Sensorimotor induction of auditory misattribution in early psychosis

Running title: Sensorimotor basis of source monitoring errors

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Abstract

Dysfunction of sensorimotor predictive processing is thought to underlie abnormalities in self-monitoring producing passivity symptoms in psychosis. Experimentally induced sensorimotor conflict can produce a failure in bodily self-monitoring (Presence hallucination, PH), yet it is unclear how this is related to auditory self–monitoring and psychosis symptoms. Here we show that the induction of sensorimotor conflict in early psychosis patients induces presence hallucinations and impacts auditory-verbal self-monitoring. Participants manipulated a haptic robotic system inducing a bodily sensorimotor conflict. In exp.1 the PH was measured. In exp.2 an auditory-verbal self-monitoring task was performed during the conflict. 51 participants (31 early psychosis patients, 20 matched controls) participated in the experiments. The PH was present in all participants. Psychosis patients with passivity experiences (PE+) had reduced accuracy in auditory-verbal self-other discrimination during sensorimotor stimulation, but only when sensorimotor stimulation involved a spatio-temporal conflict (F(2, 44)=6.68, p=.002). These results show a strong link between robotically-controlled alterations in sensorimotor processing and auditory misattribution in psychosis and provide evidence for the role of sensorimotor processes in altered self-monitoring in psychosis.

Keywords: Early Psychosis, Source monitoring, Sensorimotor conflict, Predictive Processing, Sense of Agency
**Introduction**

Psychosis is characterized by abnormal mental states including hallucinations and delusions and a deterioration of cognitive and emotion processing\(^1\). Moreover, people with psychosis often report strange and compelling feelings that other people or external forces are influencing them\(^2\). They may experience external forces as having a direct and uncontrollable influence on their inner experience, leading to intriguing symptoms such as auditory-verbal hallucinations (AVH) in form of voices commenting the person’s thoughts, delusional ideas that thoughts are inserted or stolen from their minds, or sensations of external control over their bodies and actions. These symptoms, termed “Passivity Experiences (PE)” and formerly known as schneiderian first rank symptoms, reflect a diminished demarcation of self-other boundaries and misattributions of self-generated thoughts and actions to external sources\(^3-5\). It has been suggested that these deficiencies are due to abnormal sensorimotor prediction mechanisms, causing erroneous self-monitoring\(^6-9\). The putative mechanism for self-monitoring is based upon the comparison of predictions regarding sensory outcomes of self-generated actions (i.e. efference copy) with afferent sensory signals\(^10,11\). When predictions and afferent sensory signals match, actions are self-attributed and the sensory signal is attenuated\(^12-15\). If a discrepancy is found, sensory signals are less attenuated, indicating a probable external origin\(^5,16\). Several studies have shown aberrant sensorimotor prediction and reduction of normal sensory attenuation for self-generated actions in schizophrenia\(^17-24\), providing evidence linking deficient sensorimotor prediction with self-other discrimination symptoms of psychosis\(^6,7,25\). While the neurobiological basis for AVH is still debated\(^25-28\) it has been suggested that AVH may arise from misattribution of inner speech to an external source due to defective efference copy mechanisms\(^28-31\). Studies have found correlational evidence for deficits in self-monitoring of
auditory-verbal content in schizophrenia\textsuperscript{29,32,33} as well as decreased sensorimotor prediction in patients with AVH\textsuperscript{17,34,35}.

However, this previous evidence is correlational, showing that deficient sensorimotor predictions in schizophrenia often correlate with positive symptom severity. Thus, more causal evidence employing an experimentally generated sensorimotor conflict to induce a psychosis-like mental state able to impair auditory-verbal processing in an online fashion is still lacking. Here, we used a custom robotic device\textsuperscript{36} to investigate this issue. By directly manipulating sensorimotor contingencies, the device induces a mental state characterized by an illusory feeling that a person is standing behind them (i.e. Presence Hallucination, PH)\textsuperscript{37}. Here, standing and blindfolded participants control the “lead” robot\textsuperscript{38} by moving their arm. Their movements are sent to a ”follow” robot, which applies tactile feedback in real time to the participants’ backs. In order to manipulate sensorimotor contingency, in the asynchronous condition, a 500 ms delay is introduced between the participants’ movements and the haptic feedback. We previously showed that, in healthy individuals, this asynchronous condition induces the feeling that a person is standing behind them, i.e. PH\textsuperscript{37}. Here we investigated whether the PH effects could be reproduced in psychosis patients with (PE+) and without (PE-) passivity symptoms, and critically, whether the same manipulation would also induce deficits in self-monitoring of auditory-verbal stimuli. To this aim, healthy participants and psychosis patients were presented during the sensorimotor stimulation with pre-recorded words, spoken either by themselves or by another person, and were asked to discriminate between the two\textsuperscript{30}. Thus, in the present study, we directly tested (1) whether robotic induction of a sensorimotor conflict, able to induce a mental state characterized by PH in healthy participants, induces the same effect in psychosis patients (PH Experiment), and (2) whether it impacts self-monitoring for auditory-verbal information, as is thought to occur for AVH in psychosis (Self-
Monitoring experiment). Thus, contrary to previous investigations, our design allows for online and active induction of self-monitoring errors through task irrelevant sensorimotor conflicts.

**Methods**

**Participants**

Thirty-one individuals in the early phase of psychosis having met criteria for a psychotic episode according to the Comprehensive Assessment of At-Risk Mental States (CAARMS) criteria (mean age= 26.4 SD= 4.6 years; 9 women) were recruited among the Treatment and early Intervention in Psychosis Program (TIPP), a 3-year program launched in 2004 in Lausanne (CHUV, Switzerland). Twenty age-matched control subjects (mean age= 25.95, SD=4.7 years; 5 women) from the control cohort of the TIPP program were also recruited. Exclusion criteria for the study were history of neurological illness or trauma, non-psychiatric visual or auditory disorders, diagnosis of psychosis related to intoxication or organic brain disease, intelligence quotient <70, age below 18 and above 35 at the TIPP inclusion, or antipsychotic medication more than 6 months before the inclusion. 90.3% of the subjects were right-handed. The participant group included all participants who were available and willing to participate in this study and met the inclusion criteria from the TIPP cohort at the time of the study. Controls were assessed and selected with the Diagnostic Interview for Genetic Studies. Major mood, psychotic, or substance-use disorder and having a first-degree relative with a psychotic disorder were exclusion criteria for controls. All were clinically stable during testing and gave an informed written consent. The study was approved by the Ethical Committee of Clinical Research of the University of Lausanne, Switzerland, and was run in accordance with the ethical guidelines of the ethical committee and the declaration of Helsinki.

5
The patients underwent an in depth clinical assessment by a trained psychiatrist the same day as the behavioural experiment. Symptom severity and classification were assessed in the patient group using the Positive and Negative Syndrome Scale (PANSS) for schizophrenia with a mean of the total score of 52.52 (SD=13.67) (13.90 (SD=4.96) on the positive subscale, 13.72 (SD=4.71) on the negative subscale and 24.62 (SD=7.2) on the general psychopathology subscale). Passivity symptoms were assessed using specific items of the Scale for the Assessment of Positive Symptoms (SAPS)\textsuperscript{39} (item 2: voices commenting; item 3: voices conversing; item 15: delusions of being controlled; item 18: though insertion; item 19: thought withdrawal). Patients were considered PE+ (N=19) if they had presented at least one of these five PE during the psychotic episodes\textsuperscript{35,40}. 12 patients never presented passivity symptoms and were thus included in the PE- group. The two groups were demographically similar in terms of age (p =.94) and level of education (p =.84). However, although there were no differences between the two groups in the global clinical features (total PANSS scores) (p =.41) and treatment duration (p =.24), there was an expected significant difference in the positive PANSS subscore (related to PE) (p =.02) and the chlorpromazine equivalent medication (used to treat passivity symptoms) (p =.01) (see Table 1 & supplementary material for full clinical details).

Initial diagnosis of the patient based on their first episode symptoms revealed that at the time of the experiment, 19 patients satisfied the ICD-10 criteria for schizophrenia, 2 for schizoaffective disorders, 6 for acute and transient psychotic disorders, 3 for severe depressive episode with psychotic symptoms and 1 for bipolar affective disorder.
Procedure

Presence Hallucination experiment

The procedure of the PH experiment was identical to that previously used in healthy participants\textsuperscript{37}. While standing and blindfolded, participants manipulated the robot for three minutes and were presented with synchronous or asynchronous haptic feedback. At the end of each session, participant filled an 8-item questionnaire regarding the sensations elicited by the manipulation (see Figure 1, top), including 3 experimental questions aimed at assessing illusory states (see Q2, Q6,Q8; see supplementary material), and 5 control questions (Q1,Q3,Q4,Q5,Q7), added to control for spurious effects, such as general susceptibility and response biases. Synchronous and asynchronous conditions were presented in a randomized order.

Auditory Misattribution experiment

Word recordings (50 in participant’s voice and the same 50 words in a gender matched voice) were randomly grouped in four blocks of twenty-five words. Blindfolded participants freely manipulated the robot with their left hand throughout the experimental session (Figure 1, bottom). These movements were replicated in real time to the participants’ backs by the “follow” robot. After 1 minute moving the lead robot while they received synchronous or asynchronous (500 ms delay) haptic feedback, and while still stroking their backs, participants heard the prerecorded words and were required to report if the word was spoken in their own voice (Self condition) or in another person’s voice (Other condition). An interstimulus interval of 0.5, 0.6 or 0.7 sec was presented between words, and the next word were presented only after the participant’s response. Reaction time and accuracy were measured for each trial. The synchrony of sensorimotor feedback was
pseudo-randomized between subjects. There were four blocks in total (2 synchronous and 2 asynchronous). Thus, all participants heard the same words in both their own voice and another’s voice and under both synchronous and asynchronous sensorimotor conditions.

**Statistical analysis**

**Presence Hallucination experiment**

Ordinal responses to the PH questionnaire were analyzed using linear mixed effects models, with synchrony (synchronous/ asynchronous), group (Control/PE+/PE-) and question type (experimental vs. control) as fixed effects, and subjects and questions as random effects. This was followed by nonparametric Mann-Whitney-Wilcoxon (MWW) tests on the PH experimental questions.

**Auditory Misattribution Experiment**

We used signal detection theory to analyze the responses to the auditory misattribution task. Responses were categorized as hits, misses, false alarms and correct rejections for synchronous and asynchronous conditions. D’ measures of perceptual sensitivity, as well as beta measures for subjective criterion changes, were then calculated for each condition separately and analyzed using a Synchrony(Synchronous/Asynchronous) X Group(Control/PE+/PE-) Analysis of Variance (ANOVA), with synchrony as a within subject variable and Group as a between subject factor.
Results

Presence Hallucination experiment

As expected the mixed model ANOVA indicated a significant interaction between synchrony and question type (F(1,808)=5.62, p=.01), showing that experimental questions measuring the PH were more modulated by synchrony than the control questions. A main effect of synchrony (F(1,808)=5.82, p=.01) was also found, showing overall higher ratings in the synchronous, as compared to asynchronous condition; no other effects or interactions were significant (all F<.75). Contrary to our expectations no differences in the PH were found between the PE+ and control groups. The analysis of experimental questions indicated that participants experienced the touch in the synchronous condition to be self-generated (Q2, p=.003) and experienced the asynchronous touch as originating from another person (Q6, p=.0005). Critically, participants reported a higher presence hallucination in the asynchronous condition than in the synchronous condition (Q8, p=.000007). No other effects reached significance (all p>0.15).

Auditory Misattribution Experiment

Overall accuracy rates were high (M=84.99, 95% CI=+/− 0.05) and did not differ between the groups (F(2, 44)=1.28, p=.28), indicating that all participants were able to perform the self-recognition task with good accuracy. The ANOVA on d’ scores revealed a significant interaction of Group and Synchrony (F(2, 44)=6.68, p=.002, partial η²=0.23). Post hoc analysis indicated that this was driven by reduced d’ for the PE+ group in the asynchronous condition (M=2.25, 95%
CI=±/− 0.45) compared to the synchronous condition (M=2.77, 95% CI=±/− 0.47, p=.0003). Thus, the asynchronous stimulation decreased the PE+ patients’ ability to discriminate their own voice from another’s voice. This difference in d’ for the PE+ group in the asynchronous condition, resulted from both a reduction in the hit rate (0.85 to 0.79) as well as a rise in the rate of false alarms (0.1 to 0.13). This indicates a reduction of auditory self-discrimination for both self and other voices. There was no significant main effect of Group (F(2, 44)=0.95, p=.39), nor any main effect of Synchrony (F(1, 44)=1.1, p=.28; Figure 2), indicating that this decrease was specific to the PE+ group during the asynchronous condition. This difference in sensitivity could not be explained as a consequence of a subjective change in the criterion, as the ANOVA run on beta values showed no main effect of Group (F(2, 44)=0.57, p=.56), no main effect of Synchrony (F(1, 44)= .59, p=.44) nor a significant interaction (F(2, 44)=0.24, p=.78). While not a primary outcome, reaction times were also recorded and analyzed (see Supplementary material and supplementary figure 1 for further analysis).

Discussion

Here we show that robotically-induced sensorimotor conflicts can experimentally generate the PH, extending previous findings from healthy participants to early psychosis patients. Critically, the same sensorimotor conflict induced a decrease self-monitoring during the auditory-verbal task, only observed in PE+ patients, who present symptoms of external control such as AVH and other passivity experiences.

These findings indicate that experimentally induced, task irrelevant sensorimotor conflict can cause source-monitoring errors for auditory-verbal stimuli in PE+ patients, supporting cognitive models
of schizophrenia which state that deficits in self-related processing, are related to deficient sensorimotor prediction\textsuperscript{5,23,42}. While influential previous research has shown that schizophrenia patients have reduced self-monitoring and sensory attenuation for self-generated actions\textsuperscript{17,19,20,31}, this is, to the best of our knowledge, the first study to show experimentally a causal link between a task irrelevant sensorimotor conflict and the induction of an online auditory-verbal misattribution. Interestingly, this effect is in line with a decreased ability for self-other discrimination, as discrimination errors were present for both one’s own voice and for the voice of another person. Importantly, this effect was not due to changes in the response criterion or to general differences in performance between the psychosis and healthy individuals, and was not observed in the experimental control condition. This work has some limitations that should be mentioned. While the PH was found in all participants, no differences were found in the subjective experience of the PH between groups. This may stem from reduced sensitivity of subjective, explicit reports of the PH versus more objective measures of its effects.

Previous work has shown that the brain network related to PHs is comprised of temporoparietal, insular and fronto-parietal nodes, which are key regions in a network of multisensory and sensorimotor areas underlying bodily self-consciousness\textsuperscript{43}. This was revealed using lesion mapping in patients with PHs caused by neurological disease\textsuperscript{37}. In particular, the temporoparietal junction is a multisensory region critical for self-location and first-person perspective, and neurologically or electrophysiologically induced alterations of this area have been shown to induce altered states of self-consciousness such as so-called out-of-body experiences and PH\textsuperscript{44,45}. The insular cortex is considered a fundamental cortical hub integrating interoceptive and exteroceptive stimuli\textsuperscript{45–49} and is involved in bodily self-representation\textsuperscript{49,50} and source monitoring\textsuperscript{51}. Insular lesions have been linked to distortions in bodily experience, such as somatoparaphrenia\textsuperscript{50} and heautoscopy\textsuperscript{52}. The
insular cortex is also altered in schizophrenia\textsuperscript{51,53,70,71} and is related to auditory hallucinations\textsuperscript{26,54}. Although not tested directly in the present study, future studies may reveal that the regions causing PHs in neurological patients, are also of relevance for PH in psychiatric patients as well as self-motoring deficits and related symptoms, including AVHs in schizophrenia.

To conclude, this work presents a novel link between sensorimotor conflicts and self-monitoring in psychosis. We show that sensorimotor conflicts can induce not only a sensation of an external agent (PH), but also that such conflicts propagate to other sensory and cognitive processes causing self-monitoring errors in the auditory-verbal domain. Using a novel approach combining robotics, behavioral paradigms, and psychophysics we show for the first time that psychotic-like symptoms can be induced by sensorimotor conflicts in patients and thus relate to processes known to alter the representation of the bodily self. These findings open the way for novel explorations of the mechanisms of different levels of self-disorder in psychosis.

References


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Author Contributions: Dr. Salomon & Dr. Progin had full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis.

Study concept and design: Salomon, Progin, Blanke, Conus, Griffa, Serino, Rognini, Marchesotti.

Acquisition of data: Progin, Salomon, Blanke, Conus, Griffa, Do, Hagmann, Serino, Rognini, Marchesotti.

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Figure Captions

Figure 1. Experimental design of the Presence Hallucination and the Self-Monitoring experiment.

Top figure. Presence Hallucination (PH) experimental design. Participants moved the master robot of the robotic system while receiving haptic feedback on their back from the “follow” robot. Critically, the tactile feedback could be either temporally synchronous to their movements or asynchronous through the introduction of a 500 ms delay. After 180 seconds of stroking participants responded to the PH questionnaire. This was repeated for both synchronous and asynchronous conditions, in randomized order.

Bottom figure. Auditory misattribution experimental design. Participants moved the master component of the robotic system while receiving haptic feedback on their back from the “follow” component. Again, the tactile feedback could be either temporally synchronous to their movements or asynchronous. After 60 seconds of stroking participants heard words either in their own voice or in the voice of another person and were required to judge if the word was in their own voice or that of another person. Each block included 25 words and was repeated four times.

Figure 2. Induction of auditory-verbal Self-monitoring deficits by sensorimotor conflict.

d’ scores in Self-Monitoring for synchronous and asynchronous feedback conditions for all experimental groups are shown. Sensorimotor conflicts induced a decrease of self-other discrimination for auditory-verbal stimuli, but only for the PE+ group. Note that overall discrimination was not different between patients and controls.
**Figure 3. Induction of a Presence Hallucination by sensorimotor conflict.**

Subjective ratings indicated that the sensorimotor conflict in the asynchronous versus synchronous feedback condition induced a feeling of a presence in all experimental groups (left plots, controls; middle plot PE+ patients; right plot, PE- patients), extending previous findings in healthy participants to psychosis patients. Contrarily, synchronous stimulation induced a feeling of self-touch.

**Table 1**

Demographic and clinical information for all participant groups.

**Supplementary Figures**

**Supplementary Figure 1**

Title. Reaction times for all groups and all experimental conditions. Errorbars denote standard error of the mean (SEM).