



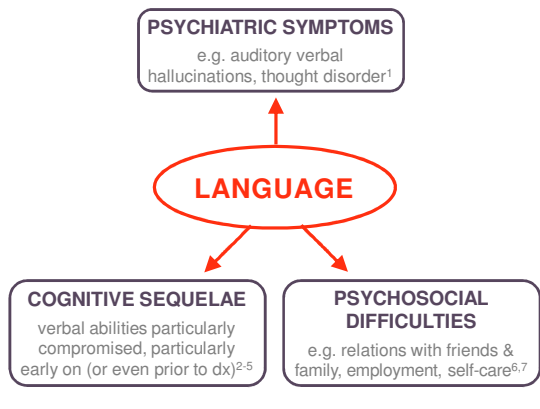
From speech to meaning: Abnormal predictive processing in schizophrenia



Meredith Brown & Gina R. Kuperberg

MGH/MIT/HMS Athinoula A. Martinos Center for Biomedical Imaging; Department of Psychiatry, Massachusetts General Hospital; Department of Psychology, Tufts University

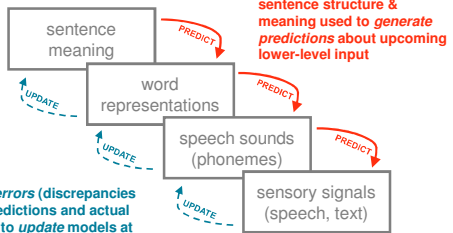
Why study language in schizophrenia?



Hierarchical generative framework

Central role for **predictions** in normal language processing⁸⁻¹¹

Goal: Optimal inference of intended message, given available information



Prediction errors (discrepancies between predictions and actual input) used to update models at successively higher levels

Explains:

- how we simultaneously take multiple sources of context (such as visual scene, discourse history, who we are talking to) into account during language processing
- how we rapidly and flexibly adapt to (and keep up with) new speakers & situations
- **abnormalities in multiple aspects of language processing in schizophrenia?**

References & acknowledgments

1. American Psychiatric Association (2013). DSM-V.
2. Bowers & Henry (2005). *The Psychiatric Clinics of North America*, 28, 613-633.
3. Fuller et al. (2002). *American Journal of Psychiatry*, 159, 1183-69.
4. Kell et al. (1999). *American Journal of Psychiatry*, 156, 1154-61.
5. Simon et al. (2007). *Schizophrenia Bulletin*, 33, 761-71.
6. Bowers & Henry (2009). *Schizophrenia Research*, 102, 234-7.
7. Razoustan et al. (1999). *The Journal of Nervous and Mental Disease*, 187, 281-9.
8. Farmer et al. (2013). *Behavioral and Brain Sciences*, 36, 211-2.
9. Kuperberg (2013). In Miller, Cullins & McClelland (Eds.), *Unraveling the Behavioral, Neurobiological, & Genetic Components of Reading Comprehension* (p. 176-92). Kuperberg (in press). *Cognition*.
10. Liberman (2014). Ph.D. dissertation, University of Rochester.
11. Kuperberg (2014). *Language and Linguistic Theory*, 30, 576-609.
12. Kuperberg (2010). *Language and Linguistic Theory*, 26, 459-604.
13. Kuperberg et al. (1998). *Journal of Abnormal Psychology*, 107, 423-34.
14. Liberman et al. (1994). *Biological Psychiatry*, 34, 864-77.
15. Dinan & Kuperberg (2010). *Journal of Neurolinguistics*, 23, 254-69.
16. Chapman (1960). *The Journal of Abnormal and Social Psychology*, 60, 412-4.
17. Tross et al. (2005). *Journal of Abnormal Psychology*, 110, 761-7.
18. Marschreck et al. (1998). *Schizophrenia Research*, 1(1), 61-6.
19. Sabin et al. (1994). *Biological Psychiatry*, 34, 864-77.
20. Marz et al. (2001). *Schizophrenia Research*, 48, 301-5.
21. Weisbard et al. (1998). *The British Journal of Psychiatry*, 172, 142-6.
22. Kuperberg et al. (2013). 28th CUNY Conference on Human Sentence Processing.
23. Tumbarello et al. (2009). *Schizophrenia Research*, 105, 27-37.
24. Todd et al. (2013). *Frontiers in Psychiatry*, 4, 171.
25. Cienfuegos et al. (1998). *Biological Psychiatry*, 43, 82-8.
26. Butler et al. (2001). *American Journal of Psychiatry*, 158, 1126-33.
27. Holcomb et al. (1998). *Psychiatry Research*, 57, 75-82.
28. Polunowicz et al. (2005). *Archives of General Psychiatry*, 57, 1149-65.
29. Kasa et al. (2002). *American Journal of Psychiatry*, 159, 546-53.
30. Javitt (2005). *Annual Review of Clinical Psychology*, 5, 249-75.
31. Javitt & Friedman (2010). *American Journal of Psychiatry*, 172, 17-31.
32. Lamm et al. (2005). *Biological Psychiatry*, 58, 56-61.
33. Kariwala et al. (2014). *Psychological Medicine*, 44, 25-36.
34. Reichen et al. (2014). *American Journal of Psychiatry*, 171, 349-59.
35. Uhlhaas & Singer (2009). *Psychological Bulletin*, 131, 618-52.
36. Silvestri et al. (1996). *Journal of Abnormal Psychology*, 105, 663-7.
37. Hemsley (1995). *Journal of the Acoustical Society of America*, 29, 117-23.
38. Liberman (1997). *Journal of the Acoustical Society of America*, 29, 117-23.
39. Liberman et al. (1984). *British Journal of Psychology*, 74, 431-61.
40. Liberman & Jungman (1981). *Psychological Review*, 74, 431-61.
41. Liberman et al. (1984). *British Journal of Psychology*, 74, 431-61.
42. McMurray & Jongman (2000). *Nature Reviews Neuroscience*, 3, 489-95.
43. Curo et al. (2000). *Human Brain Mapping*, 3, 183-91.
44. Ford & Matheson (2005). *International Journal of Psychophysiology*, 56, 179-89.
45. Ford et al. (2007). *American Journal of Psychiatry*, 164, 658-66.
46. Allen et al. (2007). *International Review of Psychiatry*, 19, 407-15.
47. Fiechter & Fries (2009). *Nature Reviews Neuroscience*, 12, 48-56.
48. Wykes et al. (2011). *American Journal of Psychiatry*, 168, 472-485.
49. Paddy et al. (2014). *Current Treatment Options in Psychiatry*, 1, 121-33.
50. Medalla & Choi (2009). *Neuropsychology Review*, 19, 353-564.
51. Rowe et al. (2012). *American Journal of Psychiatry*, 169, 710-9.
52. Abouk et al. (2009). *Schizophrenia Bulletin*, 35, 112-41.
53. Fisher et al. (2015). *Schizophrenia Bulletin*, 41, 250-6.

This work was funded by the National Institute of Mental Health (R01MH071635 to GKR).

Understanding language processing abnormalities across domains

Interpreting sentence & word meanings

An apparent paradox:

- Patients have difficulty interpreting sentence and word meaning in context, compared to healthy controls¹²⁻¹⁷

e.g. interpreting "bank" as a river bank vs. a financial institution

- But patients (particularly those with thought disorder) exhibit **faster** automatic spreading activation within networks of semantically related words¹⁸⁻²²

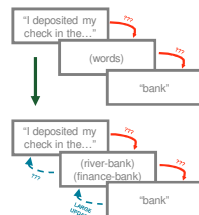
Explanation with generative model framework:

Healthy adults



Predictions constrain interpretation of words to **contextually relevant** meanings and "explain away" the lower-level signal (when accurate)

Patients with schizophrenia



Activation of word meanings is **unchecked** by expectations from sentence or discourse context

Implications for **time-course** of sentence processing: Reliance on slower non-predictive mechanisms likely to disrupt processing under time pressure (as in most normal communicative situations)²³

Perceiving speech sounds

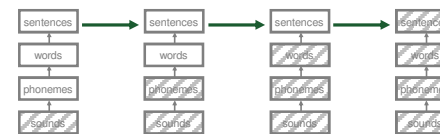
Low-level **sensory & perceptual** changes in schizophrenia, for both speech & non-speech stimuli²⁴⁻³⁰

- behavioral: decreased contrast sensitivity, increased stimulus detection thresholds
- neural: reduced amplitude of evoked responses to speech & non-speech stimuli

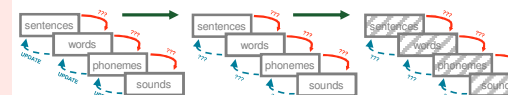
How do perceptual abnormalities relate to higher-order processing*?

*not much work has looked at this

Possibility #1: core problem = perception³¹⁻³⁵



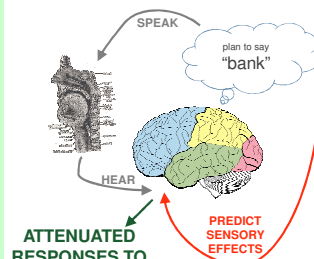
Possibility #2: core problem = generative models



- abnormalities in schizophrenia are much more pronounced when perceiving stimuli in context than when perceiving isolated stimuli³⁶⁻³⁹
- and speech sounds, in particular, are extremely context-dependent⁴⁰⁻⁴²

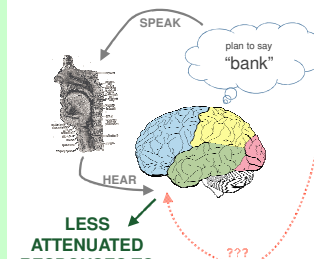
Relating action to perception

Healthy adults



ATTENUATED RESPONSES TO OWN SPEECH⁴³
compared to someone else's speech

Patients with hallucinations



LESS ATTENUATED RESPONSES TO OWN SPEECH⁴⁴⁻⁴⁵

Hypothesis: disruptions in generative models linking self-action to self-perception

- auditory verbal hallucinations may arise from failure to recognize self as source of "inner speech"⁴⁶⁻⁴⁷
- disruptions in these generative models might reflect more general disruptions to abilities to attribute speech to its source (whether internal or external, as with different speakers)
- might also scale up to disordered monitoring of higher-level language production in thought disorder

Implications & directions

Emphasis on **interfaces** between domains

- effects of higher-level context on speech perception
- relations between all these abnormalities *within the same patients*

Implications for cognitive remediation

- cognitive remediation programs **consistently somewhat effective** despite vastly different approaches⁴⁸⁻⁵⁰

approaches focusing on high-level cognition⁵⁰ rebuild generative models via **predictive pathways**

(particularly when linked to higher-order goals via combination with psychosocial therapy or skills training)^{49,51}

approaches focusing on perceptual abilities⁵²⁻⁵⁹ rebuild generative models via **model updating pathways**

possible that an **integrated approach** would have **synergistic benefits**