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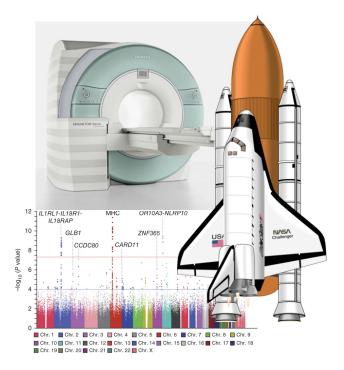


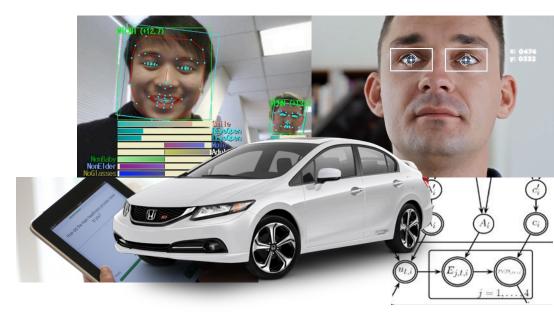
CrossMark

Challenges and promises for translating computational tools into clinical practice Woo-Young Ahn¹ and Jerome R Busemeyer²

Woo-Young (Young) Ahn PI, Computational Clinical Science Lab The Ohio State University June 16, 2016

Mission of the Computational Clinical Science Lab





To maximize prediction accuracy

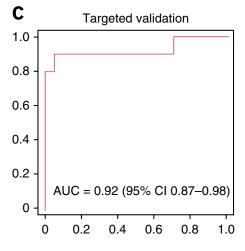
Affordable markers for clinical settings

Can we translate research into practice?

- Current diagnosis system
 - Interviews
 - Behavioral symptoms
- Identify who's at risk?
 Diagnostic tools







Mapstone et al (2014) Nature Medicine

What are the challenges and promises?

A. Precise characterization of latent neurocognitive processes



Models of multiple decision-making systems

Advanced data analytical methods including hierarchical Bayesian analysis

Integration of behavioral and neural data

B. Optimal assays (i.e., paradigms) for assessing psychiatric conditions

Theory-driven approach

Clinical insights w/ external validity

Adaptive design optimization (ADO)

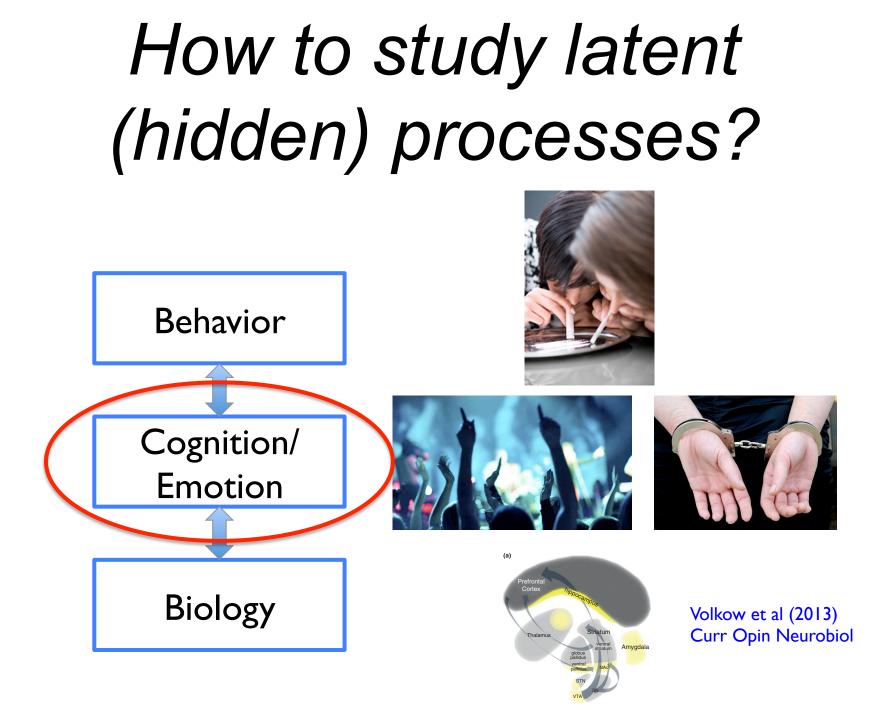
C. Developing large-scale longitudinal studies & generating predictions from multi-modal data



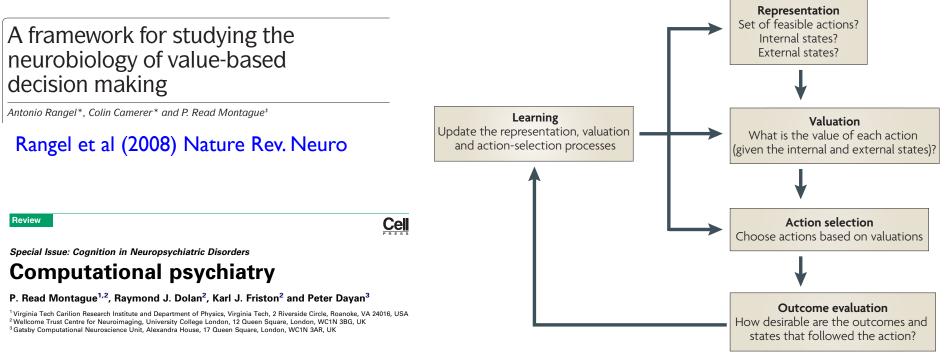
Collaborative consortia and workgroups

Machine learning

(A) Precise characterization of latent neurocognitive processes



Is there a single framework for understanding the mind?

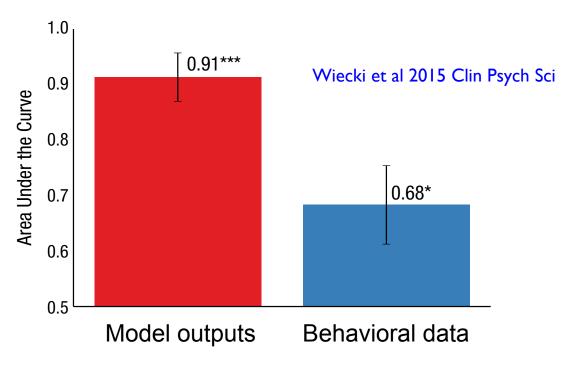


Montague et al (2012) Trends in Cog Sci

Rangel et al (2008) Nature Rev. Neuro

Computational Modeling

More information and greater prediction accuracy



Wiecki et al 2015 Clin Psych Sci Wiecki et al 2016 Plos ONE Fridberg et al 2010 JMP

First started in Mathematical Psychology



William Batchelder Roger Ratcliff Richard Neufeld Teresa Treat Jerome Busemeyer

Batchelder (1998) Psychol. Assessment McFall & Townsend (1998) Psychol. Assessment Ratcliff (2000) PBR + many papers on aging, etc. Neufeld (2002) Psychol. Assessment Treat et al (2001) Psychol. Assessment Busemeyer & Stout (2002) Psychol. Assessment Yechiam et al (2005) Psych. Sci.

Utility of computational modeling for assessing clinical populations

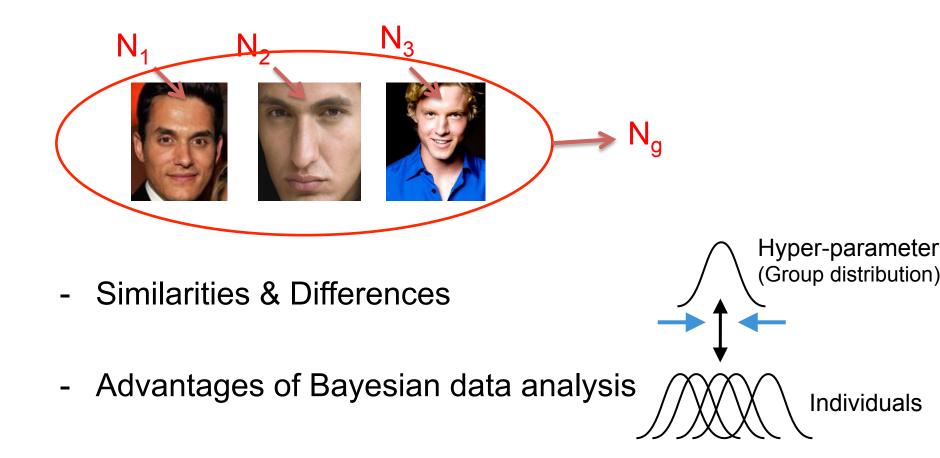
"Computational Psychiatry"



Montague (2012) Trends Cog Sci Dayan et al (2016) Curr Opinion Behav Sci Friston et al (2014) Lancet Huys et al (2015) Clin. Psychol. Sci. Maia & Frank (2011) Nat. Neurosci Wamg & Krystal (2014) Neuron Stephan et al (2015) Neuron

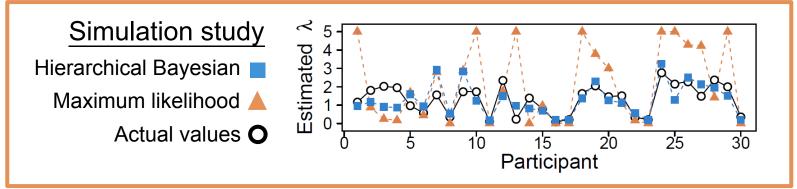
Computational accounts of abnormal cognition & its biological underpinnings

Hierarchical (Bayesian) analysis



Small amount of data from each subject but large N

Hierarchical (Bayesian) analysis

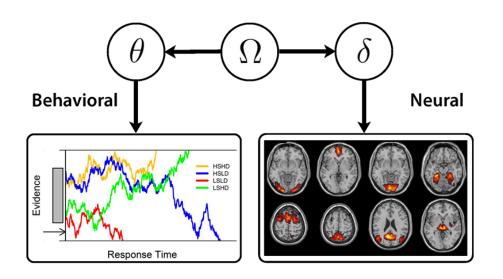


Ahn et al. (2011, JNPE)

(A lot) more room for improvement

- Single vs. multiple hyper groups
- Non-parametric Bayesian
- Data-driven clustering
- Accessibility

We can integrate behavioral and neural data in a single framework



Integrating behavioral and neural data in a single framework

Turner et al (2015) Psych Review Turner et al (2013) Neuroimage

(B) Optimal assays (tasks) for assessing psychiatric conditions

Laboratory tasks



To mimic real-life decisions

Two different camps

- 1. Clinicians/psychologists
 - Develop/adopt emotionally engaging tasks
 - Mimic naturalistic risk-taking behaviors

- 2. Economists/neuroscientists
 - Understanding specific constructs

Clinicians/psychologists

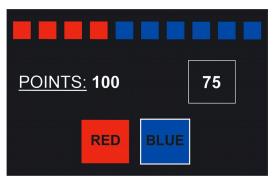
You won \$0.50, but lost \$0.75



Iowa Gambling Task Bechara et al (1999) Cognition



Balloon Analogue Risk Task Lejuez et al (2002) JEP: Applied



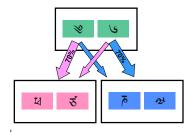
Cambridge Gambling Task

Rogers et al (1999) Neuropsychpharm.

Economists/neuroscientists



Orthogonal Go/Nogo Task Guitart-Masip et al (2012) Neuroimage



Two-Step Task Daw et al (2011) Neuron Voon et al (2014) Mol. Psychiatry



Loss Aversion Task Tom et al (2007) Science De Martino et al (2010) PNAS

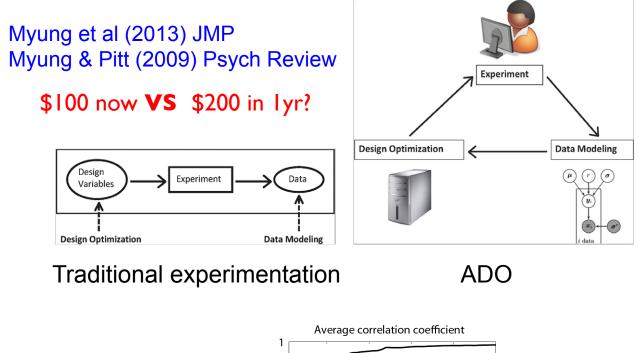
What are the pros and cons?

- Can we disentangle underlying processes?
- Emotionally engaging?
- Realistic for patient populations?

Both clinical expertise & knowledge of neurobiology

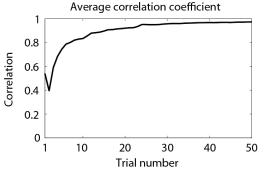
Schonberg et al (2011) TiCS

Can we optimize experimental design?



ADO in action

Cavagnaro et al (2012) Management Sci Cavagnaro et al (in press)



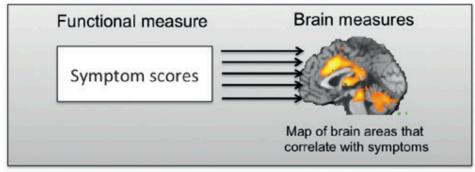
Test-retest reliability Over 0.9 after 15 trials.

Hou et al (2016)

(C) Predictions using multimodal & multi-dimensional data

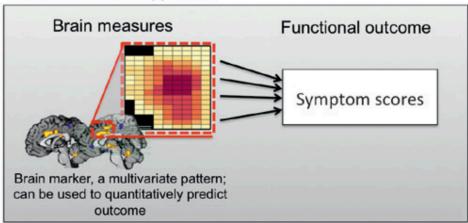
How to make precise predictions?

Traditional brain mapping approach



Wager (2015) Woo & Wager (2015) Pain Wager & Woo (2015) Science Trans Med

Brain-as-marker approach



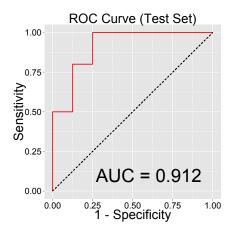
Ahn et al (2014) Curr Biol Finn et al (2015) Nature Neuro Gabrieli et al (2015) Neuron Norman et al (2006) TiCS Pereira et al (2009) Neuroimage Poldrack (2008) Curr Opin Neurobiol Wager et al (2013) NEJM Whelan et al (2014) Nature

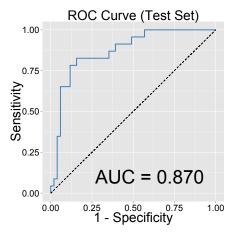
Example

Neuropsychosocial profiles of current and future adolescent alcohol misusers

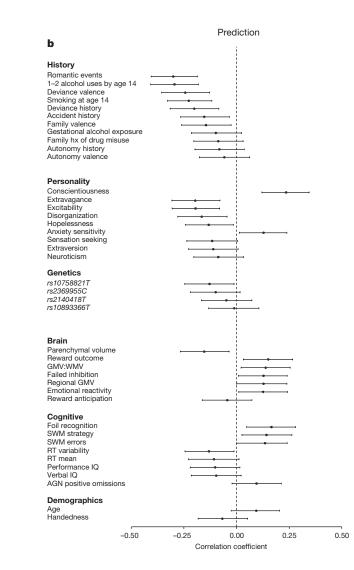
Robert Whelan^{1,2}, Richard Watts³, Catherine A. Orr⁴, Robert R. Althoff^{5,6}, Eric Artiges^{7,8}, Tobias Banaschewski⁹, Gareth J. Barker¹⁰, Arun L. W. Bokde¹¹, Christian Büchel^{12,13}, Fabiana M. Carvalho¹⁰, Patricia J. Conrod^{10,14}, Herta Flor⁹, Mira Fauth-Bühler^{9,15}, Vincent Frouin¹⁶, Juergen Gallinat^{12,17}, Gabriela Gan¹⁸, Penny Gowland¹⁹, Andreas Heinz¹⁷, Bernd Ittermann²⁰, Claire Lawrence²¹, Karl Mann⁹, Jean-Luc Martinot^{7,22}, Frauke Nees⁹, Nick Ortiz^{1,23}, Marie-Laure Paillère-Martinot^{17,22}, Tomas Paus^{24,25}, Zdenka Pausova²⁶, Marcella Rietschel⁹, Trevor W. Robbins²⁷, Michael N. Smolka¹⁸, Andreas Ströhle¹⁷, Gunter Schumann^{10,28}, Hugh Garavan^{1,6,11} & the IMAGEN Consortium[†]

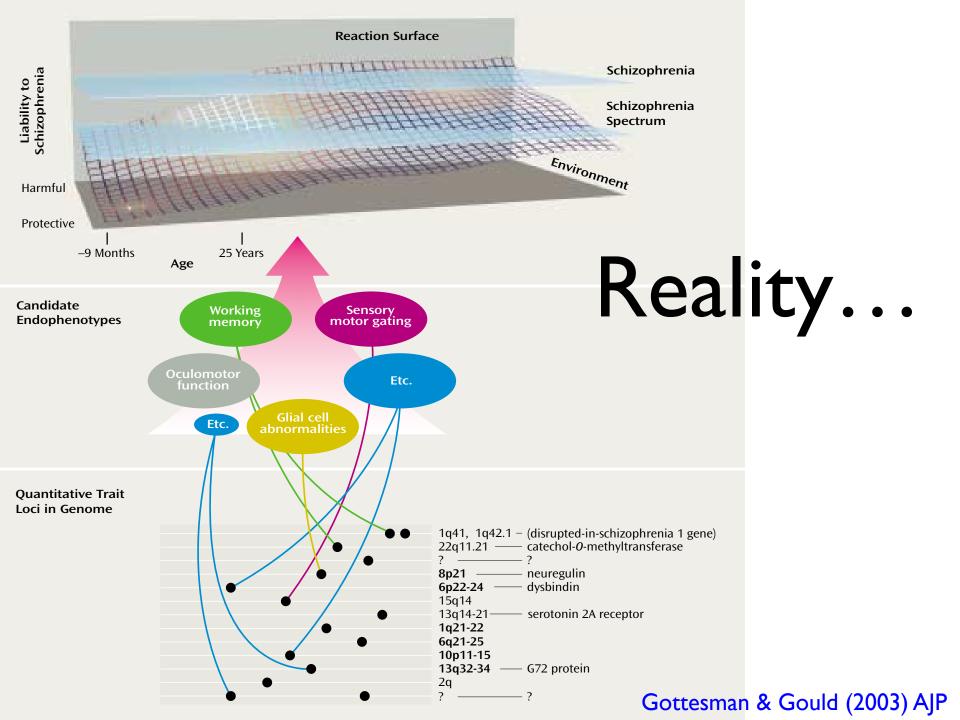
Whelan et al (2014) Nature





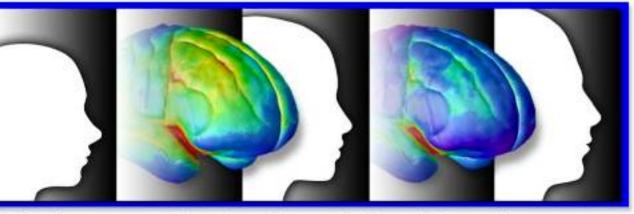
Ahn et al (2016) Frontiers Psychiatry Ahn & Vassileva (2016) DAD





Longitudinal study





Adolescent Brain Cognitive Development

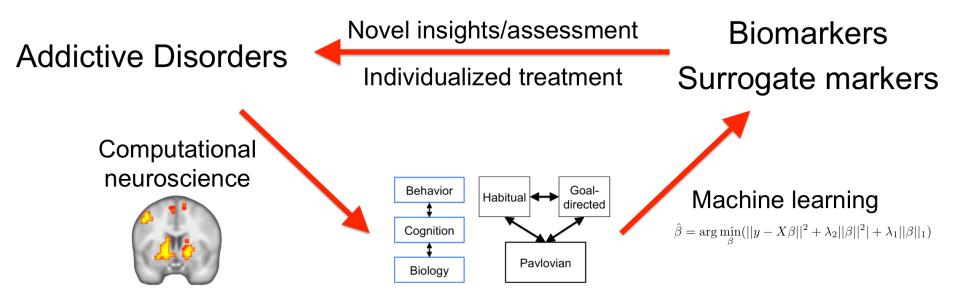
- 10,000 adolescents over 10 years
- Data sharing

Conclusions

- Cost-effective/surrogate markers
- Data/code sharing
- Education
- Collaboration across multiple disciplines
- A long way to go...

Directions

Computational Clinical Science (CCS) Laboratory



Thank you!