Brief article

Title: Neurocognitive insight and executive functioning in schizophrenia

Running head: Neurocognitive insight in schizophrenia

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Abstract

Introduction: This study was aimed at exploring whether integrity of executive functioning is required to have a good neurocognitive insight in persons with schizophrenia.

Method: Neurocognitive insight was measured by subtracting executive difficulties (errors in the Modified Card Sorting Task) to executive cognitive complaints (Subjective Scale To Investigate Cognition in Schizophrenia) in forty persons with schizophrenia and forty-two normal controls. The schizophrenia sample was a priori divided into two subgroups on the basis of executive level. Multivariate analyses were conducted to compare groups and to control for potential confounding factors.

Results: Only the schizophrenia dysexecutive subgroup had a poorer neurocognitive insight compared to normal controls. Group differences remained significant after adjustment for potential confounding factors (education, depression, anxiety, self-esteem).

Conclusion: These results provide support for the hypothesis that executive dysfunctioning is a limiting factor for neurocognitive insight, independently from depressive and anxiety symptoms.

Key words: cognition, cognitive complaint, self-awareness, mental flexibility

Introduction

Neurocognitive deficits in schizophrenia, occurring in several domains of neurocognition (Fioravanti, Carlone, Vitale, Cinti, & Clare, 2005), have been consistently linked to functional outcomes (Prouteau et al., 2005). In this perspective, awareness of neurocognitive difficulties, i.e. neurocognitive insight (NI), is of particular importance for optimizing efficacy of cognitive remediation aimed at enhancing neurocognitive functioning. For instance, self-perception of cognitive dysfunction, has been reported to be a strong predictor of long-term occupational outcome (Verdoux, Monello, Goumilloux, Cougnard, & Prouteau, 2010).

In the first study dealing explicitly with NI in schizophrenia, Medalia and Lim (2004) reported that outpatients are poor at accurately classify their cognitive status (impaired or not) in memory and attention. In subsequent studies, 28% to 52 % of cognitively impaired participants are considered to have no NI, depending on the used tools (self-reports versus clinician ratings) (Medalia & Thysen, 2008; Medalia, Thysen, & Freilich, 2008). Additionally, even if a majority of people with schizophrenia underestimates their impairment, some overestimate it (Gilleen, Greenwood, & David, 2011; González-Suárez et al., 2011). Hence, several subprofiles of NI could co-exist within included samples.

In the domain of NI, two theoretical levels of analyses and interpretation have to be differentiated. Indeed, cognitive complaints reflect difficulties relevant from the subject's viewpoint, but also reflect the efficiency of underlying mechanisms allowing the subjects to become aware of their difficulties. Brekke et al (2001) suggested that a lower executive functioning level would prevent persons with schizophrenia from making sophisticated judgments about themselves, taking into account their past history and the others. Executive functions such as mental flexibility are critical for

having access to mnesic representations, which constitute the pool of information on which judgments about one's own cognitive functioning are based. To date, the few studies dealing with this issue failed to find any association between NI and executive functioning (Donohoe et al., 2009; Gilleen, et al., 2011; Medalia & Thysen, 2008). However, associations were explored at the whole group level, potentially masking associations restricted to specific subprofiles.

This study was aimed at exploring in persons with schizophrenia whether integrity of executive functioning is required to have a good NI.

Method

Participants

Forty participants with schizophrenia (clinical group) were consecutively recruited during ambulatory treatment in the rehabilitation unit of a public psychiatric hospital. Included participants fulfilled the following criteria: 1) diagnosis of schizophrenia or schizoaffective disorder according to the DSM-IV criteria (APA, 2000); 2) age range between 18 and 65 years; 4) fluency in French; 5) no substance or alcohol dependence according to the DSM-IV criteria; 6) no history of neurological disease or brain injury; 7) medication and clinical status stable for at least one month. All the participants were receiving antipsychotic medication at the time of assessment.

To explore whether NI depends on executive functioning level, we split the schizophrenia group into two subgroups on the basis of executive performance. As we used the MCST score for measuring NI, we used another executive test, i.e. the TMT B-A score, as an independent indicator of executive functioning. The TMT B-A 5th percentile (Godefroy & GREFEX, 2008) was a priori used as a cut-off score to

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split the schizophrenia group into two subgroups. The first one was considered as "schizophrenia executively normal", containing participants with TMT B-A scores strictly inferior to 96 seconds. The second one was the "schizophrenia dysexecutive group", with scores ranging from 96 seconds to highest. The 5th percentile was chosen in accordance with consideration about pathological threshold in neuropsychology, and was selected according to schizophrenia group's mean level of education.

Forty two control participants (control group) were recruited through announcement in medical centers. Their psychiatric status was assessed using the Mini International Neuropsychiatric Interview (MINI), to exclude subjects with DSM-IV diagnoses of mental disorders (Lecrubier et al., 1997). The other inclusion criteria were the same as for patients.

The investigation was conformed to the French bioethics and clinical research legislation and all participants provided written informed consent to participate. Local ethical committee approved the procedures.

Assessments

A trained psychologist conducted all assessments. Executive functioning was assessed with the French validated versions (Godefroy & GREFEX, 2008) of the Modified Card Sorting Test (MCST) (Nelson, 1976) and the Trail Making Test (TMT) (Reitan, 1958). Cognitive complaints were measured using the French validated version of the Subjective Scale To Investigate Cognition in Schizophrenia (SSTICS, Stip, Caron, Renaud, Pampoulova, & Lecomte, 2003). The SSTICS is 21-items self-report questionnaire providing cognitive complaints scores in several domains: memory, attention, executive function, praxia and language. Subjects have to rate

each item on a 5-point Likert-type scale (from 0: « never » to 4: « very often »). The SSTICS sustained executive function score (max = 16) was considered in the present study.

The participants completed French validated versions of self-administered questionnaires to assess depressive symptoms with the Beck Depression Inventory-II (Beck, Steer, & Brown, 1998), anxiety symptoms with the State-Trait Anxiety Inventory (Spielberger, 1983), and self-esteem with the Rosenberg Self-Esteem Inventory (Rosenberg, 1965).

Neurocognitive insight calculation

A NI variable was created for the present study to reflect adequacy between selfperceived executive functioning and real executive performance. A discrepancy score was thus calculated by subtracting executive performance (error percentage in MCST) to executive complaint (complaint percentage in the SSTICS 'sustained executive functioning' factor). A value close to zero reflects a good NI, whereas values far from zero indicate either lack of awareness of difficulties (negative values), or overestimation of difficulties (positive values).

Statistical analyses

Analyses were carried out with the Statistical Package for Social Science[®]. In order to explore differences between groups, the Chi-square test was used to compare categorical qualitative data, whereas *t*-test for independent samples and one-way analysis of variance (ANOVA) were used to compare quantitative continuous variables. Degrees of freedom for the independent samples *t*-tests were corrected for unequal variances based off of a Levene's test for equality of variances. Post-hoc

analyses were performed using Tukey HSD (Honestly Significant Difference) test for pairwise comparisons.

Comparisons between the three groups (control, schizophrenia executively normal and schizophrenia dysexecutive groups) were made using one-way ANOVA on the NI score. Finally, multiple regression analyses were performed in order to control for sociodemographic characteristics, depression, anxiety and self-esteem. For all tests, significance was set at p<.05, two-tailed.

Results

Participants' characteristics

The "schizophrenia executively normal" group contained N=29 (72.5%) participants with TMT B-A scores strictly inferior to 96 seconds. There was N=11 (27.5%) participants the "schizophrenia dysexecutive" group, with scores ranging from 96 seconds to highest.

Participants' characteristics are described in Table 1. Group comparisons revealed that groups were not different on gender (χ^2 =3.53, df= 2, *P*=0.17) or age (F_(2, 79)=1.54, *P*=0.22), but were different in education level (χ^2 =41.76, df=2, *P*<0.001), SSTICS' executive complaint score (F_(2, 79)=16.50, *P*<0.001), executive performance (MCST : F_(2, 79)=19.80, *P*<0.001; TMT B-A: F_(2, 78)=82.47, *P*<0.001), depression score (F_(2, 77)=18.57, *P*<0.001), state-anxiety score (F_(2, 78)=4.71, *P*<0.05) and self-esteem score (F_(2, 64)=5.12, *P*<0.01). The schizophrenia groups were not different in terms of duration of illness (F_(1, 35)=1.15, *P*=0.29).

(Insert Table 1 here)

Neurocognitive Insight differences

The Table 1 presents scores in the three groups (control, schizophrenia executively normal, schizophrenia dysexecutive). In the schizophrenia dysexecutive group, there was a majority of high positive NI scores (63% had scores superior to 6.25), contrasting with the two other groups (24% in the schizophrenia executively normal group, 9.5% in the control group). One way ANOVA on the NI score showed a significant effect of group ($F_{(2, 79)}$ =5.45; *P*<0.01, η^2 =.12). Post hoc analyses revealed that only the schizophrenia dysexecutive group significantly differed from the control group (*P*<0.05, *d*=1.31). There was no difference between the two schizophrenia executively normal groups (*P*=0.064, *d*=.63), nor between the control group and the schizophrenia executively normal group (*P*=0.42, *d*=.33). Finally, multiple regression analysis showed that the effect of group remained significant when controlled for education, anxiety, depression and self-esteem variables.

Discussion

Our results show that only the schizophrenia dysexecutive group has a poorer NI compared to normal controls, independently from education, anxiety, depression and self-esteem. This result provides support for the hypothesis of Brekke et al (2001), as executively impaired participants were less able than controls to evaluate accurately their cognitive abilities. Our results apparently contrast with those of previous studies that failed to find any correlation between awareness of cognitive impairment and executive functioning (Donohoe, et al., 2009; Gilleen, et al., 2011; Medalia & Thysen,

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2008). This inconsistency might be related to discrepancies in the used methods of analyses. Indeed, in previous studies, correlations were explored at the whole group level, thus potentially masking different profiles among participants. Our results suggest that exploring awareness of cognitive impairment among groups of cognitive functioning (for instance severe executive impairment versus no or mild impairment) could be a more sensitive method. Moreover, the link between executive functioning and judgments about one's neurocognitive abilities has also been underscored in the domain of metamemory and aging. In elderly participants, metamemory control may be largely the result of executive limitations associated with aging (Souchay & Isingrini, 2004). Altogether, these results suggest that the integrity of executive functioning has an impact on metacognition (i.e. cognition about one's cognition), whether during one particular cognitive task (i.e. metacognitive judgment and control), or more generally on what a subject knows about her/his everyday functioning (metacognitive knowledge or neurocognitive insight). Further research are necessary to understand these relationships in normal controls, as well as their specificities in schizophrenia, in which executive dysfunctions are known to be frequent (Fioravanti, et al., 2005).

Our results also show that problems for evaluating neurocognitive difficulties do not always appear in the sense of a 'lack of awareness'. Indeed, 63% of the dysexecutive participants tend to overestimate their difficulties. This result is consistent with previous studies, showing that subjects with schizophrenia could either underestimate or overestimate their difficulties (Donohoe, et al., 2009; González-Suárez, et al., 2011; Medalia & Thysen, 2008). González-Suárez et al (2011) argued that this phenomenon could be a manifestation of low self-esteem or depression. In our study however, it seems unlikely that self-esteem, depression and anxiety could have played a major role, as these factors do not explain differences in neurocognitive insight between groups. The great amount of participants presenting with an overestimation of difficulties is at first sight surprising, given the classical consideration on unawareness of difficulties (lack of insight) in schizophrenia. However, it has already been shown that in schizophrenia, several components of self-awareness should be differentiated, such as clinical insight, cognitive insight and cognitive complaints (Tastet, Verdoux, Bergua, Destaillats, & Prouteau, 2012). Furthermore, this finding also suggests that, in some dysexecutive patients, neurocognitive deficits can be clearly perceivable. In practice, overestimation of difficulties constitutes a privileged way for including these patients in remediation programs. Considering cognitive complaints may contribute to enhance intrinsic motivation. Indeed, self-reported cognitive difficulties are perceivable in daily life and could reflect some relevant limitations for an individual project (i.e. education, work, social relationships).

This study has several methodological limitations. First, our sample was composed of persons with schizophrenia treated in a rehabilitation unit; hence results may not be generalizable to the entire population of persons suffering from schizophrenia. The limited size of our sample prevented us from studying NI in some moderate or mild executive impairment groups. We thus cannot exclude that subtle NI impairments could exist in such groups. Another potential limitation is that our cut-off score for isolating a subgroup of dysexecutive subjects is severe (percentile 5). Executive difficulties, though milder, may exist for other subjects of the entire sample. Finally, the present study design was limited to mental flexibility. Further studies are needed to know whether the same phenomenon can be described in other executive dimensions.

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In conclusion, our findings suggest that executive dysfunctioning is a limiting factor for NI in schizophrenia. Interventions aimed at enhancing NI, and more globally selfawareness, should first consider the availability of cognitive processes within each subject, particularly in the domain of executive functioning.

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Tables

Table 1: score differences between groups

	Schizophrenia dysexecutive ¹ (N=11)	Schizophrenia executively normal ² (N=29)	Control (N=42) Mean SD N (%)
	Mean SD N(%)	Mean SD N(%)	
• Age	40.00 (10.96)	36.93 (9.07)	33.86 (12.56)
• Gender	7 (63.6%)	15 (51.7%)	15 (35.7%)
 Education level < 6 years 	2 (18.2%)	1 (3.4%)	0 (0%)
• 6 to 12 years	9 (81.8%)	23 (79.3%)	9 (21.4%)
• > 12 years	0 (0%)	5 (17.2%)	33 (78.6%)
• Duration of illness ³	16.81 (13.95)	12.88 (8.20)	-
• MCST ⁴ : errors	11.45 (7.15)	6.79 (6.29)	2.02 (2.33)
SSTICS ⁵ : Sustained executive complaints score	6.27 (2.69)	6.07 (2.83)	2.98 (2.16)
• TMT B-A ⁶	161.60 (63.76)	42.11 (25.39)	34.26 (16.94)
Neurocognitive insight ⁷ score	8.10 (12.57)	1.13 (10.52)	-1.49 (5,34)
• BDI-II ⁸ : total score	15.27 (12.30)	15.03 (8.52)	4.65 (5.03)

• STAI ⁹ : state score	31.55 (12.48)	38.79 (13.93)	30.21 (9.58)
• RSE ¹⁰ : total score	6.86 (2.54)	6.71 (2.34)	8.40 (1.84)

*: p<.01; **: p<.001

¹Subjects with a Trail Making B-A score $\leq 5^{th}$ percentile

² Subjects with a Trail Making B-A score >5th percentile

³ defined as the number of years since the first hospitalization

⁴ MCST: Modified Card Sorting Task

⁵ SSTICS: Subjective Scale To Investigate Cognition in Schizophrenia

⁶ TMT B-A: Trail Making Test : time B-time A (s)

⁷ defined as 'sustained executive complaint percentage – executive errors percentage'

⁸ BDI-II: Beck Depression Inventory-II

⁹ STAI: State-Trait Anxiety Inventory

¹⁰ RSE: Rosenberg Self-Esteem inventory