Automated analysis of free speech predicts psychosis onset in high-risk youths

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BACKGROUND/OBJECTIVES: Psychiatry lacks the objective clinical tests routinely used in other specializations. Novel computerized methods to characterize complex behaviors such as speech could be used to identify and predict psychiatric illness in individuals.

AIMS: In this proof-of-principle study, our aim was to test automated speech analyses combined with Machine Learning to predict later psychosis onset in youths at clinical high-risk (CHR) for psychosis.

METHODS: Thirty-four CHR youths (11 females) had baseline interviews and were assessed quarterly for up to 2.5 years; five transitioned to psychosis. Using automated analysis, transcripts of interviews were evaluated for semantic and syntactic features predicting later psychosis onset. Speech features were fed into a convex hull classification algorithm with leave-one-subject-out cross-validation to assess their predictive value for psychosis outcome. The canonical correlation between the speech features and prodromal symptom ratings was computed.

RESULTS: Derived speech features included a Latent Semantic Analysis measure of semantic coherence and two syntactic markers of speech complexity: maximum phrase length and use of determiners (e.g., which). These speech features predicted later psychosis development with 100% accuracy, outperforming classification from clinical interviews. Speech features were significantly correlated with prodromal symptoms.

CONCLUSIONS: Findings support the utility of automated speech analysis to measure subtle, clinically relevant mental state changes in emergent psychosis. Recent developments in computer science, including natural language processing, could provide the foundation for future development of objective clinical tests for psychiatry.

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INTRODUCTION

The capacity of psychiatry to diagnose and treat serious mental illness has been hampered by the absence of objective clinical tests of the type routinely used in other fields of medicine. Although recent years have seen substantial advances in understanding of the neurobiology of mental illness, these developments have yet to yield markers that reliably differentiate psychiatric health from illness at the level of the individual patient. Whereas clinical neuroscience has focused on the brain in mental illness, computer science has, in parallel, developed increasingly sophisticated automated approaches to characterize and predict human behavior. Such advances are now commonly utilized in industry (the private business sector): models combining demographic data and purchasing behavior are used to personalize advertising content and automated language assessment is employed to screen job candidates and score essays. The degree to which such technologies might also aid diagnosis and prognosis in psychiatry is only now beginning to be explored (e.g., see ref. 4).

Developments in automated natural language processing present one promising avenue for psychiatry. Although speech may present a unique ‘window into the mind’ in a variety of altered states, it is particularly relevant to psychosis. Thought disorder, a cardinal symptom of schizophrenia in which thought processes lose coherence, is typically diagnosed on the basis of clinical observation of disorganized speech. As a complement to clinical observation, automated analysis methods have previously been used to assess speech correlates of thought disorder in schizophrenia. For example, Latent Semantic Analysis (LSA), an automated high-dimensional associative analysis of semantic structure in speech, has been used to identify a reduction in semantic coherence in schizophrenia that correlates with clinical ratings and has comparable diagnostic accuracy. LSA combined with structural speech analysis was also able to accurately differentiate between first-degree relatives of schizophrenia patients and unrelated healthy individuals, suggesting that subtle differences indicative of underlying genetic vulnerabilities to schizophrenia can be distinguished with computerized speech analysis.

As yet, however, these methods have not been applied to the prediction of psychosis onset, even though clinically diagnosed subtle disorganization in speech has consistently been identified as predictive of psychosis (i.e., with classification accuracy of ∼60%) among young people identified as at clinical high risk.
elusive. Improving the capacity to predict psychosis among high-risk youths has proven up to 2.5 years in teens and young adults identifying psychosis. Specifically, we hypothesized that automated semantic and syntactic analysis of developing schizophrenia by virtue of subthreshold or attenuated symptoms could not have occurred self or others incommensurate with outpatient care, any major medical or psychological procedures, and interviews took place between 2007 and early 2012, and were transcribed by an independent company. The resultant preprocessed data consisted of a list of lemmatized words, accurately discriminated between patients and controls, we hypothesized that automated semantic and syntactic analysis of baseline interview transcripts would yield speech features capable of predicting subsequent psychosis outcome among CHR individuals. 11

MATERIALS AND METHODS

Participants

Participants were 34 help-seeking youths aged 14 to 27 years who were fluent in English (three were immigrants who learned English as children). They were referred from schools and clinicians, or self-referred through the Center of Prevention and Evaluation website. Exclusion criteria included history of threshold psychosis or Axis I psychotic disorder, risk of harm to self or others incommensurate with outpatient care, any major medical or neurological disorder, and Intelligence Quotient < 70 (assessed with the Wechsler Abbreviated Scale of Intelligence). The attenuated psychotic symptoms characteristic of the CHR participants could not have occurred solely in the context of substance use or withdrawal. Adults provided written informed consent; participants under 18 provided written assent, with consent provided by a parent. All experiments were performed in accordance with the relevant guidelines and regulations, and all procedures were approved by the Institutional Review Board at the New York State Psychiatric Institute at Columbia University. Five participants transitioned to psychosis within 2.5 years of follow-up (CHR+), whereas 29 did not (CHR–). Demographics for CHR individuals, stratified by psychosis outcome, are presented in Table 1.

Table 1. Demographics

|                  | CHR+ (N = 5) | CHR– (N = 29) |
|------------------|--------------|---------------|
| Age (in years)   | 22.2 (3.4)   | 21.2 (3.6)    |
| Gender (%) male  | 80%          | 66%           |
| Race (%) Caucasian | 40%        | 38%           |
| Medications prescribed (antipsychotics and/or antidepressants) | 20% | 21% |

Abbreviations: CHR+, clinical high-risk participants who transitioned to psychosis during follow-up; CHR–, clinical high-risk participants who did not transition to psychosis during follow-up.

Procedures

Ascertainment and prospective characterization. The Structured Interview for Prodromal Syndromes/Scale of Prodromal Symptoms (SIPS/SOPS)3–9 was used for ascertainment of CHR status, for baseline and quarterly symptom ratings,10 and to determine psychosis outcome. The SIPS/SOPS evaluates positive (subthreshold psychotic), negative, disorganized, and general symptoms.

Participants had to meet baseline criteria for one of three prodromal syndromes, assessed with the SIPS/SOPS: (i) attenuated positive symptom syndrome (≥1 SOPS-positive item in the prodromal range with symptoms beginning or worsening in the past year, and symptoms occurring ≥ once/week in the prior month); (ii) genetic risk and deterioration syndrome (psychosis in a first-degree relative or schizotypal disorder accompanied by a 30% drop in global assessment of function over the past year); or (iii) brief intermittent psychotic symptom syndrome (≥1 SOPS-positive items in the psychotic range with symptoms beginning in the past 3 months, and symptoms occurring ≥ several minutes/day). All CHR participants in this study met criteria for the attenuated positive symptom syndrome. Trained master-level research assistants administered the SIPS/SOPS, with clinical ratings achieved by expert consensus (with CC).

Participants were prospectively characterized for symptoms every 3 months for up to 2.5 years, with transition to psychosis determined using the SIPS/SOPS ‘presence of psychosis’ criteria.

Baseline interviews. Open-ended, narrative interviews of ~ 1 h were obtained from participants by interviewers trained by an expert in qualitative interviewing and phenomenological research.15 Participants were encouraged to describe changes they had experienced and the impact of these changes, what had been helpful or unhelpful for them, and their expectations for the future. Interviews took place between 2007 and early 2012, and were transcribed by an independent company. The first 27 transcripts were previously subject to thematic analysis using phenomenological procedures, finding gender differences in themes; this earlier qualitative analysis did not assess the predictive value of the interviews for psychosis outcome.16

Speech preprocessing. Interview transcripts were preprocessed as previously described6 using the Natural Language Toolkit (NLTK; http://www.nltk.org/). After discarding punctuation, each interview was automatically parsed into phrases. Words were then converted to the roots from which they are in productively related words, then the semantic similarity of two words can be computed by semantic similarity measures, we used the SIPS/SOPS positive item in the psychotic range with symptoms beginning in the past 3 months, and symptoms occurring ≥ several minutes/day). All CHR participants in this study met criteria for the attenuated positive symptom syndrome. Trained master-level research assistants administered the SIPS/SOPS, with clinical ratings achieved by expert consensus (with CC).

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(CHR) for psychosis (reviewed in ref. 10), as well as those at genetic high risk for psychosis.11 There are several reasons to test automated prediction approaches in this population. Schizophrenia, although relatively rare (lifetime prevalence ~ 1%), is among the most catastrophic mental illnesses both personally and societally. Schizophrenia and related psychotic illnesses typically emerge in young adults at the point of maximal societal and parental investment when individuals are poised to begin to contribute socially and economically.12 Although those at CHR for developing schizophrenia by virtue of subthreshold or attenuated psychotic symptoms can be identified,13 to date reliable prediction of psychosis onset among high-risk youths has proven elusive. Improving the capacity to predict psychosis among high-risk populations would have important ramifications for early identification and preventive intervention, potentially critically altering the long-term life trajectory of people with emergent psychotic disorders.

Here, we present a proof-of-principle test of automated speech analysis to predict, at the level of the individual, the later onset of psychosis. Specifically, we employed analysis of free speech at baseline to predict psychosis onset over a subsequent period of up to 2.5 years in teens and young adults identified as at CHR for psychosis.13 On the basis of earlier findings in schizophrenia,3,9,14 in which automated text analyses yielded parameters that accurately discriminated between patients and controls, we hypothesized that automated semantic and syntactic analysis of baseline interview transcripts would yield speech features capable of predicting subsequent psychosis outcome among CHR individuals.

Automated analysis of free speech

To automatically analyze speech, we used the SIPS/SOPS (with CC).

Speech analyses. We employed a novel combination of semantic coherence and syntactic assays as predictors of psychosis transition. For the semantic analyses, we used a well-validated approach to automated text analysis previously used to analyze speech in schizophrenia,3 LSA17. LSA is a high-dimensional associative model that rests on the premise that word meaning is a function of the relationship of each word to every other word in the lexicon. If semantically similar words co-occur in texts with consistent topics more frequently than do unrelated words, then the semantic similarity of two words can be quantitatively indexed by the frequency of their co-occurrence in a sufficiently large corpus of texts.17 LSA thus captures the meaning of words through linear representations in high-dimensional (300–400 dimensional) semantic spaces based on word co-occurrence frequencies. Each word in the lexicon is assigned a vector representing its semantic content; the orientation of these vectors can then be used to compare semantic similarity between words.17

Here, LSA was trained on the Touchstone Applied Science Associates (TASA) Corpus, a collection of educational materials compiled by TASA. The semantic coherence measure we developed is similar to that used by Elvégård et al.,3 which discriminated between established schizophrenia patients and controls. The present measure differs from the earlier approach in that it explicitly incorporates syntactic information: semantic trajectories are represented by similarity among pairs of consecutive phrases, or pairs of phrases separated by an intervening phrase (see Figure 1). Given the speech transcription D, the document is split into n phrases S_i and converted into a vectorial representation by replacing each word in the phrase by its corresponding LSA vector, s_i → [w_1, ..., w_k]. The
phrase vectors are then summarized by taking the mean of their components:

\[ L_i = \frac{1}{N} \sum_{k=1}^{N} l_{ik} \]

i.e., the mean of all LSA vectors of every word in the phrase.

We defined first-order coherence by taking the similarity of consecutive phrase vectors, averaged over all the phrases in the text (represented by \( \langle \rangle \) below):

\[ FOC = \langle \cos (L_i, L_{i+1}) \rangle \]

and second-order coherence by taking the similarity between phrases separated by another intervening phrase, averaged over all the phrases in the text:

\[ SOC = \langle \cos (L_i, L_{i+2}) \rangle \]

With these two features, we were able to characterize semantic coherence by measuring components of the distributions of first- and second-order coherence over the speech samples, including features such as the minimum, mean, median, and s.d.

Thus, we indexed speech coherence by: (i) automated separation of interviews into phrases; (ii) assigning phrases semantic vectors as the mean of the LSA semantic vectors for each word within the phrase; and (iii) assessing semantic similarity (i.e., the cosine) between the phrase vectors of consecutive phrases, or phrases separated by another intervening phrase.

To complement the semantic analysis, we defined another measure for processing the documents, on the basis of Part Of Speech tagging (POS-Tag). This consists of labeling every word by its grammatical function. For example, the sentence ‘The cat is under the table’ is tagged by the POS-Tag procedure as (‘The’, ‘DT’), (‘cat’, ‘NN’), (‘is’, ‘VBZ’), (‘under’, ‘IN’), (‘the’, ‘DT’), (‘table’, ‘NN’). Where DT is the tag for determiners, NN for nouns, VBZ for verbs, and IN for prepositions. For every transcript, we calculated the POS-Tag information (with NLTK\(^5\)) and used the frequencies of each tag as an additional attribute of the text. Tagging automation uses a hand-tagged corpus to train a parsing process using a variety of heuristics. NLTK uses a model called Pen Tree Bank.

Code availability. Code for speech preprocessing (WordNet lemmatizer) and POS-Tag (Pen Tree Bank) is available open access through the NLTK (http://www.nltk.org/).\(^5\)

Classification. A cross-validated classifier is a Machine Learning algorithm with two stages: in the first stage, it learns the underlying patterns of the data using a subset of samples. The learned model is used in the second stage to predict the labels of samples not used during the learning stage (Figure 2).

We used features derived from the semantic coherence analyses and the POS-Tag extraction, providing a vector of features for each participant’s text. With this information, we trained the classifier to learn the features that discriminated among participants who did not subsequently develop psychosis (CHR –) from the group who did (CHR+).

The convex hull of a set of points is the minimal convex polyhedron that contains them. A convex hull classifier was implemented as follows: during training, we sequentially excluded one CHR+ or CHR – participant to be used for testing (leave-one-subject-out cross-validation). Using the training labels, we computed the convex hull of the CHR – set, and then tested whether the left-out sample was inside the hull (predicting CHR –) or outside (predicting CHR+). Each individual was sequentially excluded from the training set used to compute the convex hull to serve as the test subject, providing accuracy of prediction data for all participants.

The semantic coherence feature that best contributed to classification of subsequent psychosis onset was the minimum coherence between two consecutive phrases (i.e., the maximum discontinuity) that occurred in the interview. The syntactic measure included in classification was the frequency of use of determiners (‘that’, ‘what’, ‘whatever’, ‘which’, and ‘whichever’), normalized by the phrase length. Because speech in emergent psychosis often shows marked reductions in verbosity (referred to clinically as poverty of speech), we also included the maximum number of words per phrase in the classification.

Validation. To further probe findings from the CHR analyses, we also conducted the following validation analyses:

Does the coherence measure index ‘disorder’in a text?: Because the concept of semantic coherence we employed does not have a
mathematical definition, in this validation we tested the coherence measure against a corpus of classic literature and assessed how the measure changed when we modified the original texts in a way that is relevant to the concept of semantic coherence.

On the basis of the hypothesis that a text that makes sense will produce a high coherence score, we applied different levels of ‘disorder’ to a range of texts to determine whether the method could detect these modifications. We defined each level of ‘disorder’ as the percent of the text that was moved from its original position. For example, a disorder level of 40% indicates that 4 of 10 sentences were moved and thus were no longer in their original position in the text. For each of 10 disorder levels (10–100%), we created 1,000 samples, randomly shuffling the order of the appropriate proportion of sentences. We performed coherence analysis on randomly selected chapters of the following six classic books: On the Origin of Species by Charles Darwin, A Study in Scarlet by Arthur Conan Doyle, Moby Dick; Or, The Whale by Herman Melville, Pride and Prejudice by Jane Austen, The Adventures of Tom Sawyer by Mark Twain, and The Count of Monte Cristo by Alexandre Dumas.

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**Table 2. Classification performance metrics**

| Classification         | PPV | NPV | Sens. | Spec. | ROC  |
|------------------------|-----|-----|-------|-------|------|
| Convex Hull 3-feature  | 100 | 100 | 100   | 100   | 1.00 |
| SIPS/SOPS              | 33  | 89  | 40    | 86    | 0.47 |

Abbreviations: NPV, negative predictive value; PPV, positive predictive value; ROC, receiver operating characteristic area under the curve; Sens, sensitivity; SIPS/SOPS, classification based on baseline scores on the Structured Interview for Prodromal Syndromes/Scale for Prodromal Symptoms; Spec, specificity.

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Are the speech features associated with symptoms assessed with standard diagnostic instruments?: To assess the extent to which the text features that best predicted clinical status at follow-up in CHR patients (minimum first-order coherence, density of determiners, and maximum phrase length) carry information with respect to standard clinical prodromal ratings, we computed the canonical correlation between these three text features (semantic coherence, phrase length and use of determiner pronouns) and two symptom measures on the SIPS/SOPS (total positive symptoms and total negative symptoms). The canonical correlation between two sets of features from the same samples, X and Y, estimates the linear combination of X features such that this combined feature has the highest correlation with an also estimated linear combination of Y features.

Ethics statement: The Institutional Review Board at the New York State Psychiatric Institute at Columbia approved these experiments, and informed consent was obtained for all subjects (parental consent with assent for minors).

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**RESULTS**

**CHR analysis**

Of the 34 participants, 5 were known to develop schizophrenia (or schizoaffective disorder) within 2.5 years. Respectively, their times to psychosis onset from time of speech sampling were 3, 4, 8, 12, and 16 months. Twenty-nine participants were known to not develop psychosis over follow-up, with 22 of these participants followed for 2.5 years, 4 participants followed for 2 years, and 3 followed for 1.5 years (these participants’ CHR status was ascertained closer to the end of the overall study). An additional participant’s transcript was not included in speech analyses because her clinical outcome was indeterminate; she remained psychosis-free over 1.5 years of follow-up, but may have subsequently developed psychosis after the study.

A graphical representation of the differentiation obtained between CHR+ and CHR− individuals using the three parameters of minimum semantic coherence, normalized use of determiners, and maximum phrase length is presented as the convex hull of the set of CHR− individuals (the minimal convex polyhedron that contains all data points) in Figure 3. The convex hull of CHR− individuals does not include any CHR+ individuals.

The convex hull classifier yielded 100% accuracy for prediction of psychosis onset. Null hypothesis tests were used to estimate the probability of obtaining this result by chance. We first partitioned the data set (N = 34) randomly, assigning five subjects to the CHR+ label and the remainder to the CHR− group. Because some assignments for this initial test included the actual CHR+ individuals, we implemented a second test by repeating the previous scheme, including only CHR− individuals. That is, using the CHR− set, we randomly assigned CHR+ labels to 5 CHR− individuals, and estimated the probability that they would all fall outside the remaining 24 individuals randomly labeled as CHR−. Finally, we repeated the same scheme by assigning random labels to the 29 CHR− individuals (matching the original number of labels), and also randomly assigning the semantic and syntactic speech features, drawing values from a Gaussian distribution with...
the same mean and s.d. as the actual values. In each scheme, the probability that all five individuals labeled as CHR+ would fall outside the convex hull of CHR− individuals was less than chance, i.e., \( P < 0.05 \).

To investigate whether standard clinical ratings could differentiate CHR+ and CHR− individuals, we entered variables from clinical ratings—the SIPS/SOPS\(^1\)3 into several classifiers. The best prediction obtained was less accurate than the automated analysis, misclassifying 3 of 5 CHR+ patients and 4 of 29 CHR− patients to yield an accuracy of 79%, consistent with prior studies (see Table 2 for classification performance metrics).

**Validations**

The coherence measure as an index of ‘disorder’ in texts. We found that two features of the semantic coherence distributions, the minimum semantic distance for first-order coherence (i.e., the minimum coherence or maximum discontinuity between two adjacent sentences within the text sampled), and the mean semantic distance for first-order coherence (i.e., the average coherence between adjacent sentences within the text) were negatively correlated with the disorder level we produced in texts, indicating that higher levels of disorder within the text produced lower coherence scores (see Figure 4).

**Associations between speech features and symptoms assessed with standard diagnostic instruments.** The canonical correlation analysis of text features versus the entire set of clinical prodromal features did not yield any significant correlation; however, restricting the analyses to the sums of subthreshold psychotic and negative symptom severity ratings (i.e., \( A_{\text{total}}, B_{\text{total}} \)) yields a correlation of \( r = 0.57 \) and \( P = 0.046 \), for the variables \( s \) (symptoms) and \( t \) (speech variables; Figure 5):

\[
\begin{align*}
    s & = 0.066 \times A_{\text{total}} + B_{\text{total}} - 0.02 \times \text{coherence} - 0.54 \times \text{determiners}. \\
    t & = -0.68 \times \max(\text{words per phrase})
\end{align*}
\]

In this equation, there are two symptom variables (sums of subthreshold psychotic and negative symptoms, respectively,
Of note, the sample size employed in this initial study was small, with five participants developing psychosis during the follow-up period. This limitation meant that we were unable to divide participants into separate training and test samples, instead using cross-validation procedures to assess the predictive algorithm. This approach, although providing important information about the potential predictive capacity of these novel speech measures, may have resulted in higher estimates of the predictive accuracy of the model than would be obtained in a larger, separate sample. Thus, replication in a larger sample will be an important future research direction.

Our findings from this proof-of-concept study, although needing to be replicated in larger samples, have several implications. First, reliable identification of individuals likely to progress to schizophrenia would greatly facilitate targeted early intervention. Second, automated speech assessment, if further validated, could provide previously unavailable information for clinicians on which to base treatment and prognostic decisions, effectively functioning as a ‘laboratory test’ for psychiatry. The ease of speech recording makes this approach particularly suitable for clinical applications. Self-report of symptoms, on which much of psychiatric assessment relies, depends on the patient’s motivation and capacity to accurately report their introspective experiences, which may be influenced by psychiatric illness. Although clinicians routinely detect disorganized speech on the basis of clinical observations, our data suggest that automated analytic methods allow for superior assessment. As a direct, objective measure, automated speech analysis could thus provide important information to complement existing methods for patient assessment. Finally, these findings support the use of advanced computational methods to characterize complex human behaviors such as speech in both normal and pathological states. Such a fine-grained behavioral analysis could allow tighter mapping between psychiatrically relevant phenotypes and their underlying biology, in essence carving nature more closely at its joints. Better mapping between the behavioral and the biological is likely to lead to greater understanding of the pathophysiology of schizophrenia and other psychiatric disorders, potentially also informing psychiatric nosology.

These findings represent the initial stages in the use of emerging computer science behavioral analysis techniques, already prominently used in industry, to characterize and predict human behavior in the context of psychiatric health and illness. Using automated approaches, we were able to extract indices of speech-semantic coherence and syntax and use these to accurately predict the subsequent development of psychosis in high-risk youths. Prognostic prediction using this approach outperformed prediction on the basis of standard psychiatric ratings. Computerized analysis of complex human behaviors such as speech may present an opportunity to move psychiatry beyond reliance on self-report and clinical observation toward more objective measures of health and illness in the individual patient.

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CONTRIBUTIONS
GB contributed to the conception of the study, the interpretation of data, and drafting the manuscript. CMC led the prospective clinical high-risk cohort study and overview all data collection, and worked on and edited iterative drafts of the manuscript. DCJ also contributed to the design and conduct of the cohort study, and contributed suggested edits to the manuscript. FC and GAC designed and performed the automated text analysis; DFS and MS contributed to the analysis of the data; NM, SR, and MC collected and preprocessed data on patients with schizophrenia and their controls. All the authors reviewed the results, edited the
manuscript, and gave final approval for submission of the manuscript. Drs Corcoran and Cecchi had full access to all the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis.

COMPETING INTERESTS
The authors declare no conflict of interest.

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Supplementary Information accompanies the paper on the npj Schizophrenia website (http://www.nature.com/npjschz)