al.* 2004; Osuji & Cullum, 2005; Tiihonen *et al.* 2005). It has also been suggested that it is the presence of psychotic symptoms rather than affective disorders *per se* that is related to low pre-morbid intelligence (Reichenberg *et al.* 2002; Hill *et al.* 2004). Depending on the magnitude and the pattern of cognitive differences between schizophrenia and affective illness, authors either opt for a continuity of these two diagnoses or for a nosological separation (Goldberg *et al.* 1993; Reichenberg *et al.* 2002; Hill *et al.* 2004; Zammit *et al.* 2004; McIntosh *et al.* 2005; Tiihonen *et al.* 2005; Green, 2006). The present study shows that future affective disorder patients pre-morbidly obtain higher scores in intelligence tests than all other diagnostic categories, including schizophrenia, but still lower than non-psychiatric controls. However, we do not think that these results justify any radical claim concerning the relationship between schizophrenia and affective illness.

The categories of 'neurotic' (essentially including all non-psychotic Axis I disorders) and personality disorders probably include a fairly heterogeneous spectrum of psychopathology (Coid, 1999). As already noted, these hospitalized patients are very likely to be dissimilar from those untreated or treated in psychotherapeutic out-patient facilities. Our ND and PD samples probably include important proportions of

individuals with significant dysfunction, abuse, maladaptive behaviour and lower IQ. The literature on personality disorder and pre-morbid intelligence or cognition is limited. However, an association with low intelligence is well documented for anti-social personality disorder (Kandel *et al.* 1988). A few clinical studies of borderline, obsessive-compulsive, histrionic and narcissistic PDs identified deficits in several neurocognitive domains (Burgess, 1992; Coid, 1999; Ruocco, 2005).

The results concerning schizophrenia and non-affective psychosis are compatible with the hypothesis of disturbed neurodevelopmental trajectories in the genesis of schizophrenia (Pantelis *et al.* 2005; Rappaport & Addington, 2005), but do not provide information on the possible neurodegenerative decline in the course of the disease (Seidman *et al.* 2006). With the exception of ND, there were significant, albeit weak, correlations between low IQ and young age at the illness onset, not inconsistent with a neurodevelopmental genesis of mental disease.

Conclusion

This large sample study demonstrates that hospitalized patients of all major diagnostic categories obtain significantly lower pre-morbid intelligence scores than healthy controls. Patients hospitalized within 1 year of intelligence testing were excluded and the shortest mean time span that elapsed from the time of testing to the first hospitalization was 10 years. Therefore, the observed IQ scores are unlikely to be just a consequence of a fully developed (but undetected) disorder or of the so-called 'prodromal' symptoms.

The IQ is an aggregate of several measures of cognitive capacities. It may be considered as an index of the history of ontogenetic interactions between genetic, environmental and complex epigenetic processes, thus lending space to an interaction between several more specific, yet still extremely complex, factors such as genetic vulnerability, perinatal circumstances, socio-economic factors, and upbringing. It is likely that similar IQ deficits may reflect differential functional patterns and temporal vicissitudes of the processes that are operative in different, specific mental disorders. We believe that the demonstration of the IQ deficits in a range of psychiatric disorders should stimulate a multiplicity of research perspectives on the pathogenetic processes involved.

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Declaration of Interest

None

References

- Bilder RM, Reiter G, Bates J, Lencz T, Szeszko 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.
- Burgess JW** (1992). Neurocognitive impairment in dramatic personalities: histrionic, narcissistic, borderline, and antisocial disorders. *Psychiatry Research* **42**, 283–290.
- Coid JW** (1999). Aetiological risk factors for personality disorders. *British Journal of Psychiatry* **174**, 530–538.
- David AS, Malmberg A, Brandt L, Allebeck P, Lewis G** (1997). IQ and risk for schizophrenia: a population-based cohort study. *Psychological Medicine* **27**, 1311–1323.
- David AS, Zammit S, Lewis G, Dalman C, Allebeck P** (2008). Impairments in cognition across the spectrum of psychiatric disorders: evidence from a Swedish conscript cohort. *Schizophrenia Bulletin* **34**, 1035–1041.
- Davidson M, Reichenberg A, Rabinowitz J, Weiser M, Kaplan Z, Mark M** (1999). Behavioral and intellectual markers for schizophrenia in apparently healthy male adolescents. *American Journal of Psychiatry* **156**, 1328–1335.
- Fearon P, Murray RM** (2002). Intellectual function and schizophrenia. *British Journal of Psychiatry* **181**, 276–277.
- Gale RG, Deary IJ, Boyle SH, Barefoot J, Mortensen LH, Batty D** (2009). Cognitive ability in early adulthood and risk of 5 specific psychiatric disorders in middle age. The Vietnam experience study. *Archives General of Psychiatry* **65**, 1410–1418.
- Goldberg TE, Gold JM, Greenberg R, Griffin S, Schulz SC, Pickar D, Kleinman JE, Weinberger DR** (1993). Contrasts between patients with affective disorders and patients with schizophrenia on a neuropsychological test battery. *American Journal of Psychiatry* **150**, 1355–1362.
- Green MF** (2006). Cognitive impairment and functional outcome in schizophrenia and bipolar disorder. *Journal of Clinical Psychiatry* **67**, 36–42.
- Heinrichs RW, Zakzanis KK** (1998). Neurocognitive deficit in schizophrenia: a quantitative review of the evidence. *Neuropsychology* **12**, 426–445.
- Hill SK, Keshavan MS, Thase ME, Sweeney JA** (2004). Neuropsychological dysfunction in antipsychotic-naïve first-episode unipolar psychotic depression. *American Journal of Psychiatry* **161**, 996–1003.
- Jørgensen A, Teasdale TW, Parnas J, Schulsinger F, Schulsinger H, Mednick SA** (1987). The Copenhagen high-risk project: the diagnosis of maternal schizophrenia

- and its relation to offspring diagnosis. *British Journal of Psychiatry* **151**, 753–757.
- Jones P, Rodgers B, Murray RM, Marmot M** (1994). Child developmental risk factors for adult schizophrenia in the British 1946 birth cohort. *Lancet* **344**, 1398–1402.
- Kandel E, Mednick SA, Kirkegaard-Sorensen L, Hutching B, Knop J, Rosenberg R, Schulsinger F** (1988). IQ as a protective factor for subjects at high risk for antisocial behavior. *Journal of Consulting and Clinical Psychology* **56**, 224–226.
- Kessing LV** (1998). Validity of diagnoses and other register data in patients with affective disorder. *European Psychiatry* **13**, 342–345.
- Kessler RC, Berglund P, Demier O, Jin R, Merikangas KR, Walters EE** (2005). Lifetime prevalence and age-of-onset distributions of DSM-IV disorders in the National Comorbidity Survey Replication. *Archives of General Psychiatry* **62**, 593–602.
- Koenen KC, Moffitt TE, Roberts AL, Martin LT, Kubzansky L, Harrington HL, Poulton R, Caspi A** (2009). Childhood IQ and adult mental disorders: a test of the cognitive reserve hypothesis. *American Journal of Psychiatry* **166**, 50–57.
- Kremen WS, Buka SL, Seidman LJ, Goldstein JM, Koren D, Tsuang MT** (1998). IQ decline during childhood and adult psychotic symptoms in a community sample: a 19-year longitudinal study. *American Journal of Psychiatry* **155**, 672–677.
- Kremen WS, Koenen KC, Boake C, Purcell S, Eisen SA, Franz CE, Tsuang MT, Lyons MJ** (2007). Pretrauma cognitive ability and risk for posttraumatic stress disorder. *Archives of General Psychiatry* **64**, 361–368.
- McIntosh AM, Harrison LK, Forrester K, Lawrie SM, Johnstone EC** (2005). Neuropsychological impairments in people with schizophrenia or bipolar disorder and their unaffected relatives. *British Journal of Psychiatry* **186**, 378–385.
- Mohamed S, Paulsen JS, O'Leary D, Arndt S, Andreasen N** (1999). Generalized cognitive deficits in schizophrenia: a study of first-episode patients. *Archives of General Psychiatry* **56**, 749–754.
- Mortensen EL, Reinisch JM, Teasdale TW** (1989). Intelligence as measured by WAIS and a military draft board group test. *Scandinavian Journal of Psychology* **30**, 315–318.
- Mortensen EL, Sorensen HJ, Jensen HE, Reinisch JM, Mednick SA** (2005). IQ and mental disorder in young men. *British Journal of Psychiatry* **187**, 407–415.
- Munk-Jørgensen P** (1985). The schizophrenia diagnosis in Denmark. A register-based investigation. *Acta Psychiatrica Scandinavica* **72**, 266–273.
- Munk-Jørgensen P, Mortensen PB** (1997). The Danish Psychiatric Central Register. *Danish Medical Bulletin* **44**, 82–84.
- Osuji IJ, Cullum CM** (2005). Cognition in bipolar disorder. *Psychiatric Clinics of North America* **28**, 427–441.
- Ott SL, Spinelli S, Rock D, Roberts S, Amminger GP, Erlenmeyer-Kimling L** (1998). The New York High Risk Project: social and general intelligence in children at risk for schizophrenia. *Schizophrenia Research* **31**, 1–11.
- Pantelis C, Yücel M, Wood SJ, Velakoulis D, Sun D, Berger G, Stuart GW, Yung A, Phillips L, McGorry PD** (2005). Structural brain imaging for multiple pathological processes at different stages of brain development in schizophrenia. *Schizophrenia Bulletin* **31**, 672–696.
- Parnas J, Cannon M, Schulsinger F, Mednick SA** (1995). Early predictors of onset and course of schizophrenia: results from the Copenhagen High-Risk Study. In *Search for the Causes of Schizophrenia*, vol. III (ed. H. Häfner and W. F. Gattaz), pp. 67–86. Springer Verlag: Berlin.
- Parnas J, Teasdale TW** (1987). A matched-paired comparison of treated versus untreated schizophrenia spectrum cases: a high-risk population study. *Acta Psychiatrica Scandinavica* **75**, 44–50.
- Rabinowitz J, Reichenberg A, Weiser M, Mark M, Kaplan Z, Davidson M** (2000). Cognitive and behavioral functioning in men with schizophrenia both before and shortly after first admission to hospital. *British Journal of Psychiatry* **177**, 26–32.
- Rappaport JL, Addington AM** (2005). The neurodevelopmental model of schizophrenia: update 2005. *Molecular Psychiatry* **10**, 434–449.
- Reichenberg A, Weiser M, Caspi A, Knobler HY, Lubin G, Harvey P, Rabinowitz J, Davidson M** (2006). Premorbid intellectual functioning and risk of schizophrenia and spectrum disorders. *Journal of Clinical and Experimental Neuropsychology* **28**, 193–207.
- Reichenberg A, Weiser M, Rabinowitz J, Caspi A, Schmeidler J, Mordechal M, Zeev K, Davidson M** (2002). A population-based cohort study of premorbid intellectual, language, and behavioral functioning in patients with schizophrenia, schizoaffective disorder, and nonpsychotic bipolar disorder. *American Journal of Psychiatry* **159**, 2027–2035.
- Reichenberg A, Weiser M, Rapp MA, Rabinowitz J, Caspi A, Schmeidler J, Knobler HY, Lubin G, Nahon D, Harvey PD, Davidson M** (2005). Elaboration on premorbid intellectual performance in schizophrenia: premorbid intellectual decline and risk for schizophrenia. *Archives of General Psychiatry* **62**, 1297–1304.
- Ruocco AC** (2005). The neuropsychology of the borderline personality disorder: a meta-analysis and review. *Psychiatry Research* **137**, 191–202.
- Seidman LJ, Buka SL, Goldstein JM, Tsuang MT** (2006). Intellectual decline in schizophrenia: evidence from a prospective birth cohort 28 year follow-up study. *Journal of Clinical and Experimental Neuropsychology* **28**, 225–242.
- Teasdale TW, Owen DR** (1989). Continuing secular increase in intelligence and a stable prevalence of high intelligence level. *Intelligence* **13**, 255–262.
- Tiihonen J, Haukka J, Henriksson M, Cannon M, Kieseppä T, Ilmo L, Sinivuo J, Lönnqvist J** (2005). Premorbid intellectual functioning in bipolar disorder and schizophrenia: results from a cohort study of male conscripts. *American Journal of Psychiatry* **162**, 1904–1910.
- van Os J, Jones P, Lewis G, Wadsworth M, Murray R** (1997). Developmental precursors of affective illness in a general population birth cohort. *Archives of General Psychiatry* **54**, 625–631.
- Walker NP, McConville PM, Hunter D, Deary IJ, Whalley LJ** (2002). Childhood mental ability and lifetime

- psychiatric contact: a 66-year follow-up study of the 1932 Scottish Mental Ability Survey. *Intelligence* **30**, 233–245.
- Woodberry KA, Anthony GJ, Seidman LJ** (2008). Premorbid IQ in schizophrenia: a meta-analytic review. *American Journal of Psychiatry* **165**, 579–587.
- Zammit S, Allebeck P, David AS, Dalman C, Hemmingsson T, Lundberg I, Lewis G** (2004). A longitudinal study of premorbid IQ score and risk for developing schizophrenia, bipolar disorder, severe depression, and other nonaffective psychoses. *Archives of General Psychiatry* **61**, 354–360.