#### What kinds of algorithms would it take for a neuroscientist to understand a microprocessor? with Eric Jonas

BioRXiv: Could a neuroscientist understand a microprocessor?

#### Error messages are useful





Epistemological problem			$\times$
Your model does not describe reality.			
ignore	blame student	publish (nature)	

Reverse engineer a big biological distributed algorithm

MOS 6502





Courtesy http://visual6502.org

# How it actually works



#### Multi scale

1-bit Adder



AND gate Vdd Vdd в**--d** A-d Out Α-B-

Vss

AND gate (silicon)



logic gate primitives





#### 3 Behaviors



a. Donkey Kong (DK)

b. Space Invaders (SI)

c. Pitfall (PF)

#### Lesion studies

Lesions which impact single behavior



# How to make it work

- Problem: Complex game instead of targeted instructions
- Same as for brain
- But could work if one activated/inactivated
- And optimized stimulation so that effects are sparse

# "Spike data"



#### Tuning curves



luminance

### How to make it work

- Problem: not having understanding of "instructions"
- Same as for brain
- Run lots of programs. Relate instructions to activities.

#### Strong global correlations



#### LFPs and power law spectra







## Granger causality



a. Donkey Kong

b. Space Invaders

c. Pitfall

#### How to make these work?

• No idea!

#### Whole chip



time

# Nonnegative matrix factorization finds something



0.15

0.20

0.25

0.10

time (ms)

0.00

0.05

#### How to make these work?

- Need far more different states to be meaningful
- Far more data
- Nonlinear dimensionality reduction

#### Souped up Stochastic block model finds some network structure



Transistor distance

# How to make it work

- Problem: The network is far more complicated
- Same for the brain
- Solutions hierarchical structure inference
  - MCMC is too slow, clustering too unspecific, needs something in between
  - Big systems



Kasthuri and Lichtman



#### with Kasthuri, Xiao, Jacobsen

#### cubic mm



#### Conclusion

- We know little about how the brain works
- Data by itself won't solve the problem
- Need to ask the fundamental questions
- Countless big computational problems