Universal laws and architectures: Layering, learning, and decentralized control in neuroscience

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Outline: Laws and architectures

- Motivating case studies
 - Computers, networks
 - Cells
 - Physiology
 - Brains
- Bits of theory
 - Computation, Turing
 - Control
 - Info theory, stat mech

Emphasis

- Who, what, how, *why*
- Accident versus *necessity*

Turing on layering

The 'skin of an onion' analogy is also helpful. In considering the functions of the mind or the brain we find certain operations which we can explain in purely mechanical terms. This we say does not correspond to the real mind: it is a sort of skin which we must strip off if we are to find the real mind. But then in what remains we find a further skin to be stripped off, and so on. Proceeding in this way do we ever come to the 'real' mind, or do we eventually come to the skin which has nothing in it? In the latter case the whole mind is mechanical.

1950, Computing Machinery and Intelligence, Mind

"Universal laws and architectures?"

- Universal "conservation laws" (constraints)
- Universal architectures (constraints that deconstrain)

Fundamentals!

A rant

- Mention recent papers*
- Focus on broader context not in papers
- Lots of case studies for motivation

*try to get you to read them?



Other case studies (not today)

- Other complex tech nets, aerospace, etc
- Wildfire ecosystems
- Turbulence
- Stat mech foundations
- Synesthesia

Compute

Turing (1912-1954)

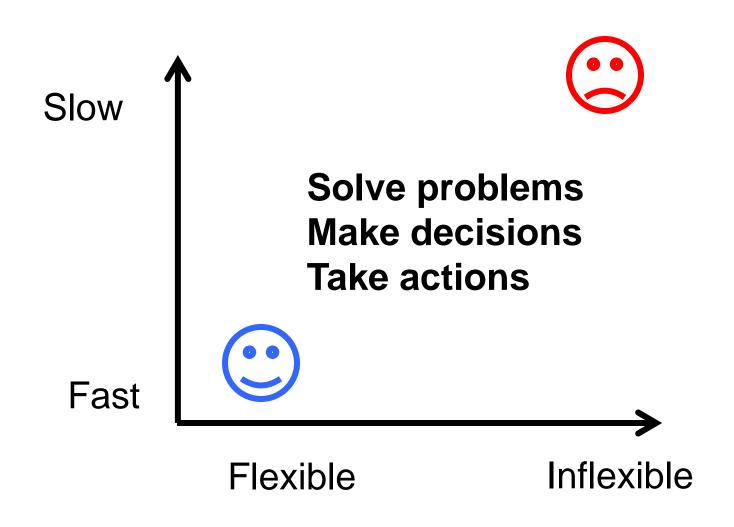
- Turing 100th birthday in 2012
- Turing
 - machine (math, CS)
 - test (AI, neuroscience)
 - pattern (biology)
- Arguably greatest*
 - all time math/engineering combination
 - WW2 hero
 - "invented" software

*Also world-class runner.

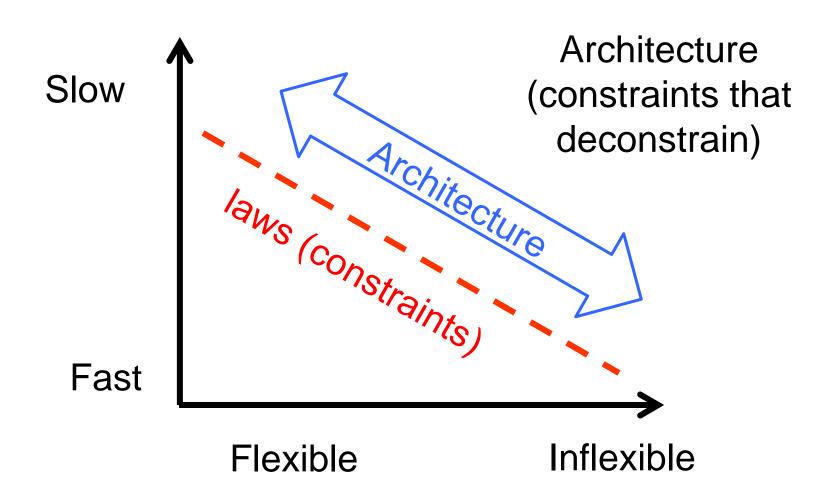
Key papers/results

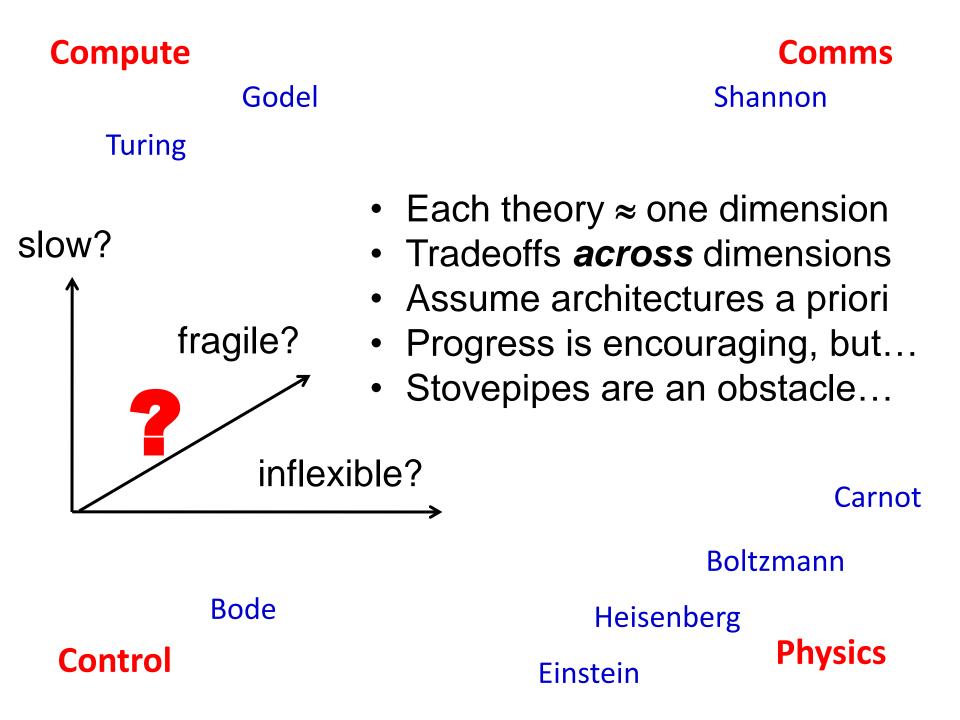
- Theory (1936): Turing machine (TM), computability, (un)decidability, universal machine (UTM)
- Practical design (early 1940s): code-breaking, including the design of code-breaking machines
- Practical design (late 1940s): general purpose digital computers and software, layered architecture
- Theory (1950): Turing test for machine intelligence
- Theory (1952): Reaction diffusion model of morphogenesis, plus practical use of digital computers to simulate biochemical reactions

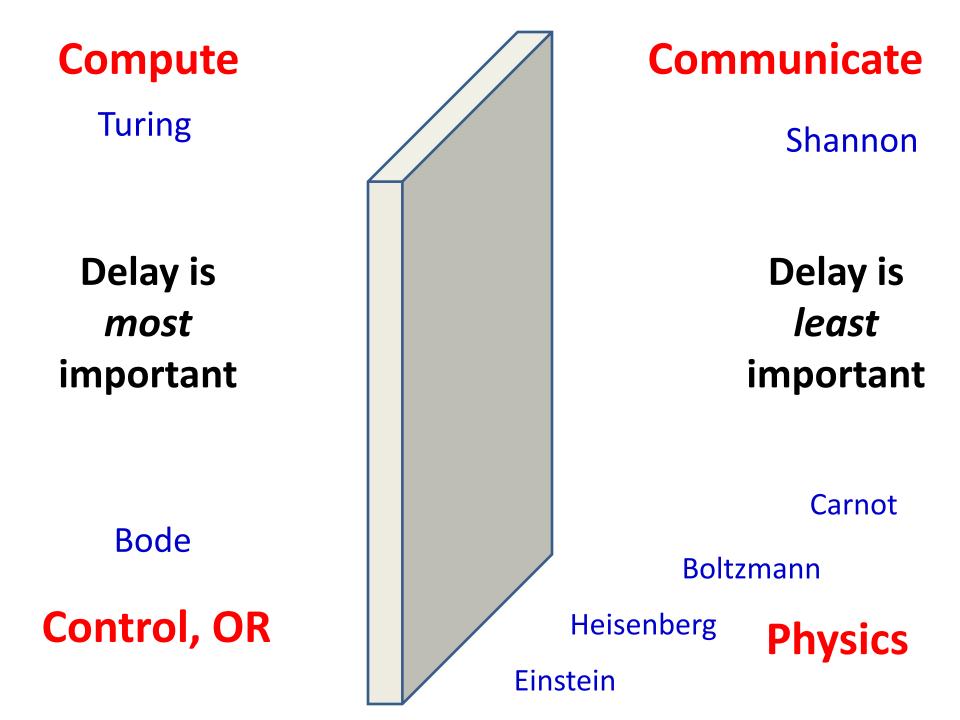
Fast and flexible

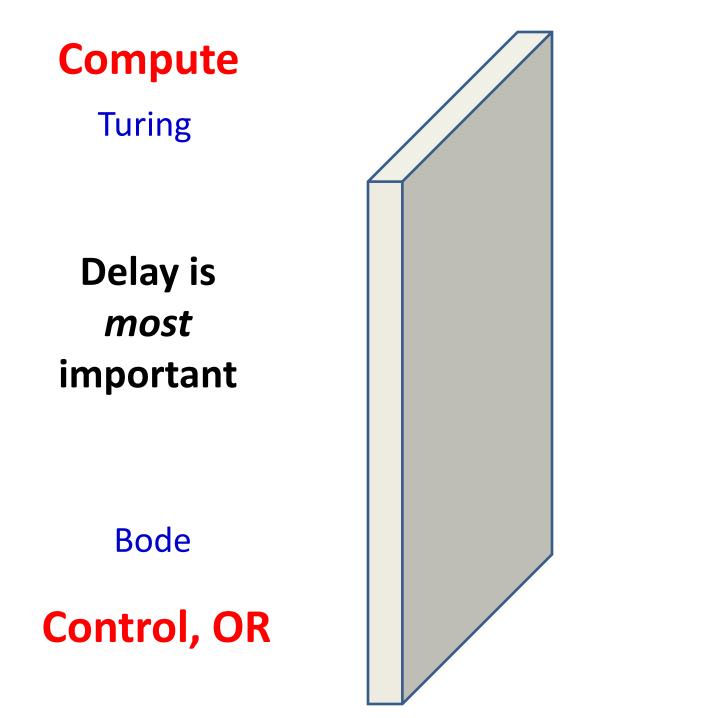


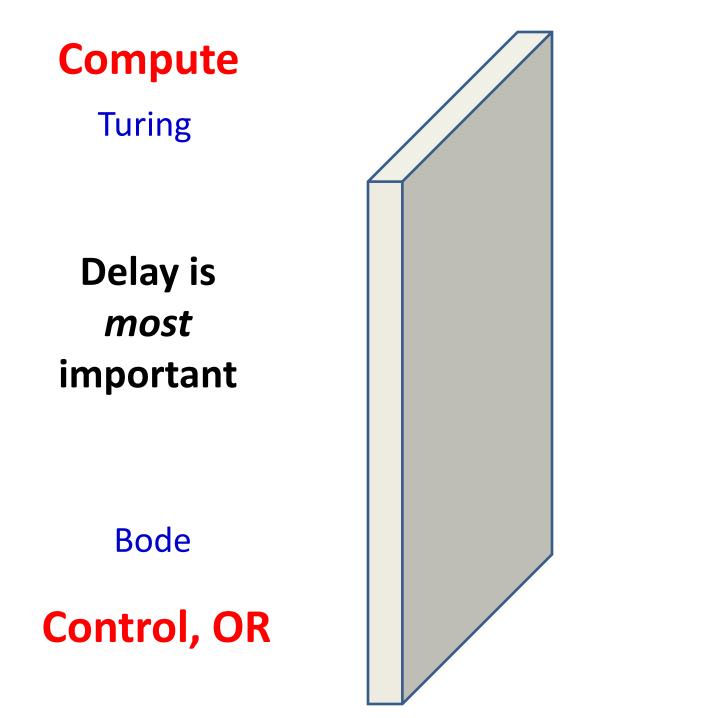
Laws and architectures











Turing as "new" starting point?

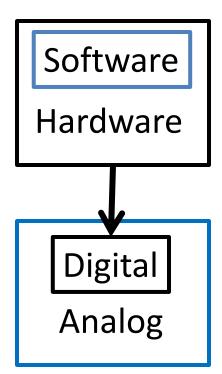


Turing

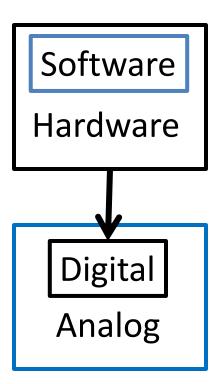




Control, OR



Turing as "new" starting point?



Essentials:

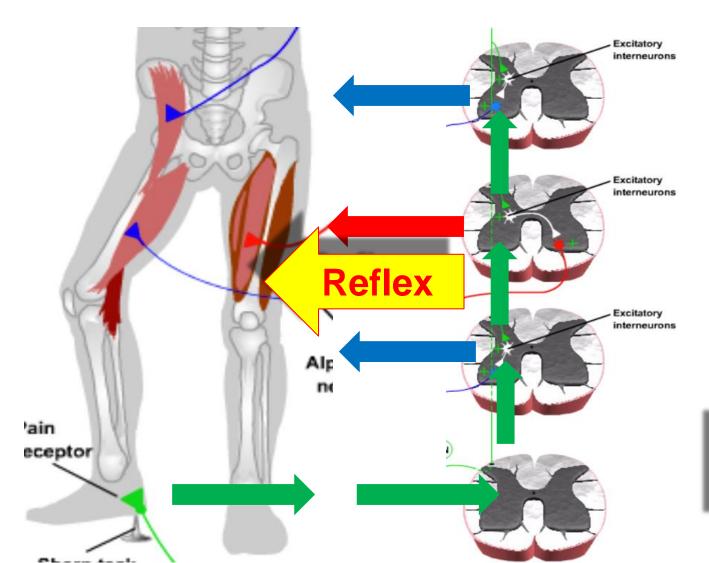
- 0. Model
- 1. Universal laws
- 2. Universal architecture
- 3. Practical implementation

Turing's 3 step research:

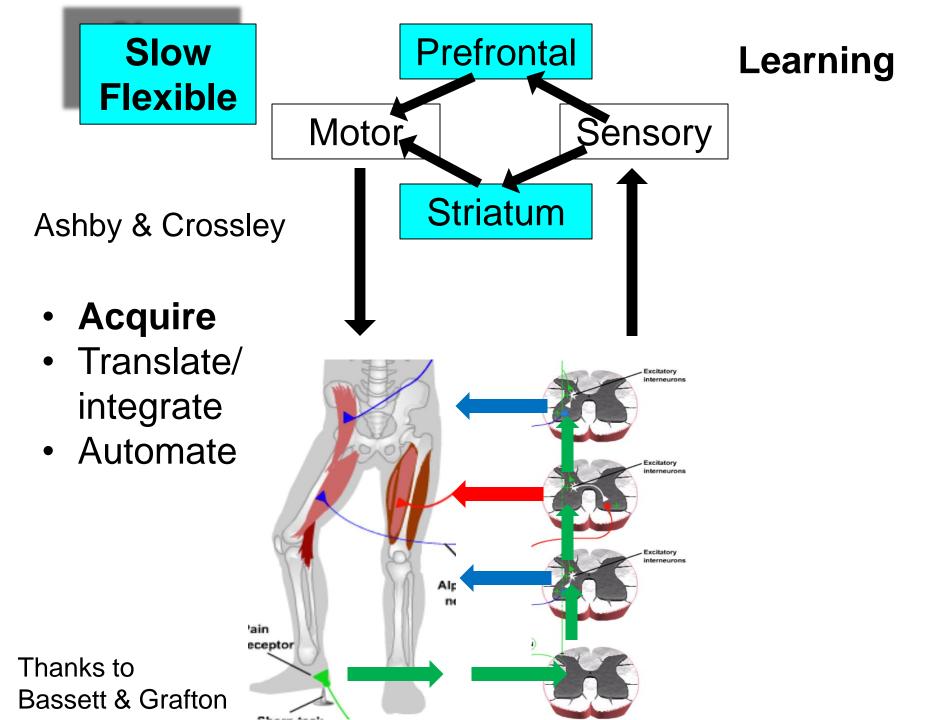
- 0. Virtual (TM) machines
- 1. hard limits, (un)decidability using standard model (TM)
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- 3. Practical implementation in digital electronics (biology?)

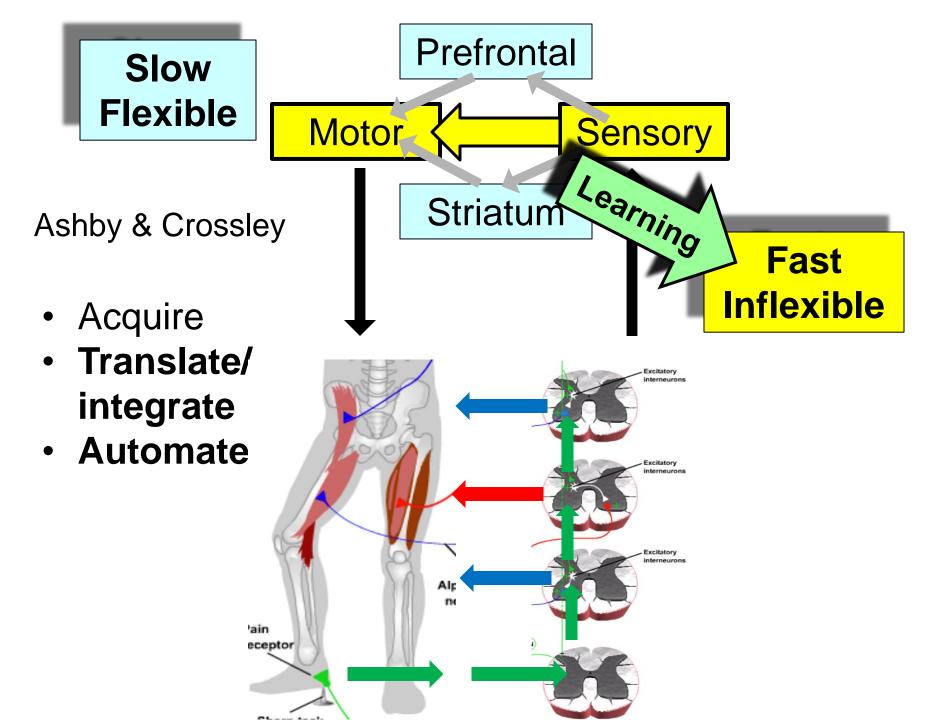
Who and what

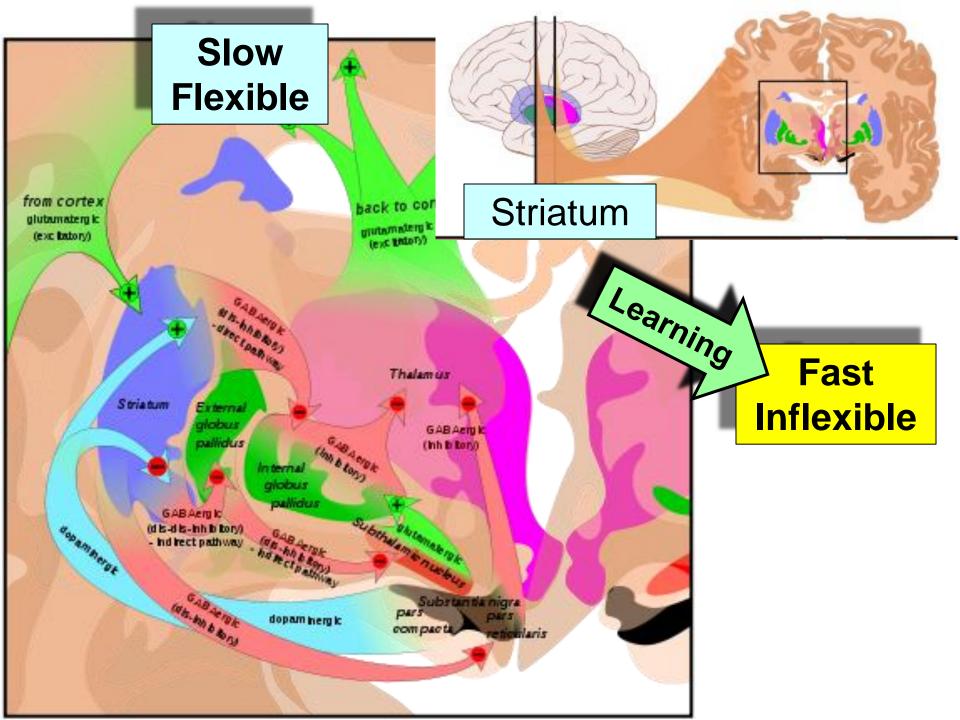
Neuro motivation

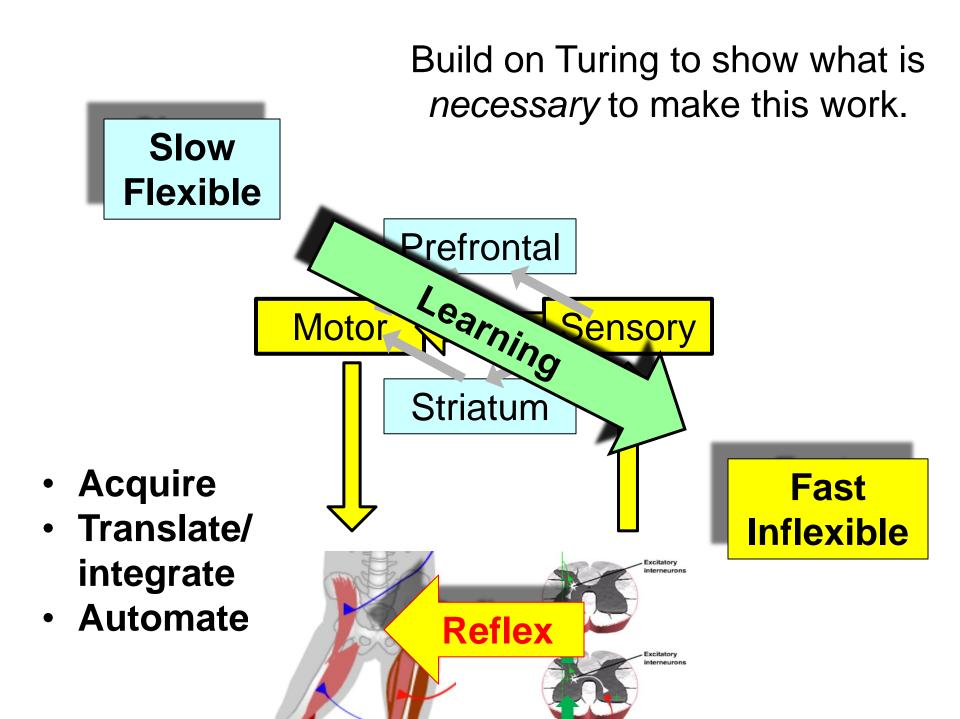


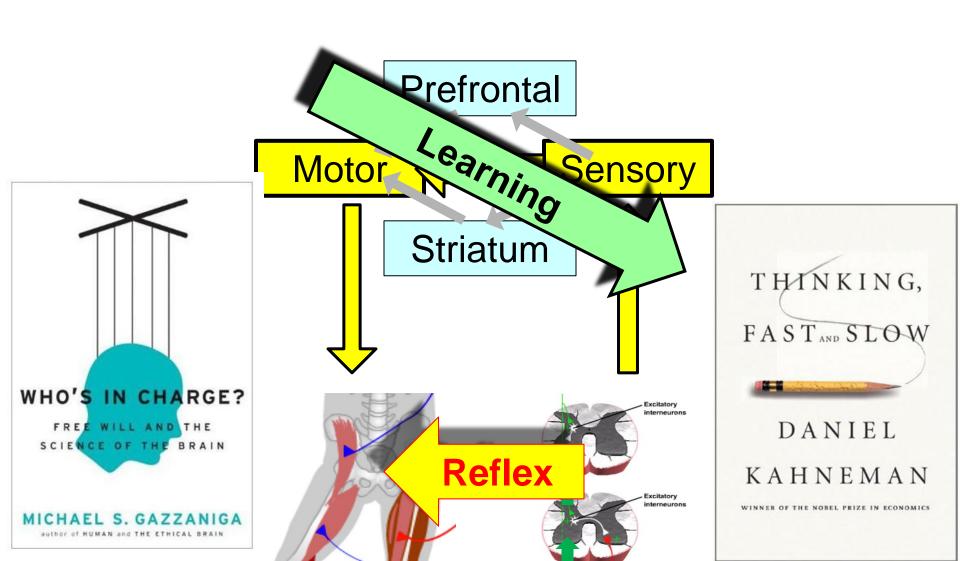


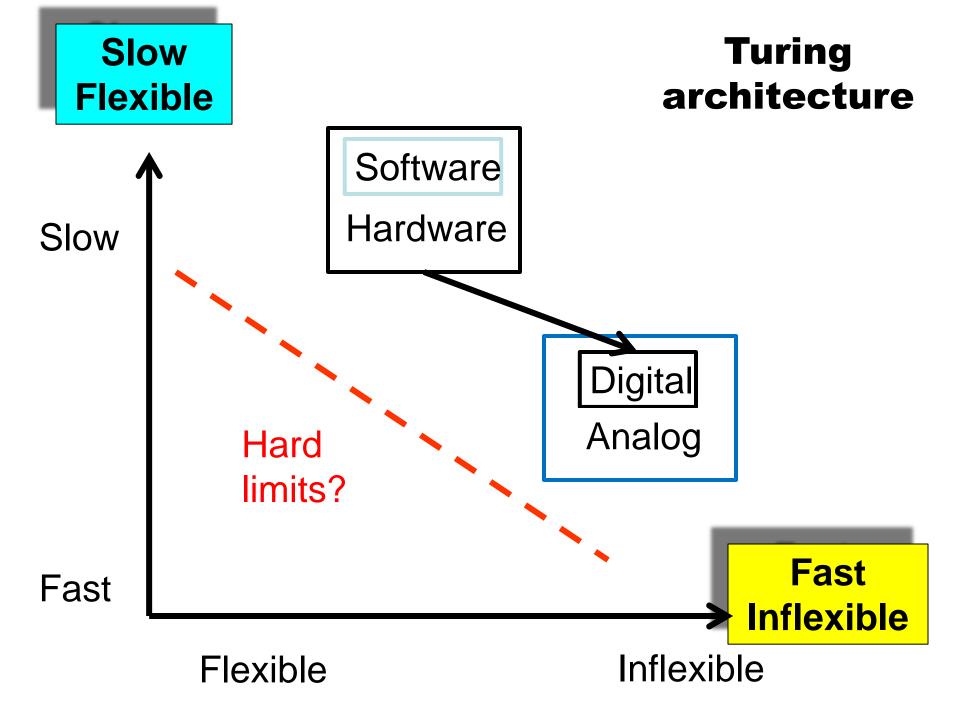




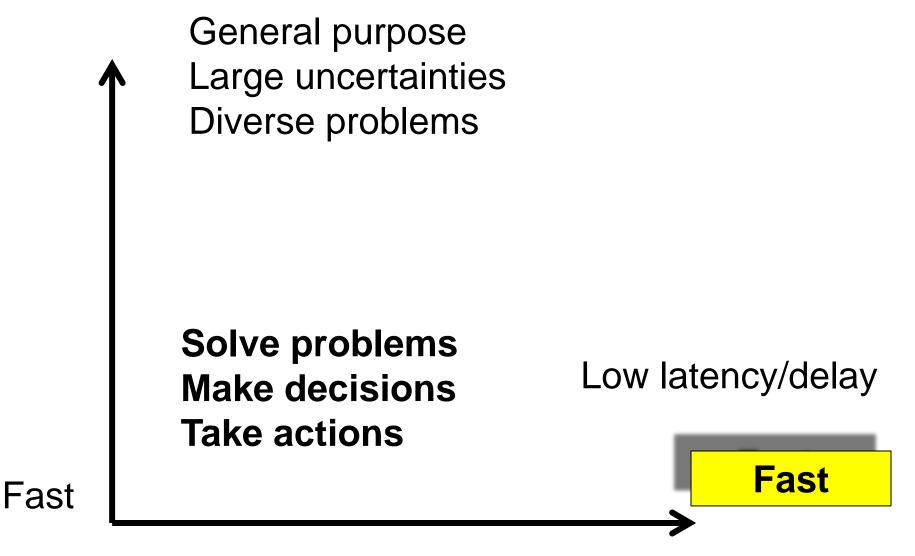




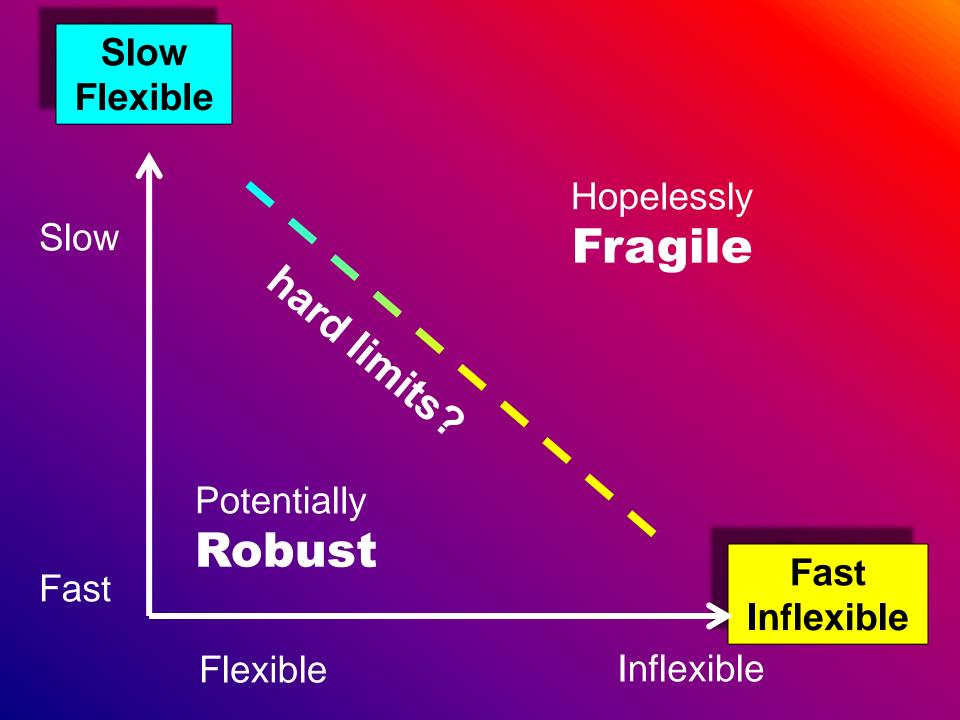








Flexible



Human complexity

Robust

- ③ Metabolism
- © Regeneration & repair
- ③ Healing wound /infect

Fragile

- Obesity, diabetes
- Cancer
- AutoImmune/Inflame

Start with physiology

Lots of triage

Benefits

Robust

- ③ Metabolism
- Constant Regeneration & repair
- ③ Healing wound /infect
 - Sefficient
 - ③ Mobility
 - Survive uncertain food supply
 - Recover from moderate trauma and infection

Mechanism?

Robust

- ③ Metabolism
- Constant Segmentation & Regeneration & Regeneration
- ③ Healing wound /infect
 - Set accumulation
 - Insulin resistance
 - Proliferation
 - ☺ Inflammation

Fragile

- Obesity, diabetes
- Cancer
- AutoImmune/Inflame
 - Sat accumulation
 - $\ensuremath{\textcircled{\otimes}}$ Insulin resistance
 - Proliferation
 - Inflammation

What's the difference?

Robust

- ③ Metabolism
- Regeneration & repair
- Healing wound /infect

Fragile

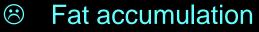
Obesity, diabetes

Cancer

- AutoImmune/Inflame
- Section 3 Fat accumulation
- Insulin resistance
- Proliferation
- Inflammation

Controlled Dynamic

Uncontrolled Chronic



- Insulin resistance
- Proliferation
- Inflammation

Controlled Dynamic

Low mean High variability

Death

Controlled Dynamic

Low mean High variability

- S Fat accumulation
- Insulin resistance
- Proliferation
- Inflammation

Uncontrolled Chronic

High mean Low variability

Restoring robustness?

Robust

- ③ Metabolism
- Regeneration & repair
- Healing wound /infect
 - Sat accumulation
 - ℬ Insulin resistance
 - Proliferation
 - Inflammation
 - Controlled Dynamic

Low mean High variability

Fragile

- Obesity, diabetes
- Cancer
- AutoImmune/Inflame
 - Section 8 Fat accumulation
 - Insulin resistance
 - Proliferation
 - Inflammation
 - Uncontrolled Chronic

High mean Low variability

Human complexity

Robust

Yet Fragile

- Metabolism
- Regeneration & repair
- Immune/inflammation
- ③ Microbe symbionts
- Seuro-endocrine
- Complex societies
- Advanced technologies
- Risk "management"

- Obesity, diabetes
- Cancer
- AutoImmune/Inflame
- Parasites, infection
- ⊗ Addiction, psychosis,...
- Epidemics, war,...
- ♦ Obfuscate, amplify,...

Accident or necessity?

Robust

Metabolism

Fragile Obesi<u>ty, diabetes</u>

- Segenerati
- Healing wo
- Set accumulation

 \odot

- Insulin resistance
- Proliferation
- Inflammation

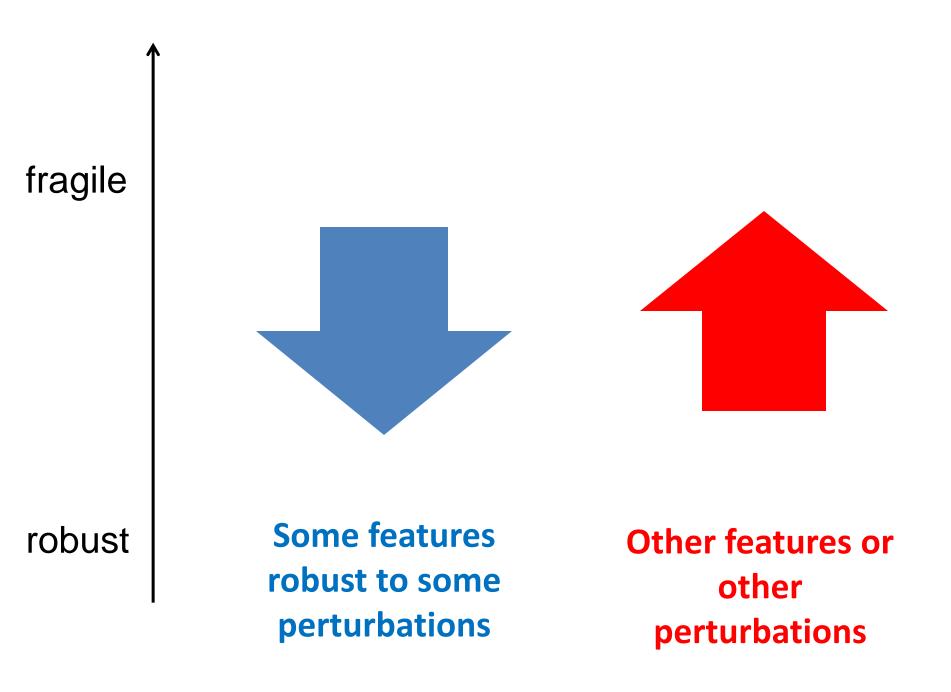
une/Inflame

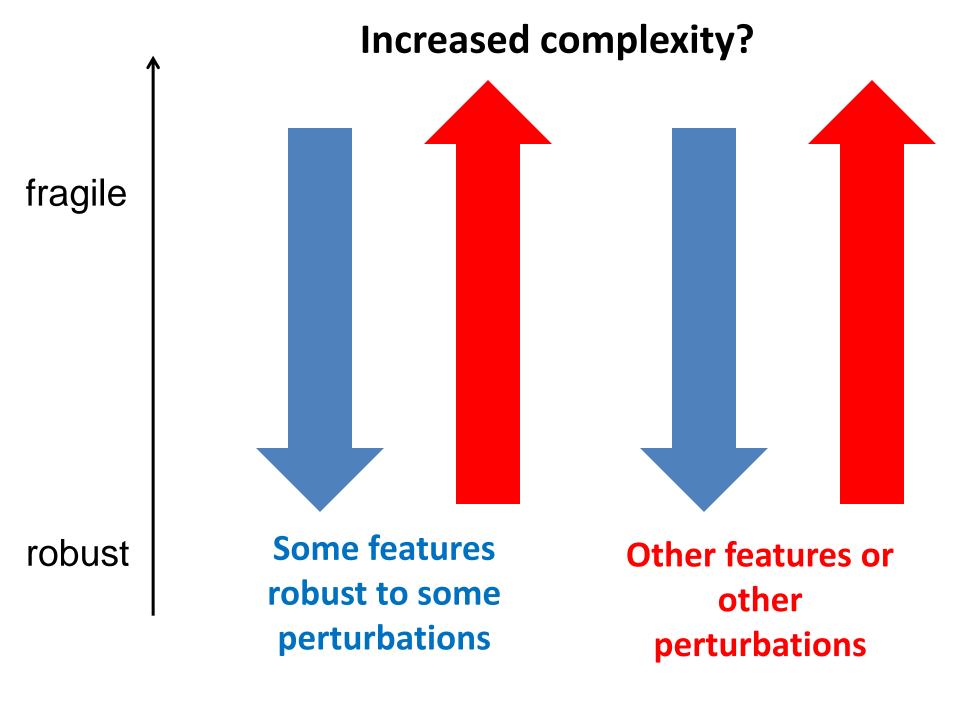
- Fragility ← Hijacking, side effects, unintended...
- Of mechanisms evolved for robustness
- Complexity ← control, robust/fragile tradeoffs
- Math: robust/fragile constraints ("conservation laws")

Both

Accident or necessity?



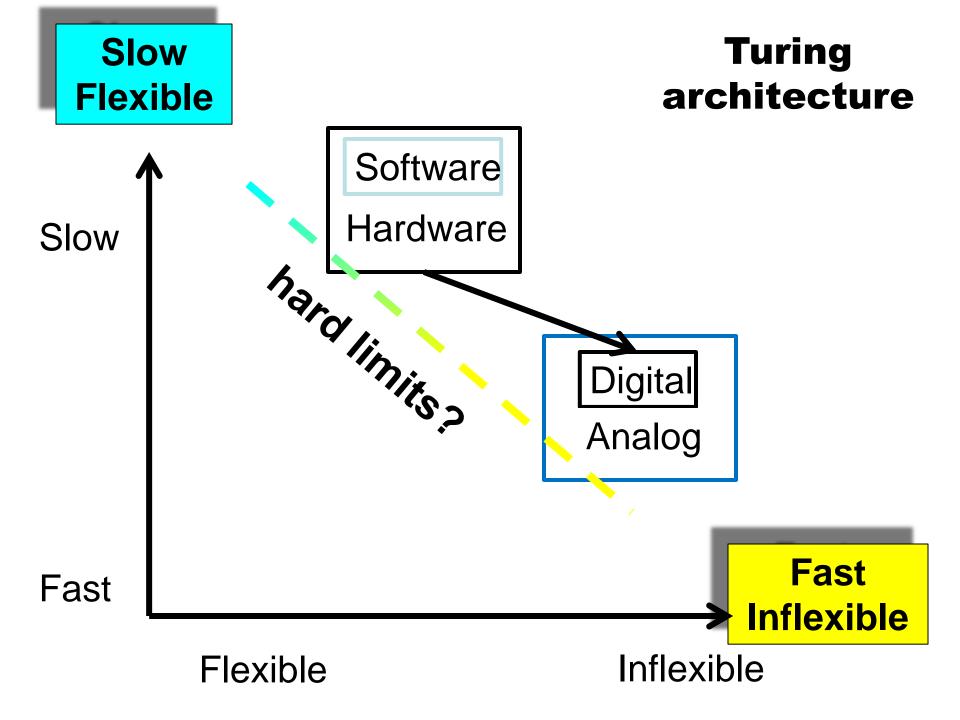


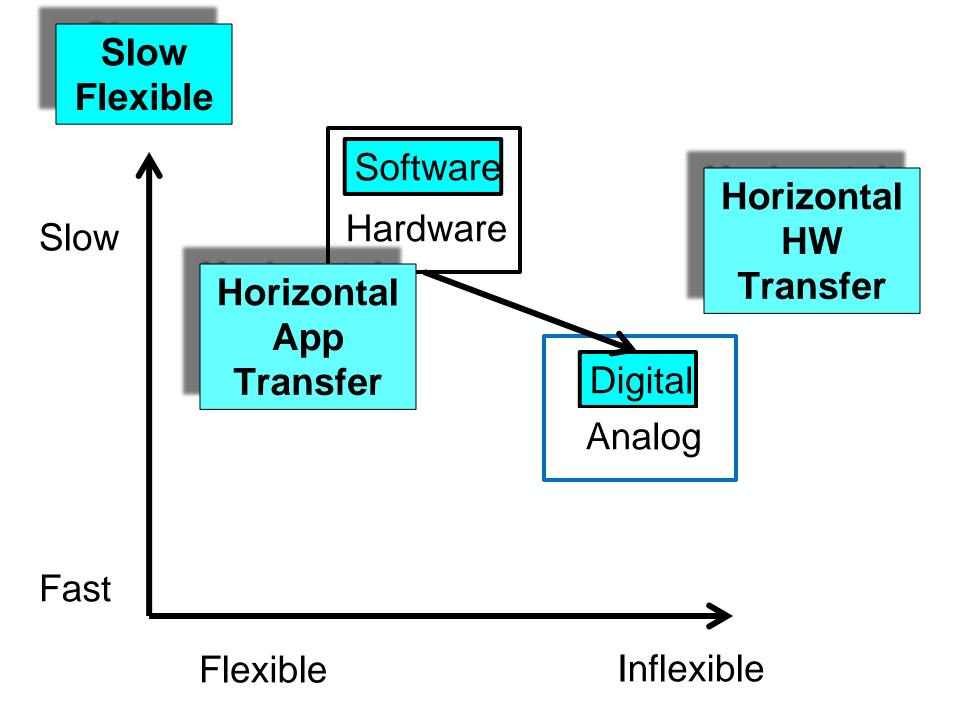


Robust Modular Simple Plastic Evolvable Fragile Distributed Complex Frozen Frozen

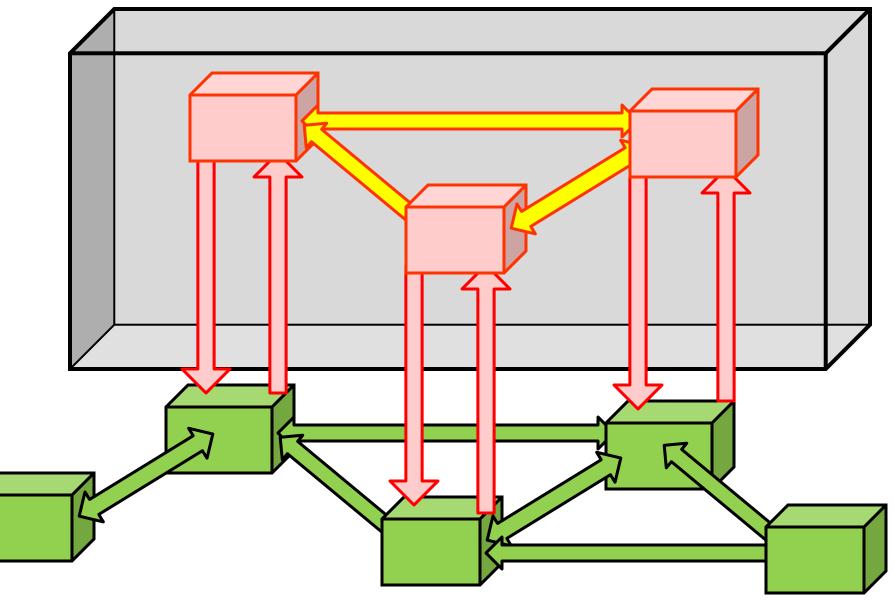
tradeoffs

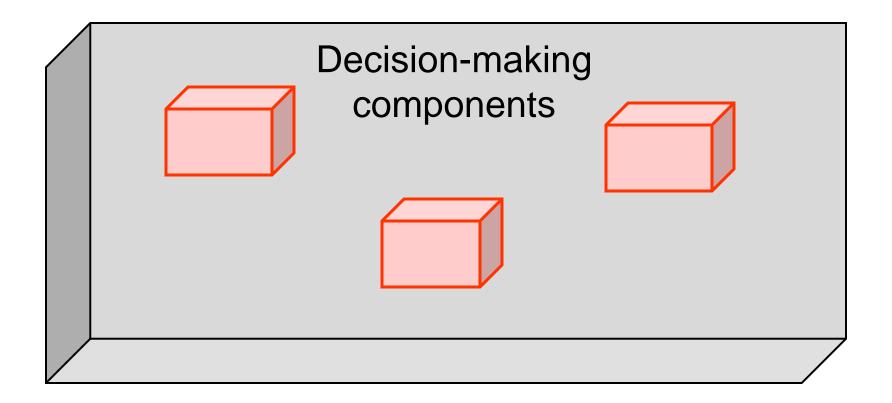
and



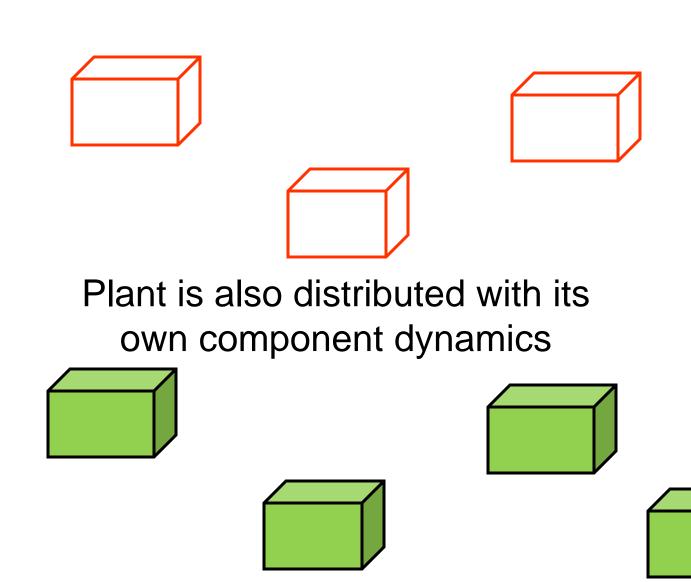


Cyber-physical: decentralized control with internal delays.

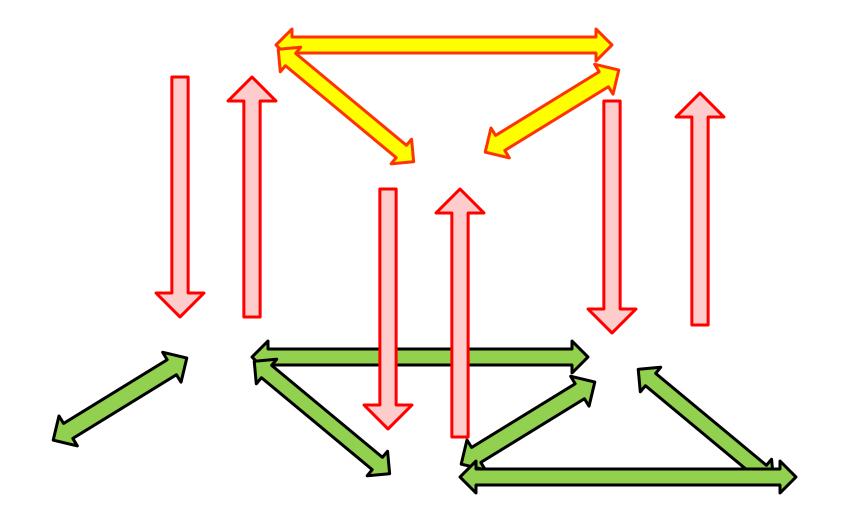




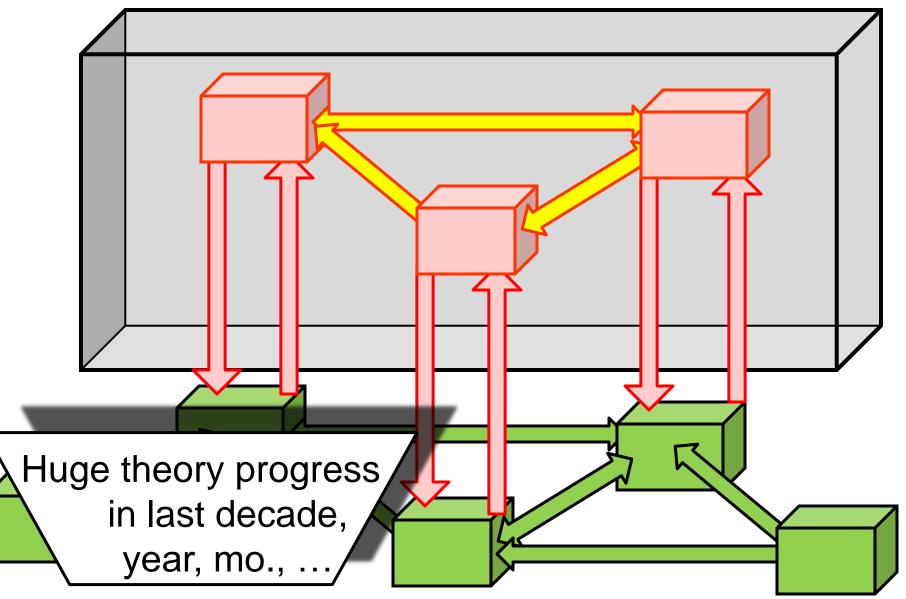
Decentralized, but initially assume computation is fast and memory is abundant.



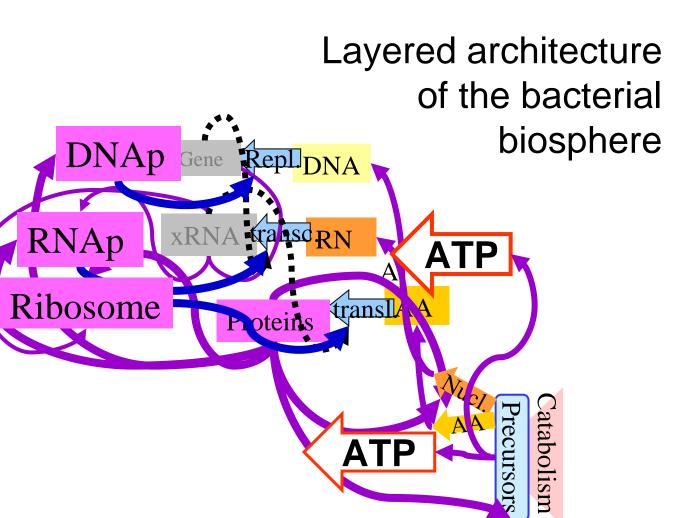
Internal delays between components, and their sensor and actuators, and also externally between plant components



Going beyond black box: control is decentralized with internal delays.



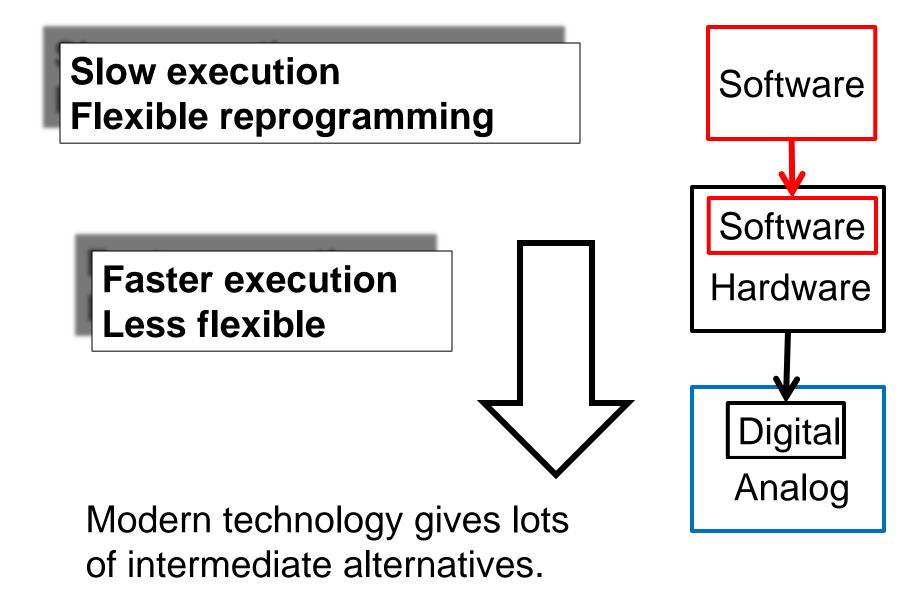
The best case study so far

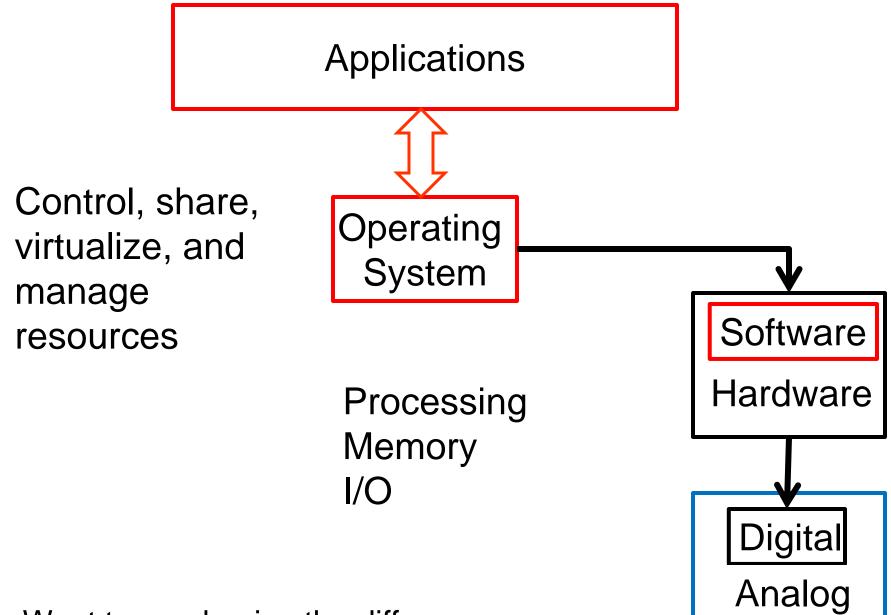


Not done here in detail, see slides elsewhere

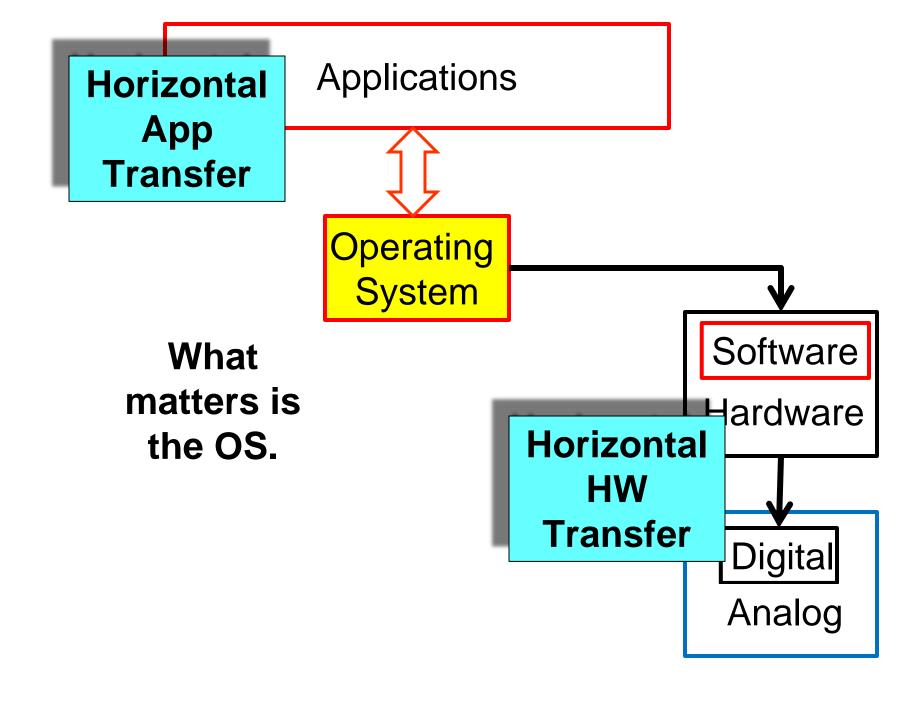
How?

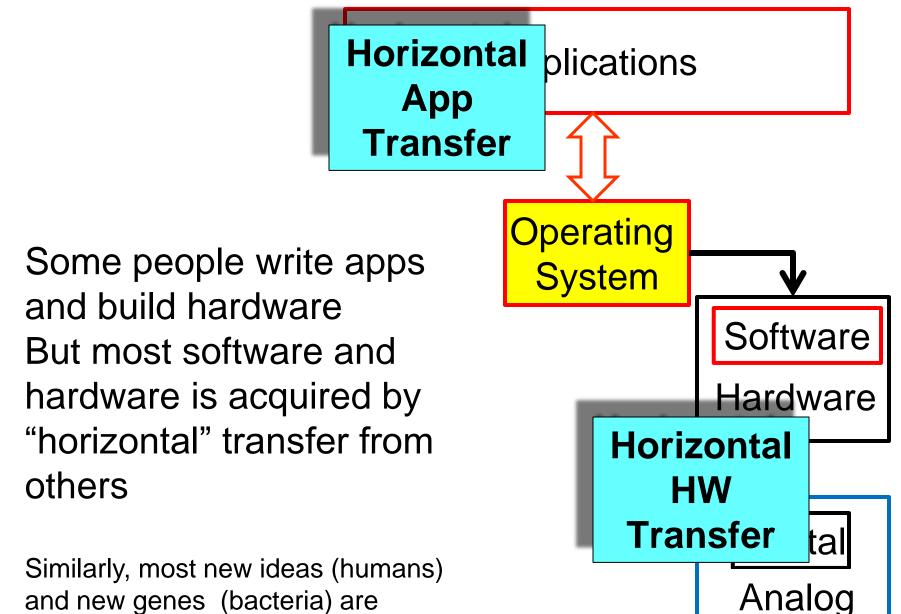
Universal architectures





Want to emphasize the differences between these two types of layering.

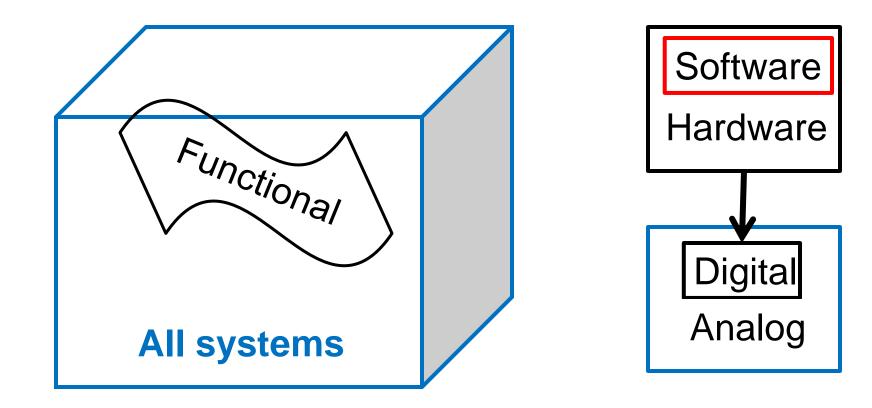




and new genes (bacteria) are acquired horizontally

"solution sets" (a la Marder, Prinze, etc)

large, thin, nonconvex



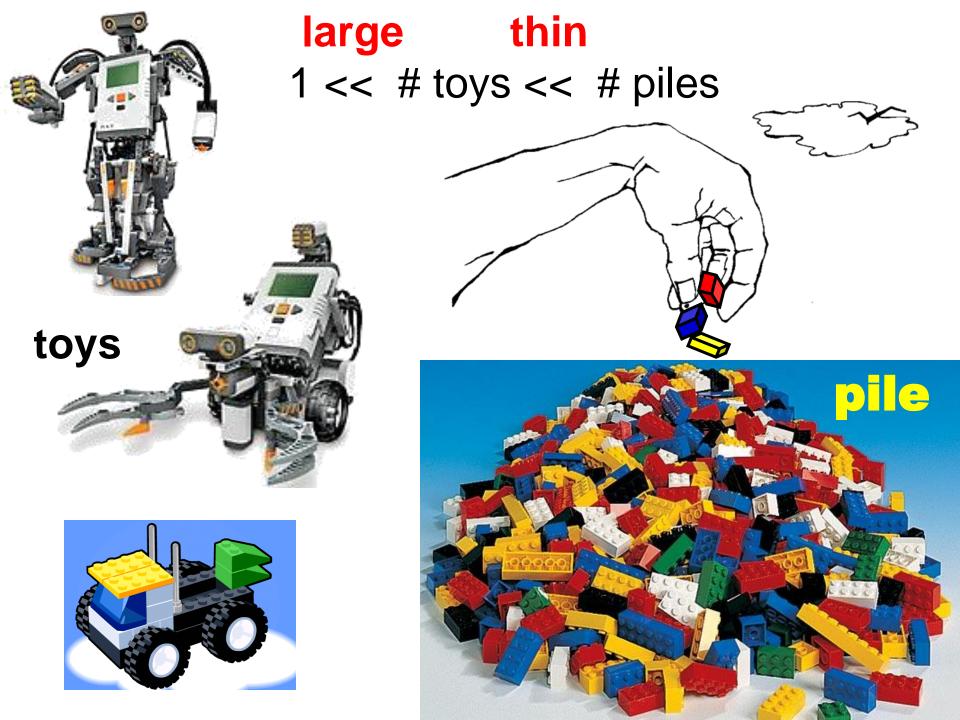
Letters and words

- 9 letters: adeginorz
- 9!= 362,880 sequences of 9 letters
- Only "organized" is a word

1 << (# words) << (# non-words) large thin

Computer programs

- Almost any computer language
- Large # of working programs
- Much larger # of non-working programs
- "Nonconvex" = simple mashups of working programs don't work
- 1 << (# programs) << (# non-programs) large thin

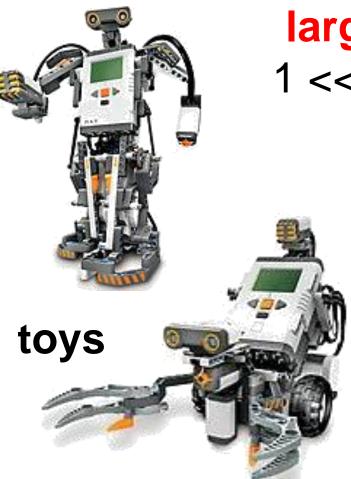




edge of chaos self-organized criticality scale-free ???

statistical physics random ensembles minimally tuned phase transitions bifurcations



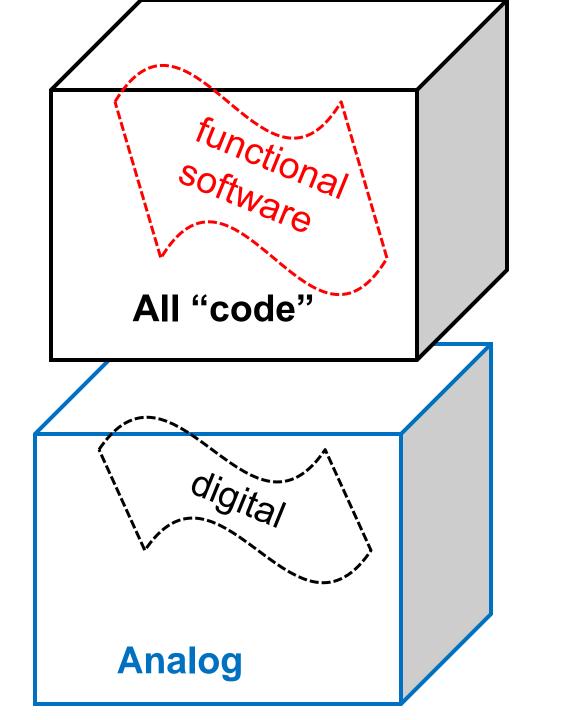




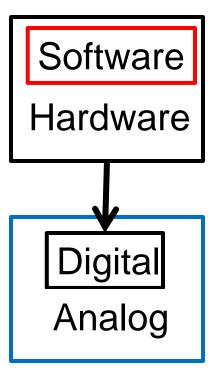
largethin1 << # toys << # piles</td>

nonconvex





large, thin, nonconvex



This paper aims to bridge progress in **neuroscience** involving sophisticated quantitative analysis of behavior, including the use of **robust control**, with other relevant conceptual and theoretical frameworks from **systems engineering**, **systems biology**, and **mathematics**.

Architecture, constraints, and behavior

Very accessible No math

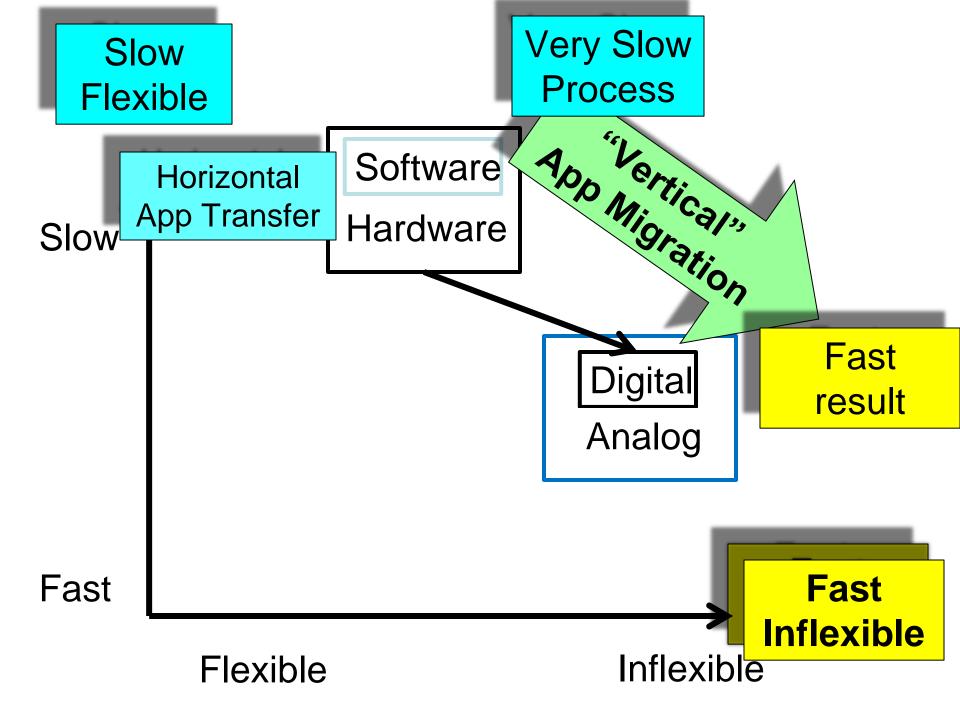
John C. Doyle^{a,1} and Marie Csete^{b,1}

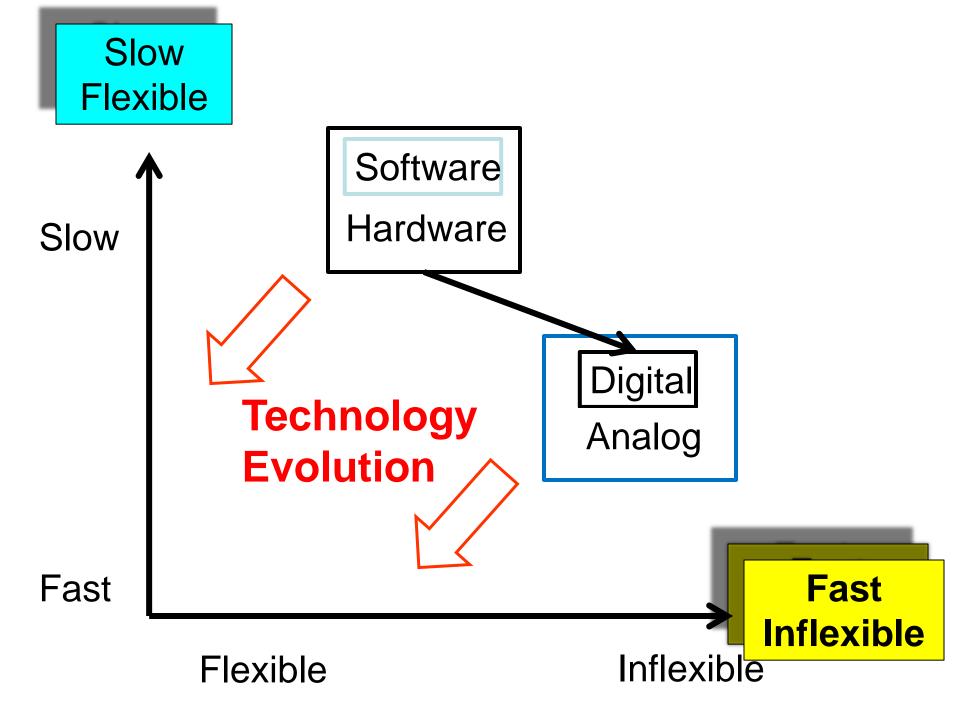
^aControl and Dynamical Systems, California Institute of Technology, Pasadena, CA 91125; and ^bDepartment of Anesthesiology, University of California, San Diego, CA 92103

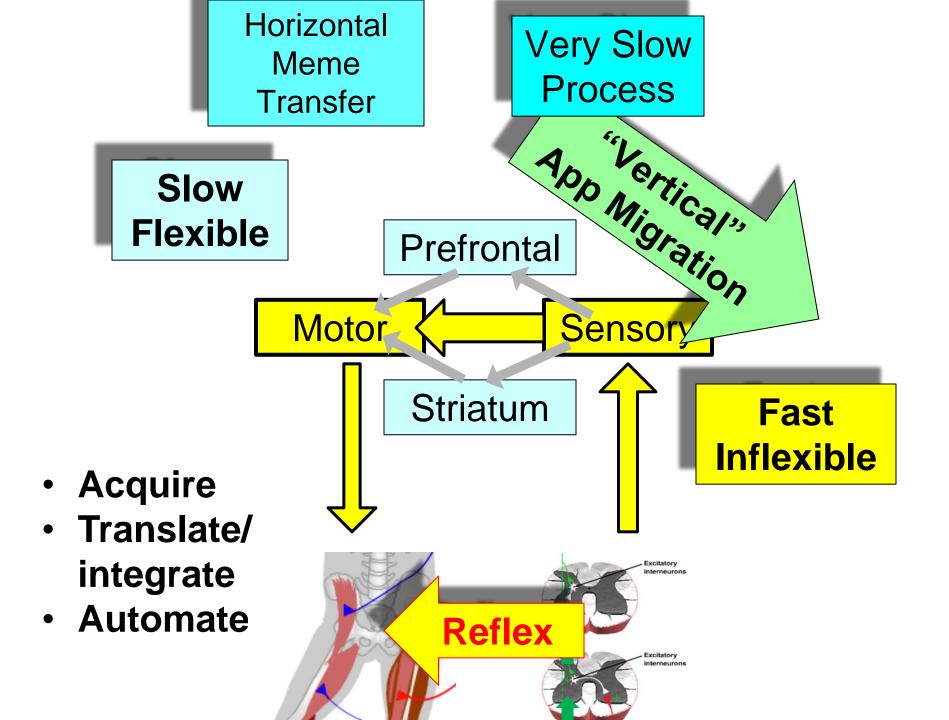
Edited by Donald W. Pfaff, The Rockefeller University, New York, NY, and approved June 10, 2011 (received for review March 3, 2011)

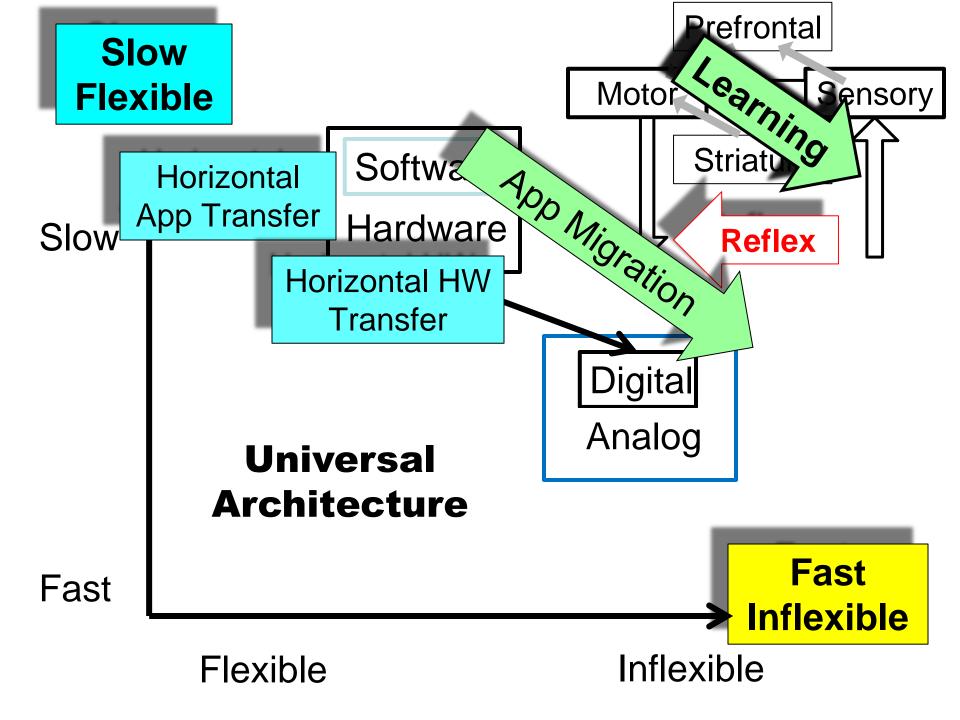
This paper aims to bridge progress in neuroscience involving sophisticated quantitative analysis of behavior, including the use of robust control, with other relevant conceptual and theoretical frameworks from systems engineering, systems biology, and mathematics. Familiar and accessible case studies are used to illustrate concepts of robustness, organization, and architecture (modularity and protocols) that are central to understanding complex networks. These essential organizational features are hidden during normal function of a system but are fundamental for understanding the nature, design, and function of complex biologic and technologic systems. evolved for sensorimotor control and retain much of that evolved architecture, then the apparent distinctions between perceptual, cognitive, and motor processes may be another form of illusion (9), reinforcing the claim that robust control and adaptive feedback (7, 11) rather than more conventional serial signal processing might be more useful in interpreting neurophysiology data (9). This view also seems broadly consistent with the arguments from grounded cognition that modal simulations, bodily states, and situated action underlie not only motor control but cognition in general (12), including language (13). Furthermore, the myriad constraints involved in the evolution of circuit

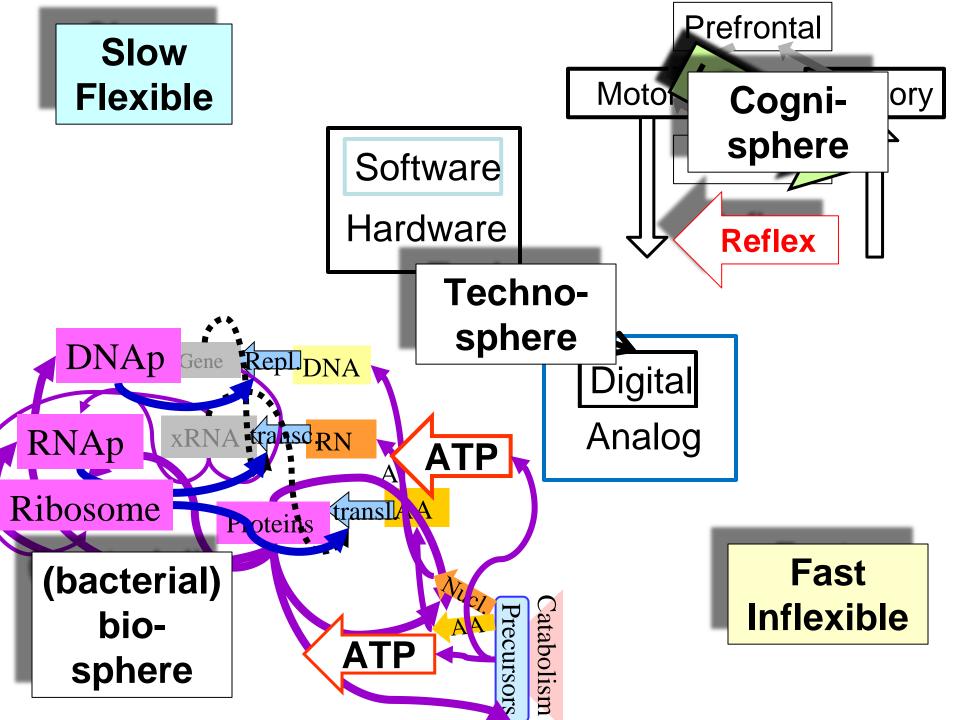
Doyle, Csete, Proc Nat Acad Sci USA, JULY 25 2011

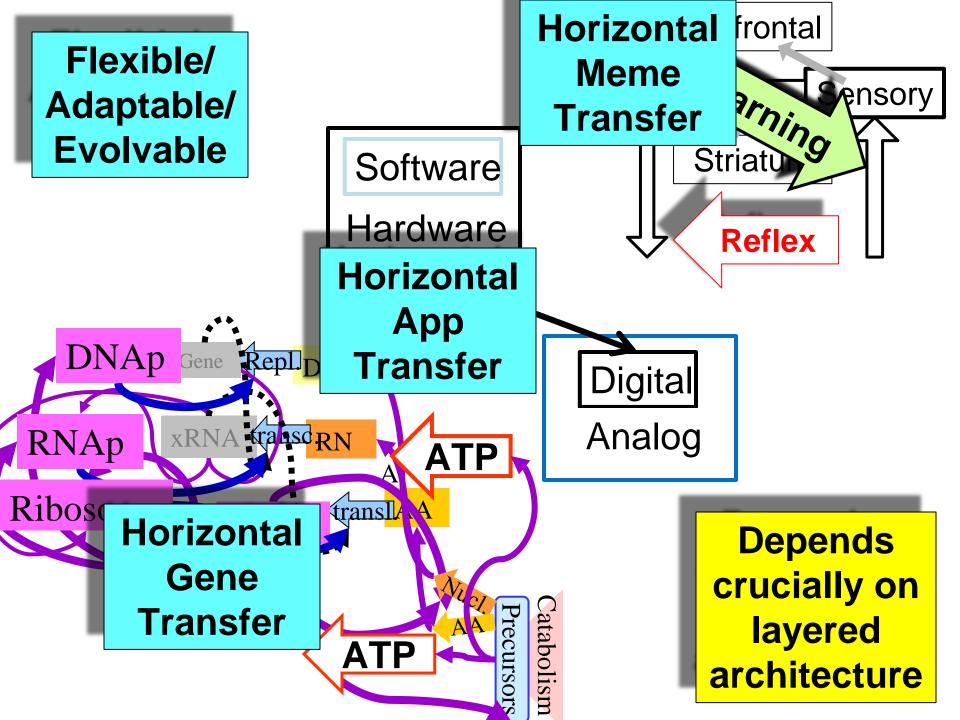


















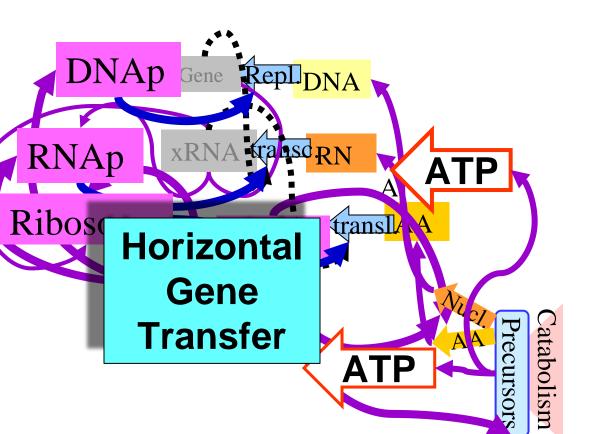
Most

- software and hardware
- new ideas (humans)
- new genes (bacteria)

is acquired by "horizontal" transfer, though sometimes it is evolved locally

Sequence ~100 E Coli (not chosen randomly)

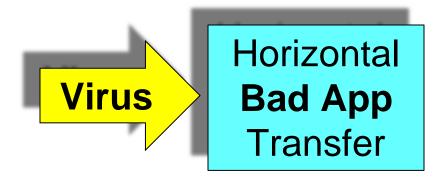
- ~ 4K genes per cell
- ~20K different genes in total
- ~ 1K universally shared genes



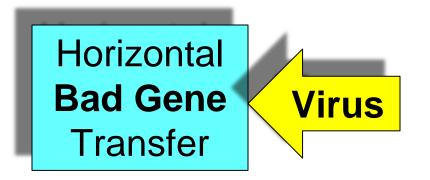
See slides on bacterial biosphere

Exploiting layered architecture

Horizontal Bad Meme Transfer









Depends crucially on layered architecture

Build on Turing to show what is *necessary* to make this work.

Acquire

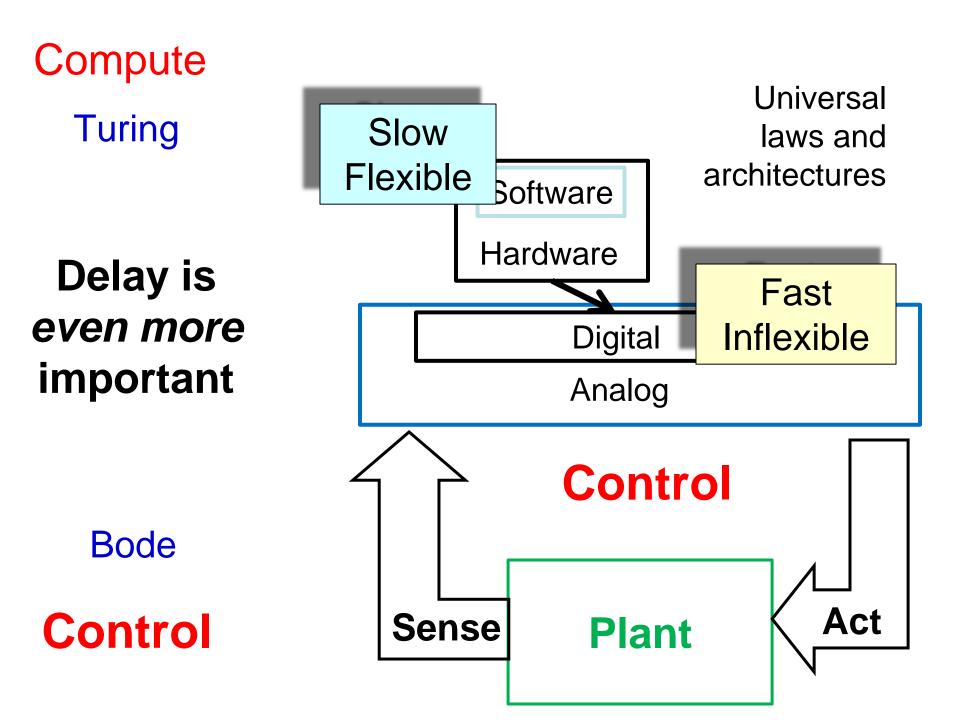
- Translate/
 integrate
- Automate

Horizontal Meme Transfer

Horizontal App Transfer

> Amazingly Flexible/ Adaptable

Horizontal Gene Transfer



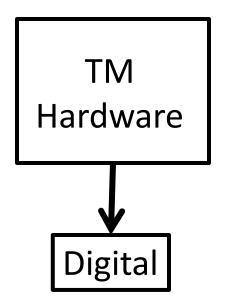


Why

Necessity

Essentials:

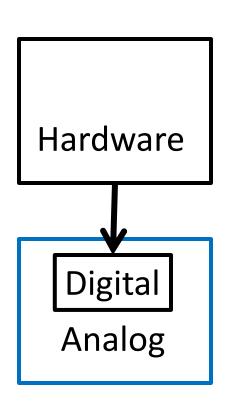
- 0. Model
- 1. Universal laws
- 2. Universal architecture
- 3. Practical implementation



Turing's 3 step research:

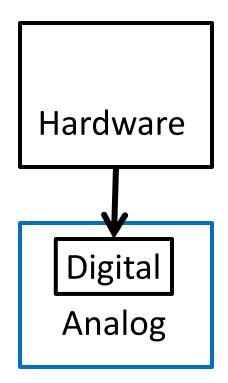
0. Virtual (TM) machines

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- 3. Practical implementation in digital electronics (biology?)



- ...being digital should be of greater interest than that of being electronic. That it is electronic is certainly important because these machines owe their high speed to this... But this is virtually all that there is to be said on that subject.
- That the machine is digital however has more subtle significance. ... One can therefore work to any desired degree of accuracy.

1947 Lecture to LMS

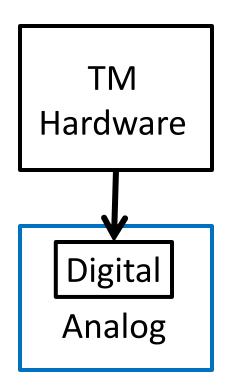


- ... digital ... of greater interest than that of being electronic ...
- ...any desired degree of accuracy...
- This accuracy is not obtained by more careful machining of parts, control of temperature variations, and such means, but by a slight increase in the amount of equipment in the machine.

1947 Lecture to LMS

Summarizing Turing:

- Digital more important than electronic...
- Robustness: accuracy and repeatability.
- Achieved more by internal hidden complexity than precise components or environments.



Turing Machine (TM)

- Digital
- Symbolic
- Logical
- Repeatable



• ... quite small errors in the initial conditions can have an overwhelming effect at a later time. The displacement of a single electron by a billionth of a centimetre at one moment might make the difference between a man being killed by an avalanche a year later, or escaping.

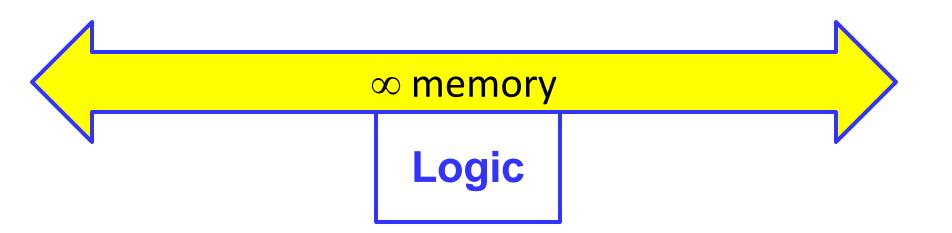
1950, Computing Machinery and Intelligence, *Mind*

• ... quite small errors in the initial conditions can have an overwhelming effect at a later time....

 It is an essential property of the mechanical systems which we have called 'discrete state machines' that this phenomenon does not occur.

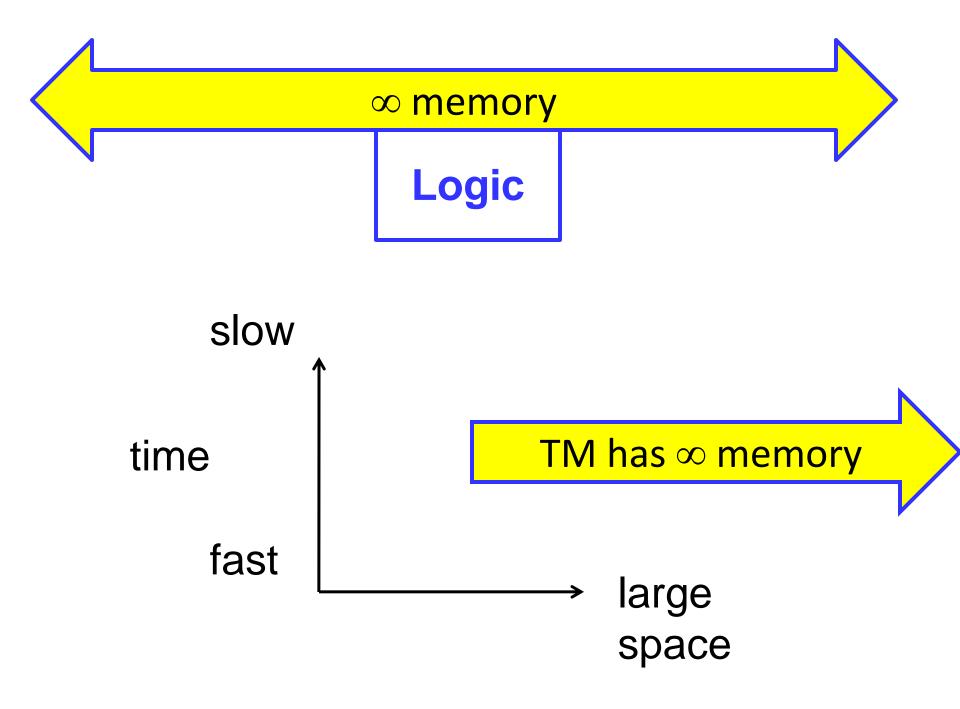
• Even when we consider the actual physical machines instead of the idealised machines, reasonably accurate knowledge of the state at one moment yields reasonably accurate knowledge any number of steps later.

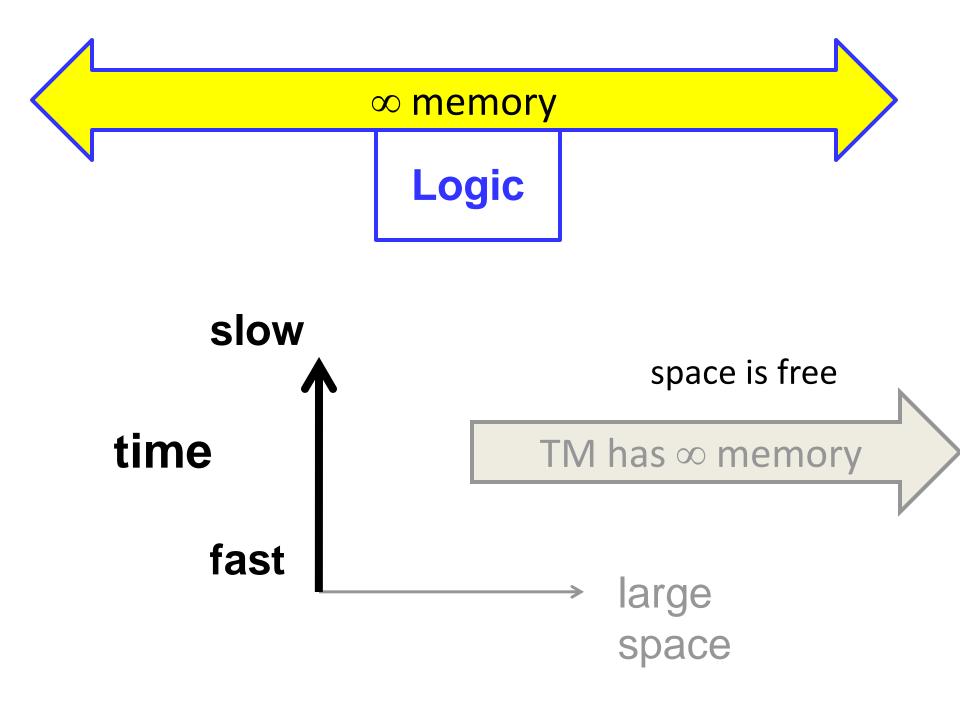
1950, Computing Machinery and Intelligence, Mind

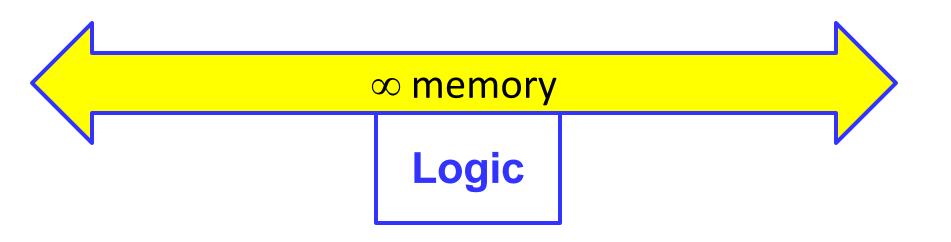


TM Hardware Turing's 3 step research:

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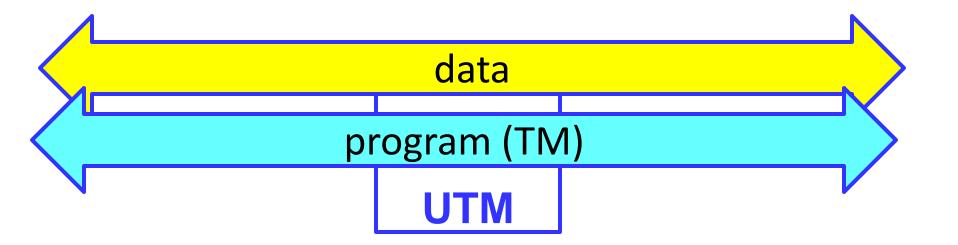




time?

Decidable problem = \exists algorithm that solves it

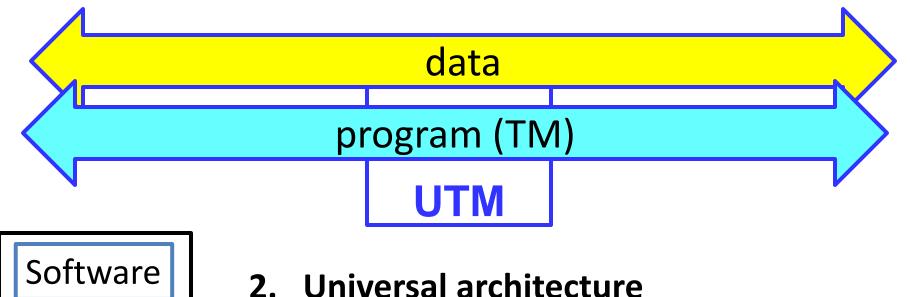
Most naively posed problems are undecidable.



Turing's 3 step research:

- 0. Virtual (TM) machines
- 1. hard limits, (un)decidability using standard model (TM)
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3. Practical implementation in digital electronics (biology?)

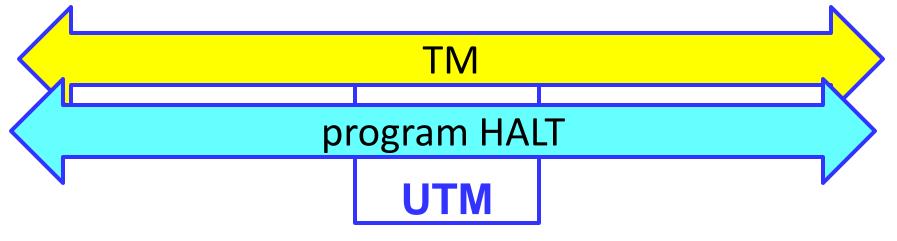


achieving hard limits (UTM)

- Software: A Turing machine (TM) can be data for another Turing machine
- A Universal Turing Machine can run any TM
- A UTM is a virtual machine.

Hardware

• There are lots of UTMs, differ only (but greatly) in speed and programmability (space assumed free)



The halting problem

- Given a TM (i.e. a computer program)
- Does it halt (or run forever)?
- Or do more or less anything in particular.
- Undecidable! There does not exist a special TM that can tell if any other TM halts.
- i.e. the program HALT does not exist. ⊗

Thm: TM H=HALT does not exist.

That is, there does not exist a program like this:

$$H(TM, input) \triangleq \begin{cases} 1 \text{ if } TM(input) \text{ halts} \\ 0 \text{ otherwise} \end{cases}$$

Proof is by contradiction. Sorry, don't know any alternative. And Turing is a god.

$$H(TM, input) \triangleq \begin{cases} 1 \text{ if } TM(input) \text{ halts} \\ 0 \text{ otherwise} \end{cases}$$

Thm: No such H exists.

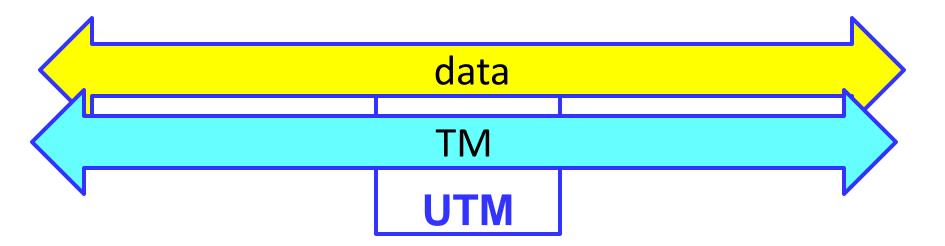
Proof: Suppose it does. Then define 2 more programs:

$$H'(TM, input) \triangleq \begin{cases} 1 \text{ if } H(TM, input) = 0\\ \text{loop forever otherwise} \end{cases}$$

$$H^{*}(TM) \triangleq H'(TM, TM)$$

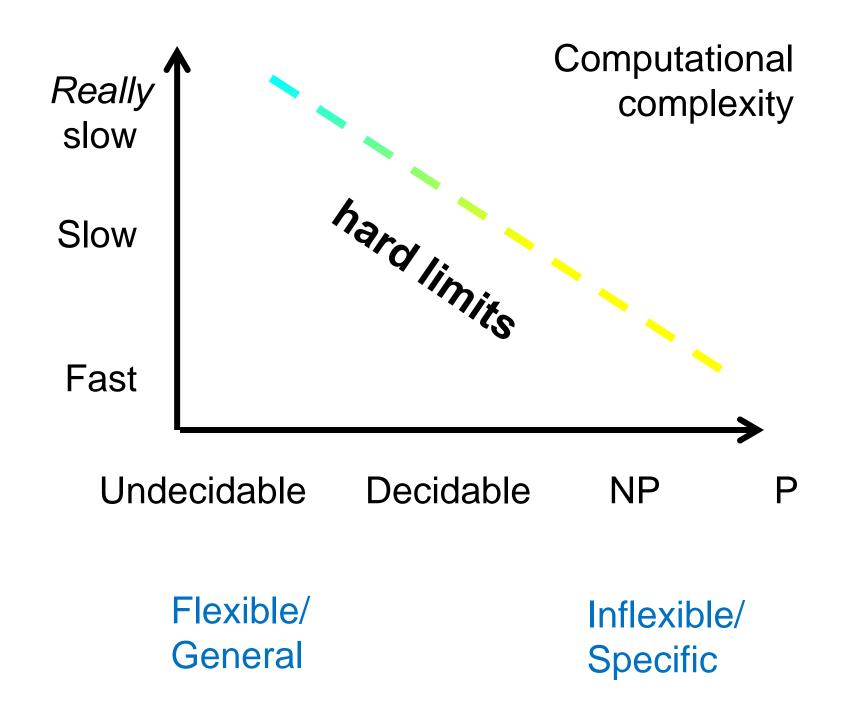
Run $H^*(H^*) = H'(H^*, H^*)$ = $\begin{cases} \text{halt if } H^*(H^*) \text{ loops forever} \\ \text{loop forever otherwise} \end{cases}$

Contradiction!

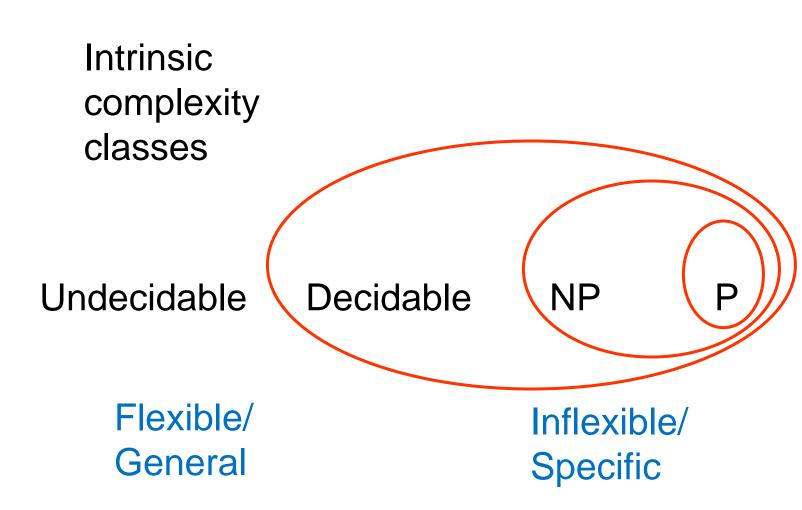


Implications

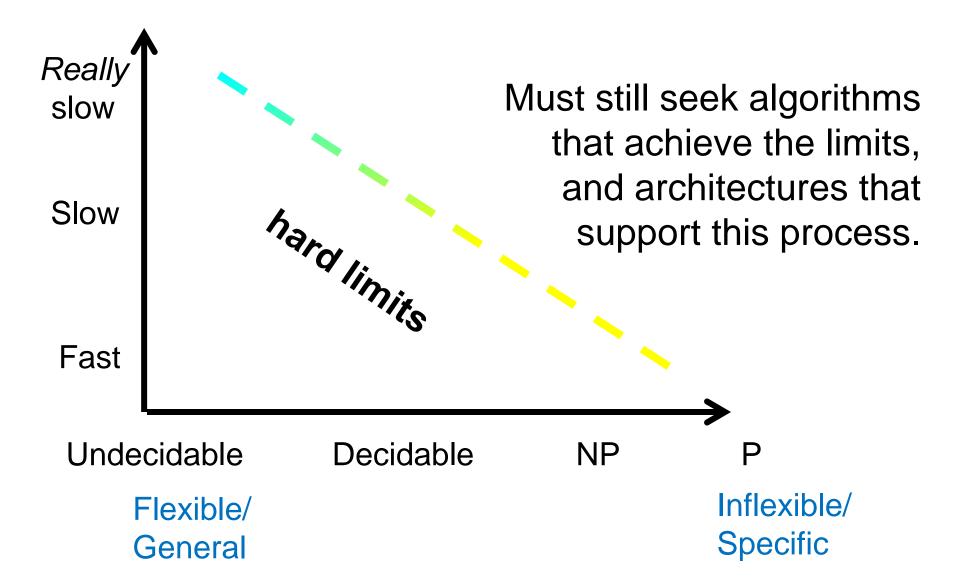
- Large, thin, nonconvex everywhere...
- TMs and UTMs are perfectly repeatable
- But perfectly unpredictable
- Undecidable: Will a TM halt? Is a TM a UTM? Does a TM do X (for almost any X)?
- Easy to make UTMs, but hard to recognize them.
- Is anything decidable? Yes, questions NOT about TMs.



Computational complexity



These are hard limits on the *intrinsic* computational complexity of *problems*.



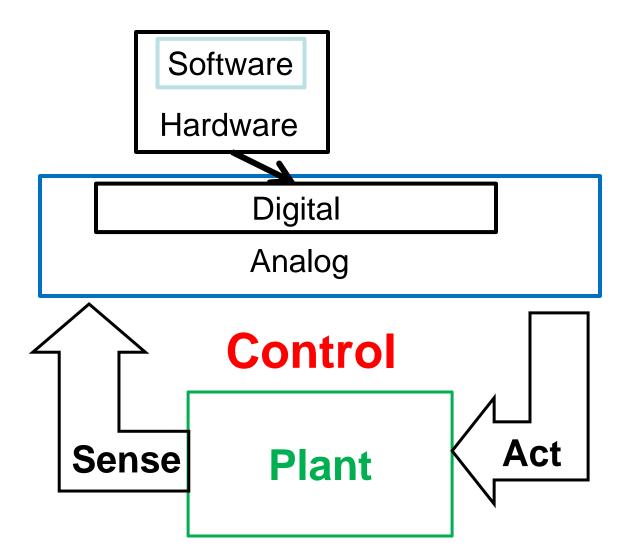
Compute

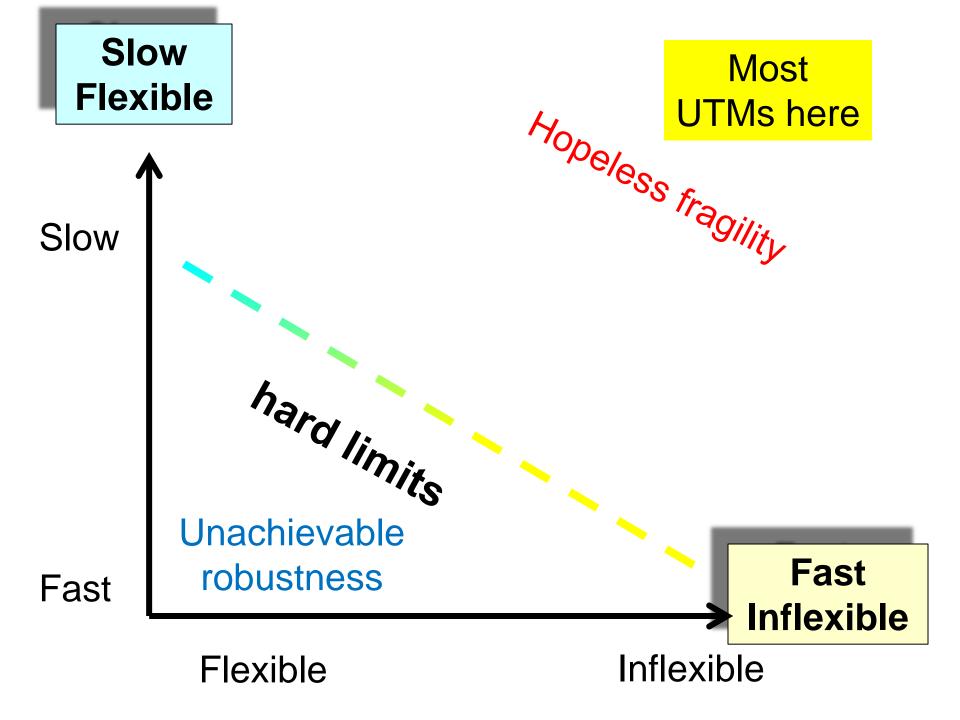
Computational complexity of

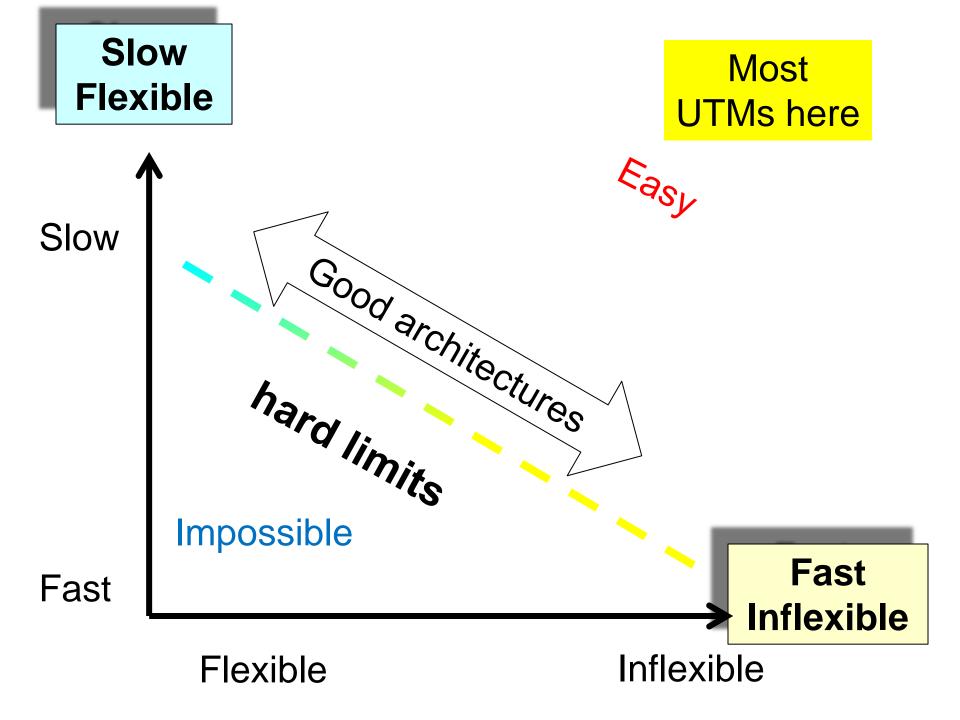
- **Designing** control algorithms
- Implementing control algorithms

Delay is even more important in control









Issues for engineering

- Turing remarkably relevant for 76 years
- UTMs are \approx implementable
 - Differ only (but greatly) in speed and programmability
 - Time/speed/delay is most critical resource
 - Space (memory) almost free for most purposes
- Read/write random access memory hierarchies
- Further gradations of decidable (P/NP/coNP)
- Most crucial:
 - UTMs differ vastly in speed, usability, and programmability
 - You can fix bugs but it is hard to automate finding/avoiding them

Issues for engineering

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 - You can fix bugs but it is hard to automate finding/avoiding them

Conjectures, biology

- Memory potential $\approx \infty$
- Examples
 - Insects
 - Scrub jays
 - Autistic Savants

Gallistel and King

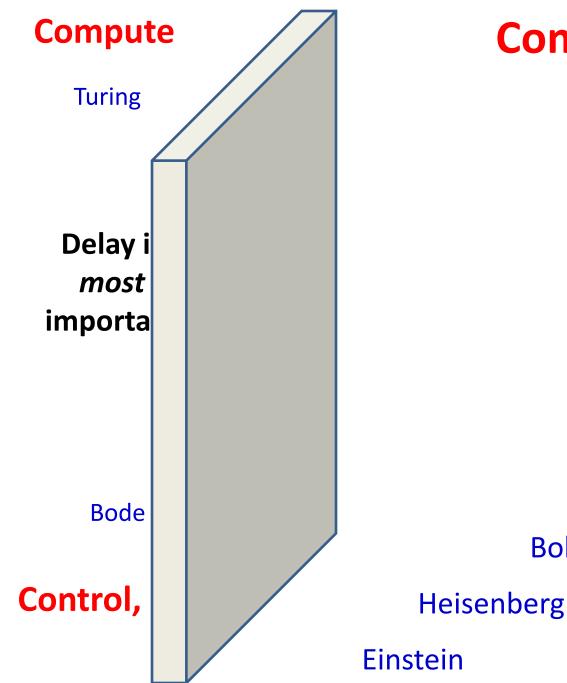


Memory and the Computational Brain

Why Cognitive Science Will Transform Neuroscience

WILEY-BLACKWELL

- But why so rare and/or accidental?
- Large memory, computation of limited value?
- Selection favors fast robust action?
- Brains are distributed (not studied by Gallistel)



Communicate

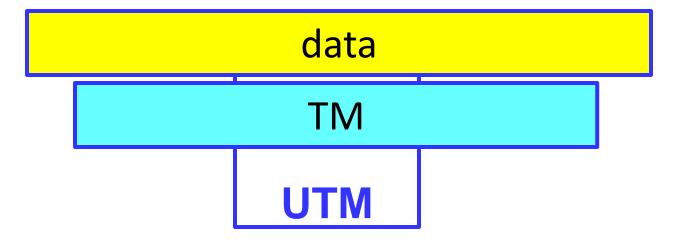
Shannon

Delay is least important

Carnot

Boltzmann





• Suppose we only care about space?

- And time is free
- Bad news: compression undecidable.
- Shannon: change the problem!



time

Communications

Shannon's brilliant insight

- Forget time
- Forget files, use *infinite random ensembles*

Good news

- Laws and architecture!
- Info theory most popular and accessible topic in systems engineering
- *Fantastic* for some engineering problems

Shannon

Communications

Shannon

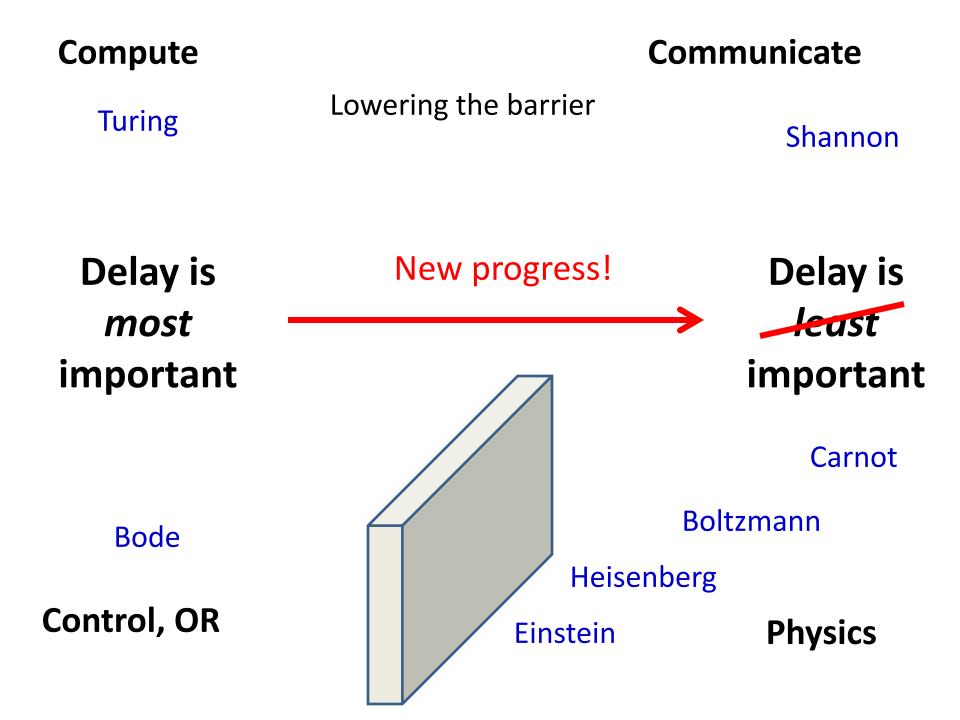
Shannon's brilliant insight

- Forget time
- Forget files, use *infinite random ensembles*

Bad news

• Laws and architecture very brittle

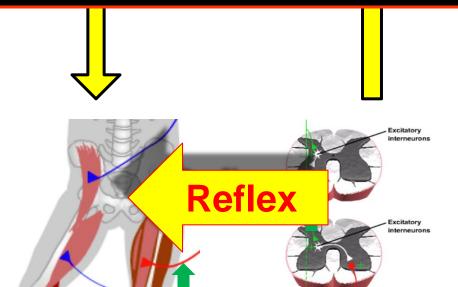
- Less than zero impact on internet architecture
- Almost useless for biology (But see Lestas et al, 2010)
- Misled, distracted generations of biologists (and neuroscientists)

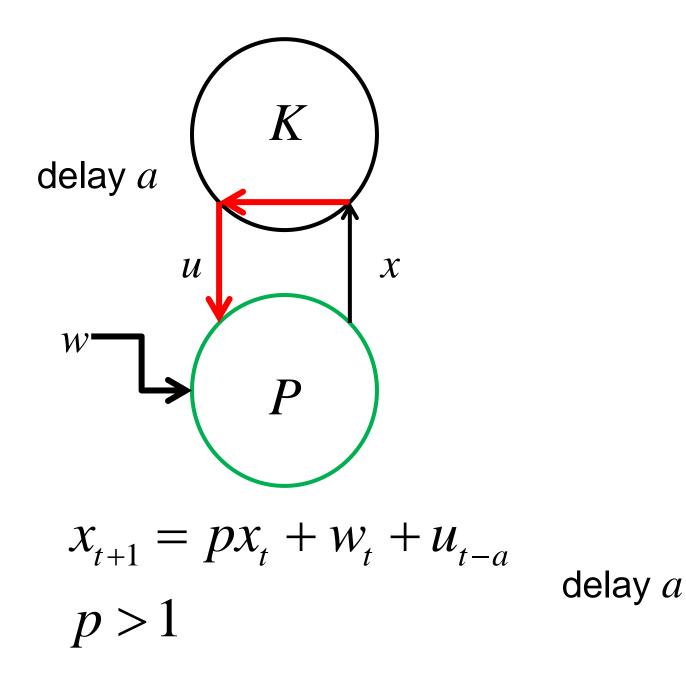


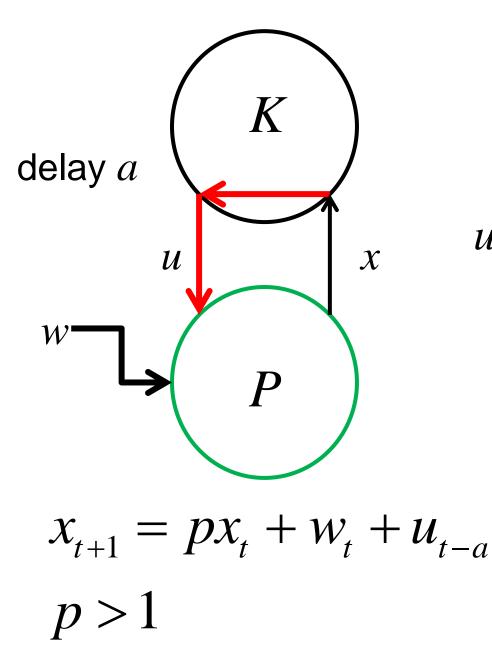
Delay is even more important

Wolpert, Grafton, etc *nobust* Brain as optimal controller

- Acquire
- Translate/ integrate
- Automate

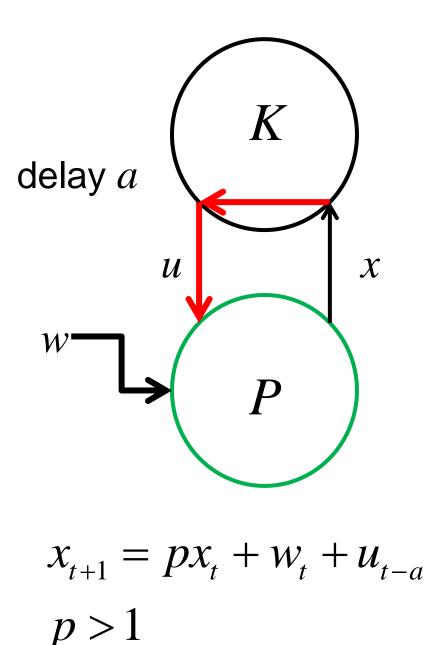






No delay or no uncertainty

$$u_{t-a} = -(px_t + w_t)$$
$$\implies ||x|| \approx 0 \quad ||u|| \approx ||w||$$

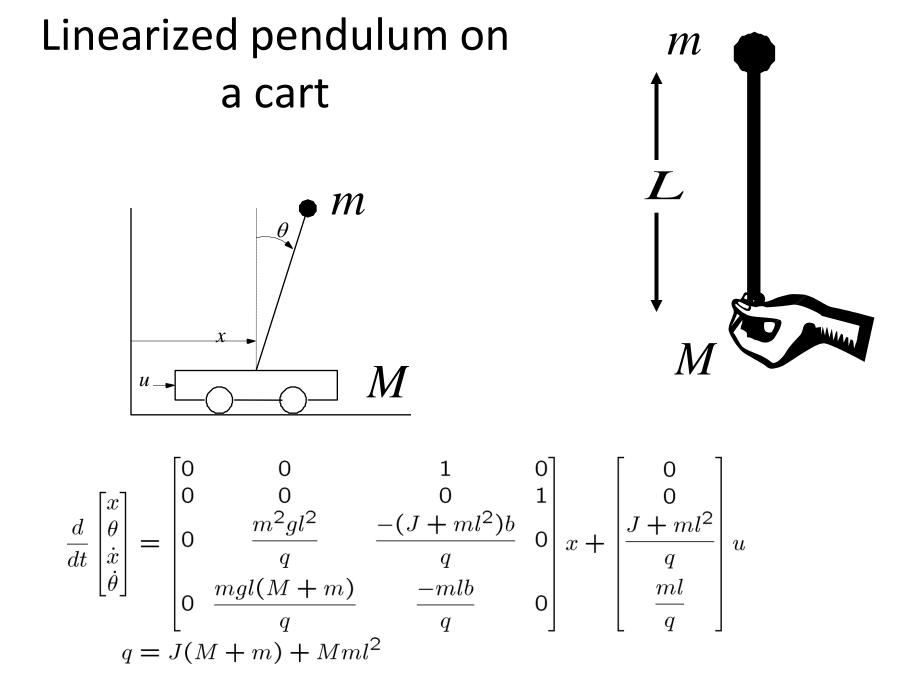


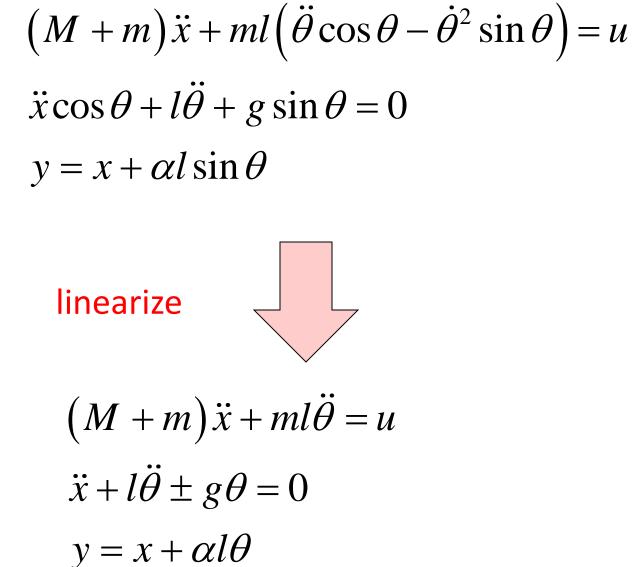
No delay or no uncertainty

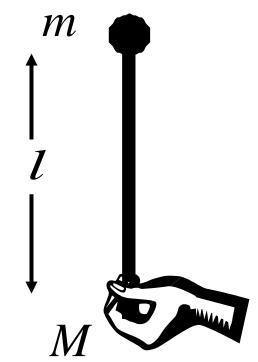
$$u_{t-a} = -(px_t + w_t)$$
$$\Rightarrow ||x|| \approx 0 \quad ||u|| \approx ||w||$$

With delay **and** uncertainty

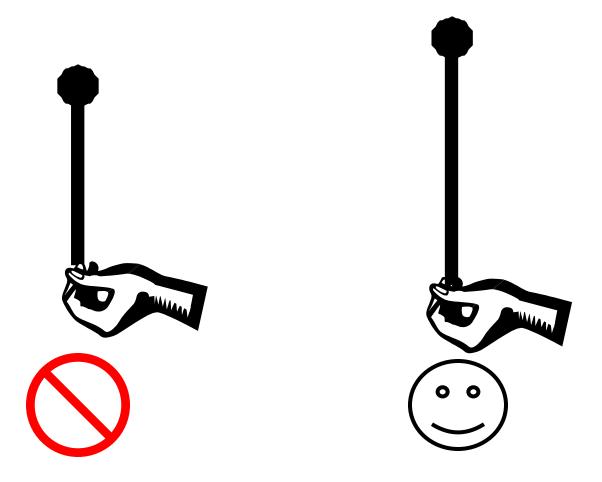
 $\Rightarrow \|x\| \approx \|u\| \approx p^a \|w\|$







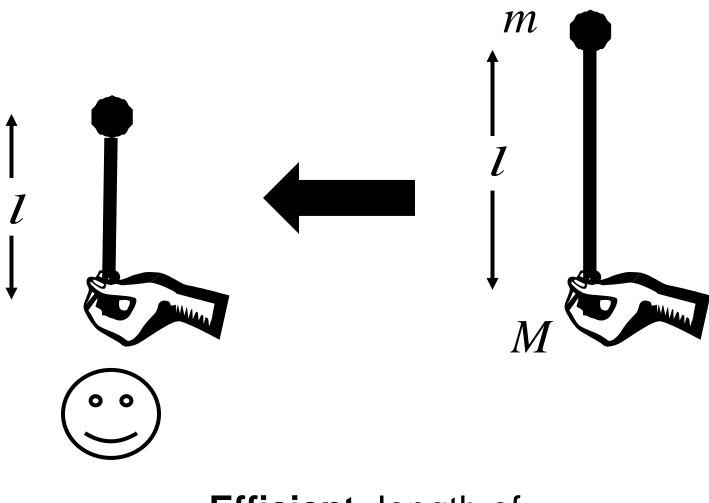
Robust =agile and balancing



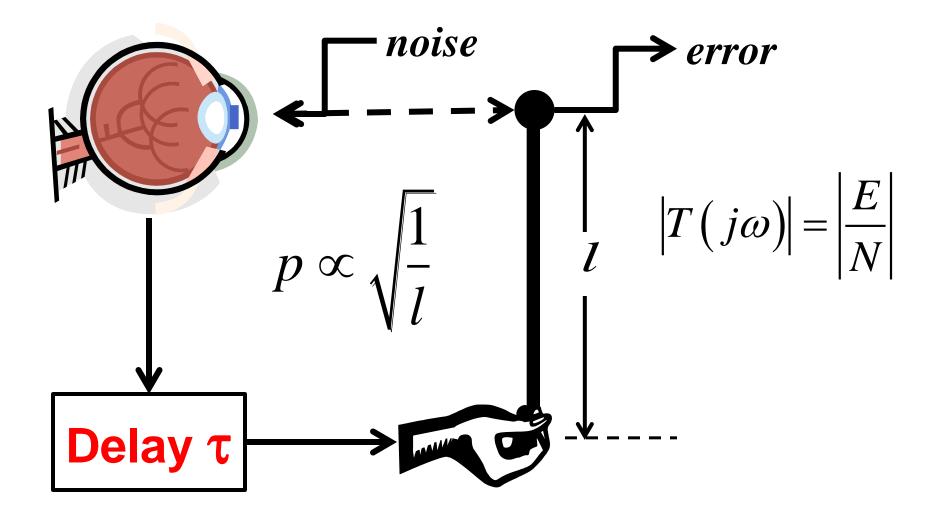
Robust =agile and balancing







Efficient=length of pendulum (artificial)





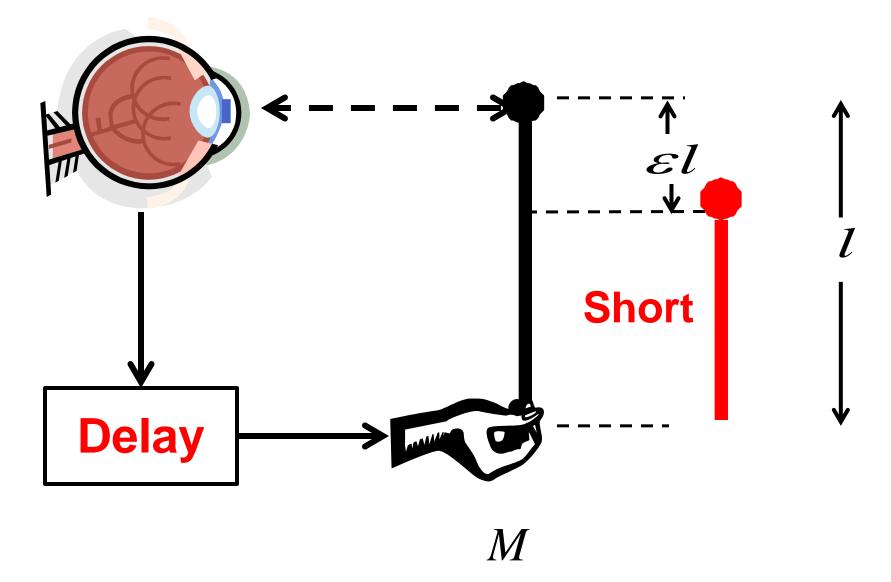
 $\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|d\omega\geq0$



Easy, even with eyes closed No matter what the length

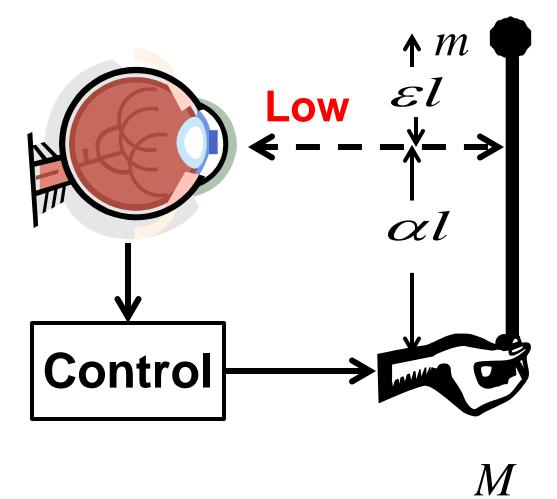
Proof: Standard UG control theory: Easy calculus, easier contour integral, easiest Poisson Integral formula

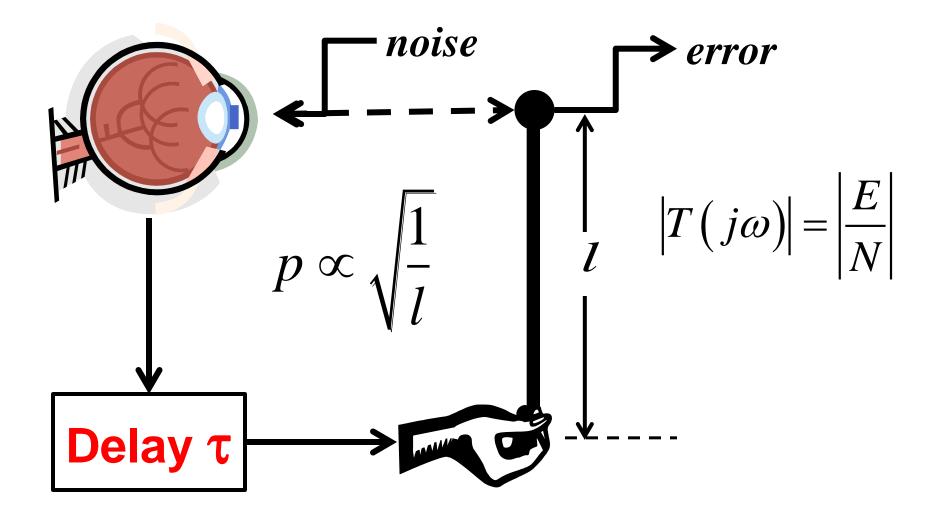
Harder if delayed or short

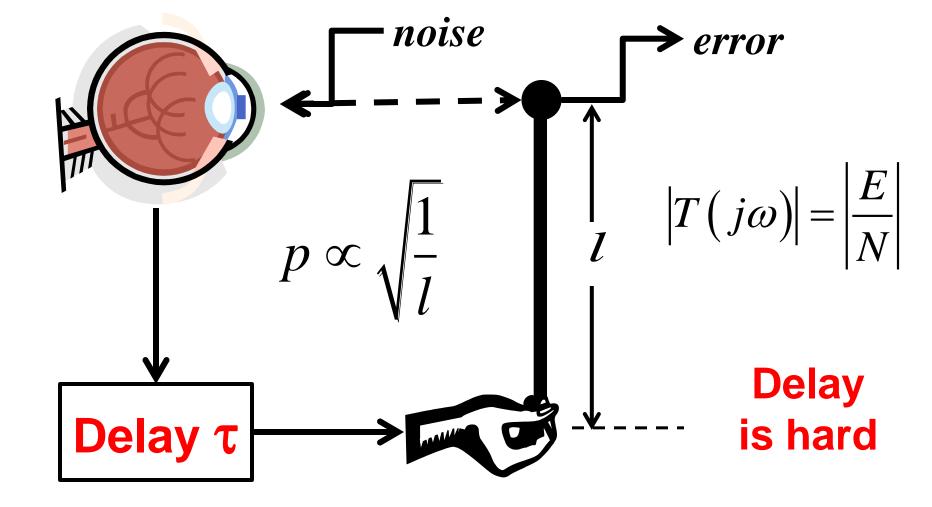


Also harder if sensed low

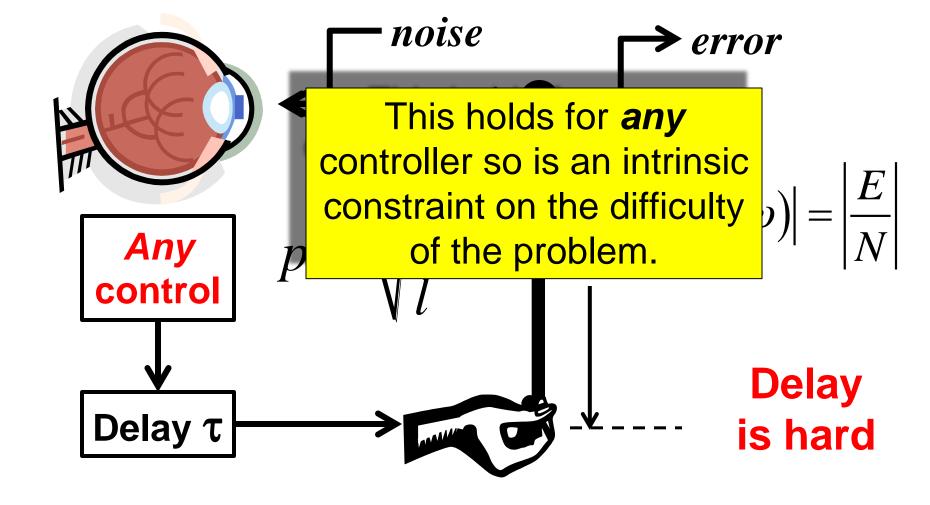
 $r = \frac{m}{M}$





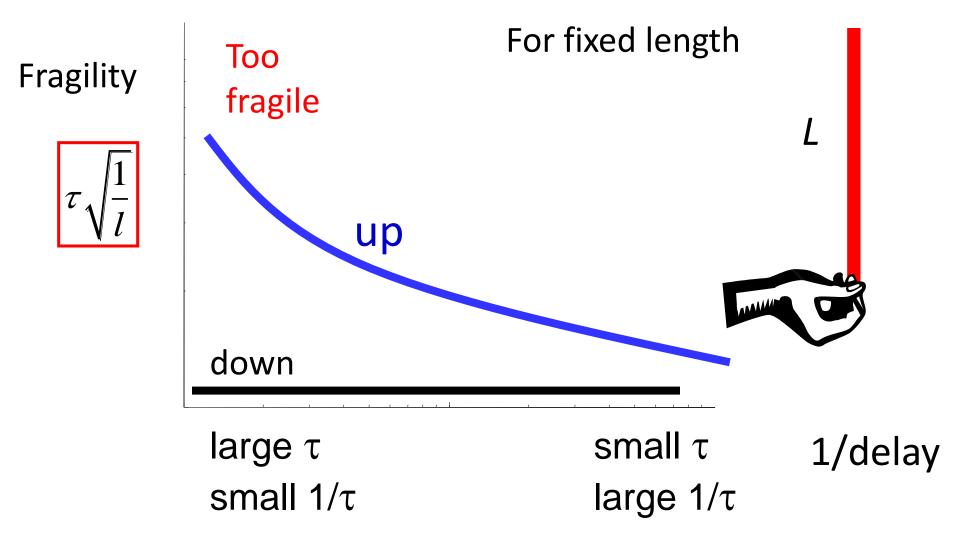


$$\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega\geq\ln\left|T_{mp}(p)\right|=p\tau\propto\tau\sqrt{\frac{1}{l}}$$



 $\frac{1}{\pi}\int_{0}^{1}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega\geq\ln\left|T_{mp}(p)\right|=p\tau\propto\tau\sqrt{\frac{1}{I}}$

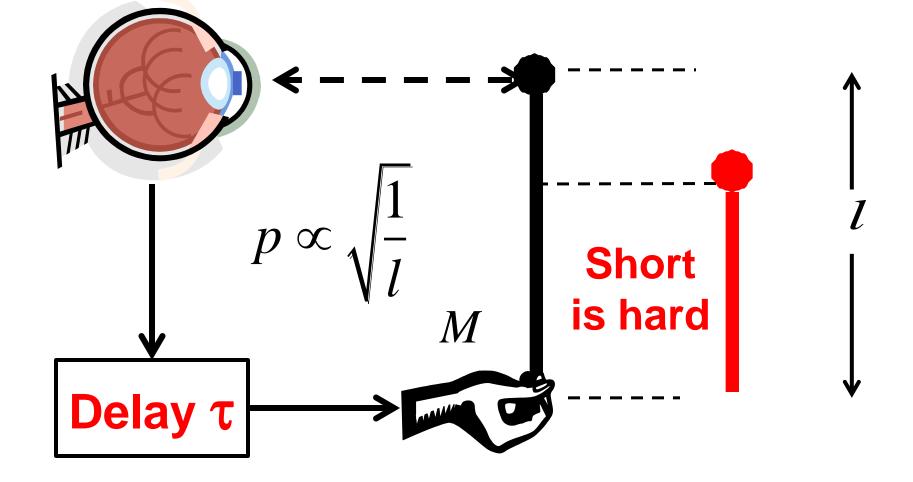
 $\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega \ge p\tau \propto \tau\sqrt{\frac{1}{l}}$



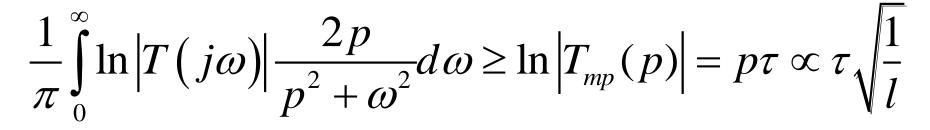
$$\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega\geq p\tau\propto\tau\sqrt{\frac{1}{l}}$$

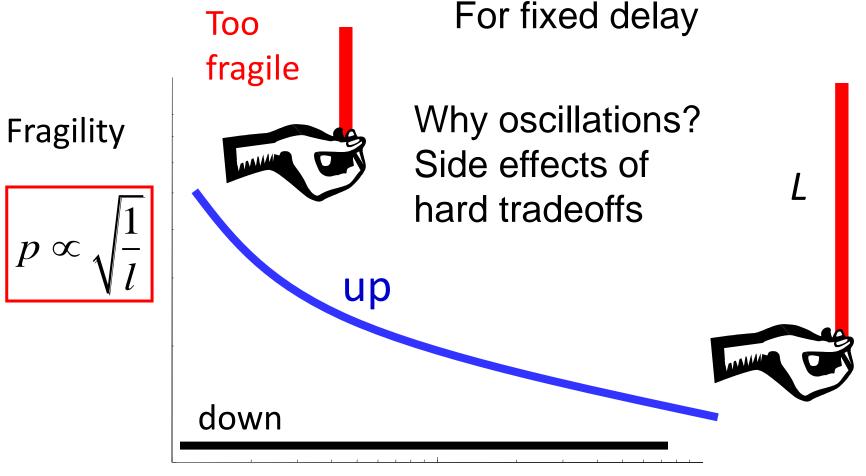
We would like to tolerate large delays (and small lengths), but large delays severely constrain the achievable robustness.

large τ small $1/\tau$ small τ large 1/ τ



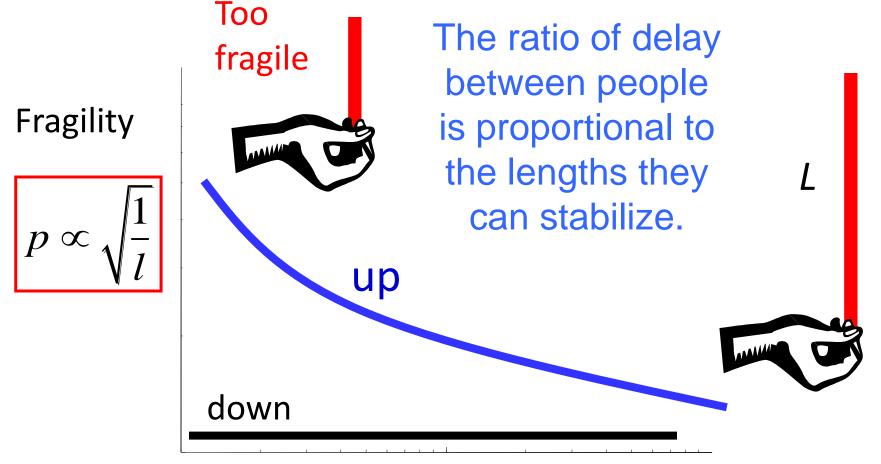
 $\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega\geq\ln\left|T_{mp}(p)\right|=p\tau\propto\tau\sqrt{\frac{1}{I}}$





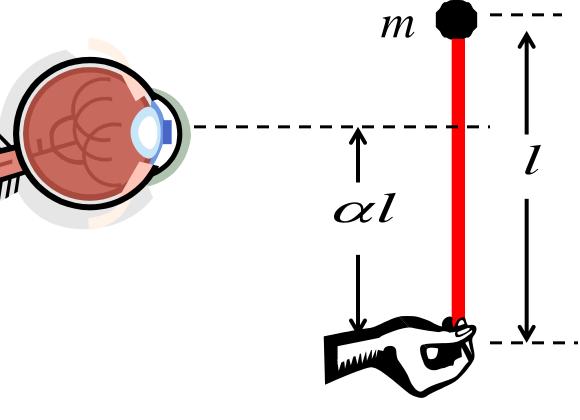
length L

 $\frac{1}{\pi}\int_{\Omega}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega\geq\ln\left|T_{mp}(p)\right|=p\tau\propto\tau\sqrt{\frac{1}{l}}$

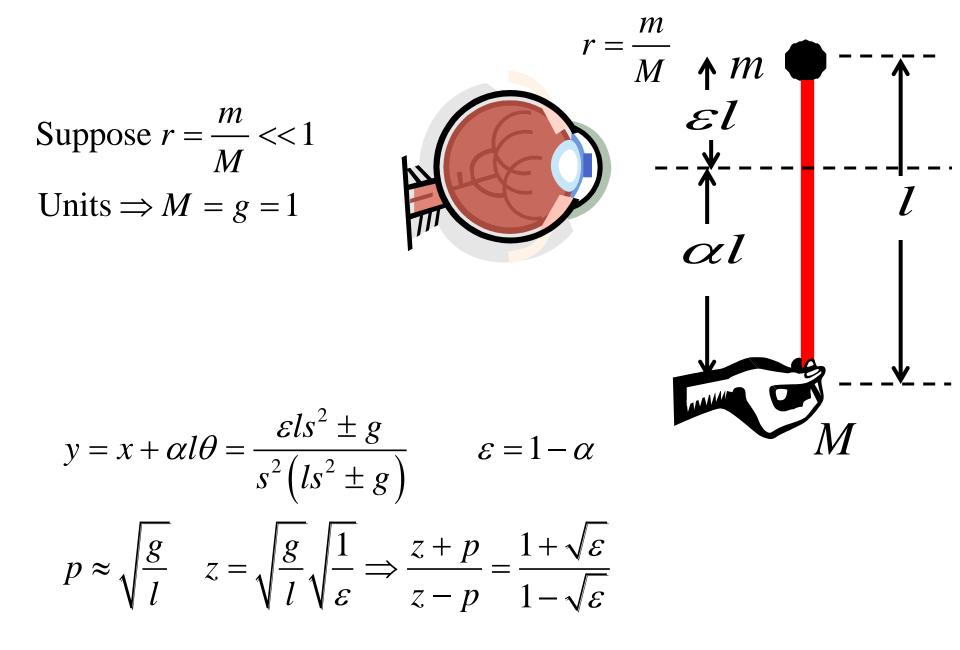


length L

Eyes moved down is harder (RHP zero) Similar to delay



M



Compare

$$p = \sqrt{\frac{g}{l(1-\varepsilon)}} \sqrt{1+r} = p_0 \sqrt{\frac{1}{(1-\varepsilon)}} \approx p_0 \left(1 + \frac{\varepsilon}{2}\right)$$

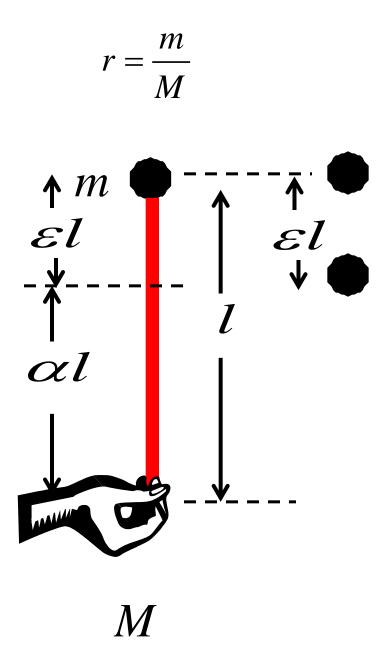
Move eyes

$$p = \sqrt{\frac{g}{l}}\sqrt{1+r} \quad r = \frac{m}{M} \quad z = \sqrt{\frac{g}{l}}\sqrt{\frac{1}{\varepsilon}}$$

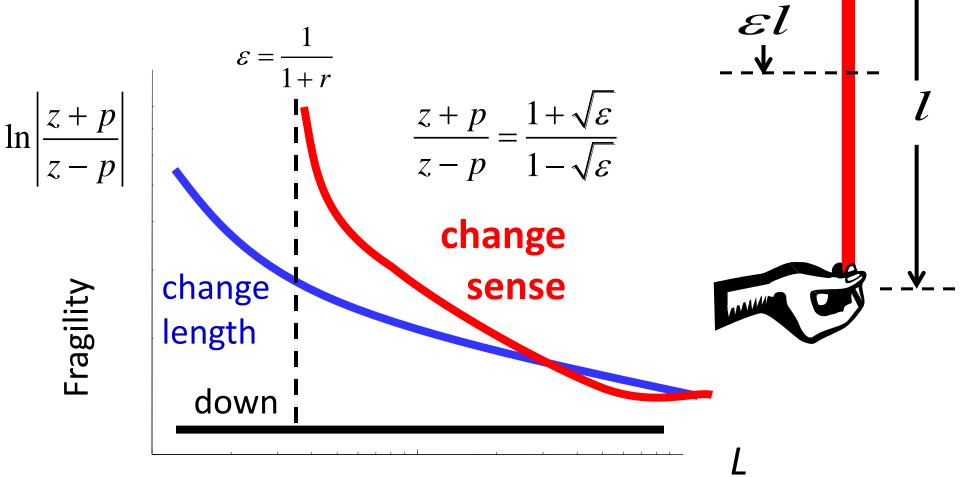
$$p = z \Longrightarrow 1+r = \frac{1}{\varepsilon} \Longrightarrow \varepsilon = \frac{1}{1+r}$$

$$p\left(1+\frac{1}{3}\frac{p^2}{z^2}\right) = \sqrt{\frac{g}{l}}\sqrt{1+r}\left(1+\frac{1}{3}\varepsilon\right) = p\left(1+\frac{\varepsilon}{3}\right)$$

$$= p\left(1+\frac{1-\alpha}{3}\right)$$

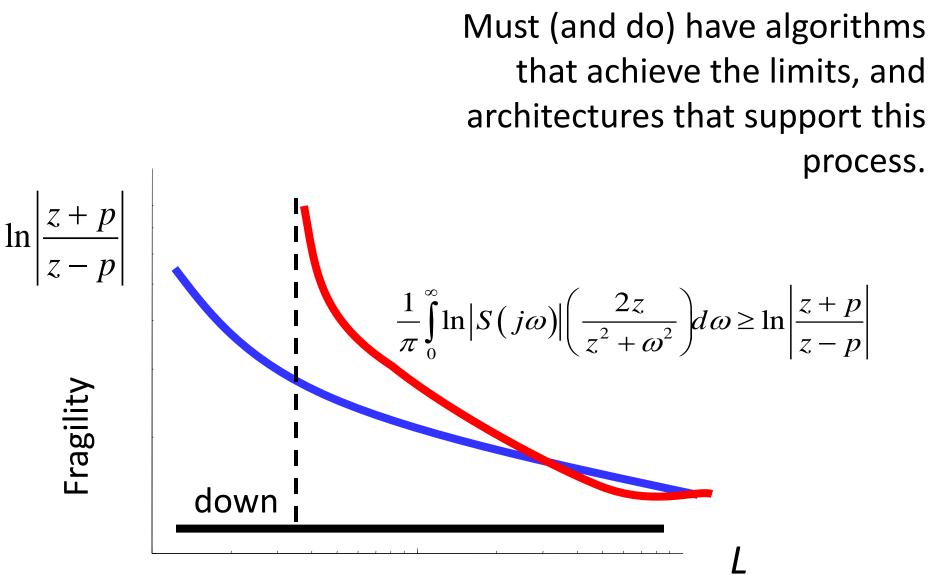


$$\frac{1}{\pi}\int_{0}^{\infty}\ln\left|S\left(j\omega\right)\right|\left(\frac{2z}{z^{2}+\omega^{2}}\right)d\omega \ge \ln\left|\frac{z+p}{z-p}\right|$$
$$\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\left(\frac{2p}{p^{2}+\omega^{2}}\right)d\omega \ge \ln\left|\frac{z+p}{z-p}\right|$$

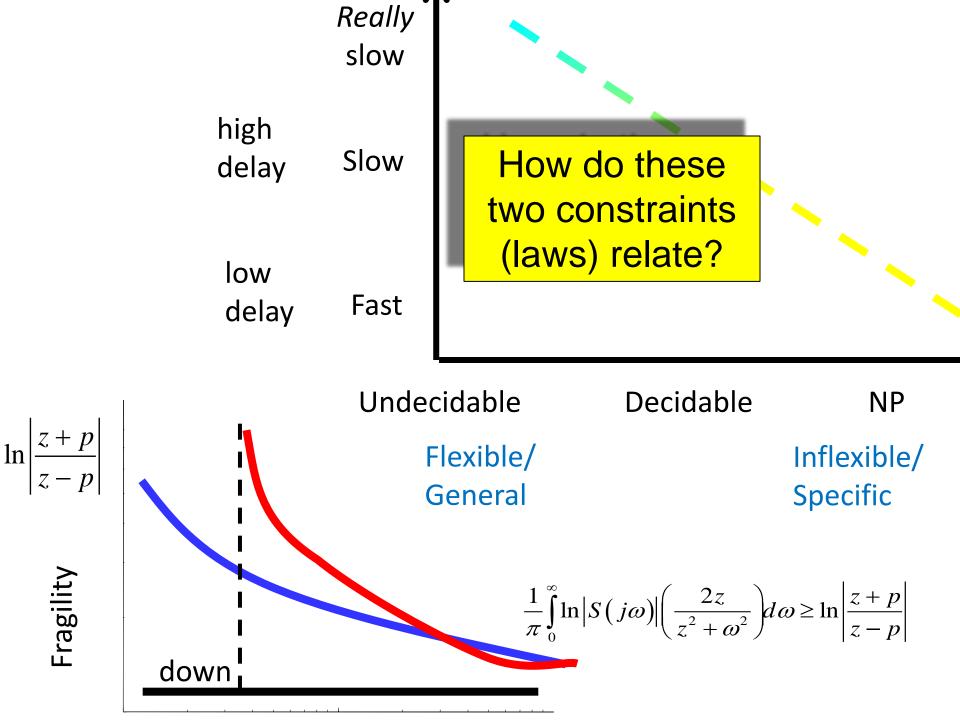


This is a cartoon, but can be made precise.

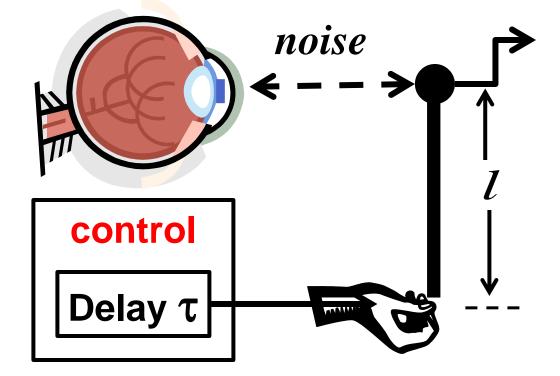
Hard limits on the *intrinsic* robustness of control *problems*.



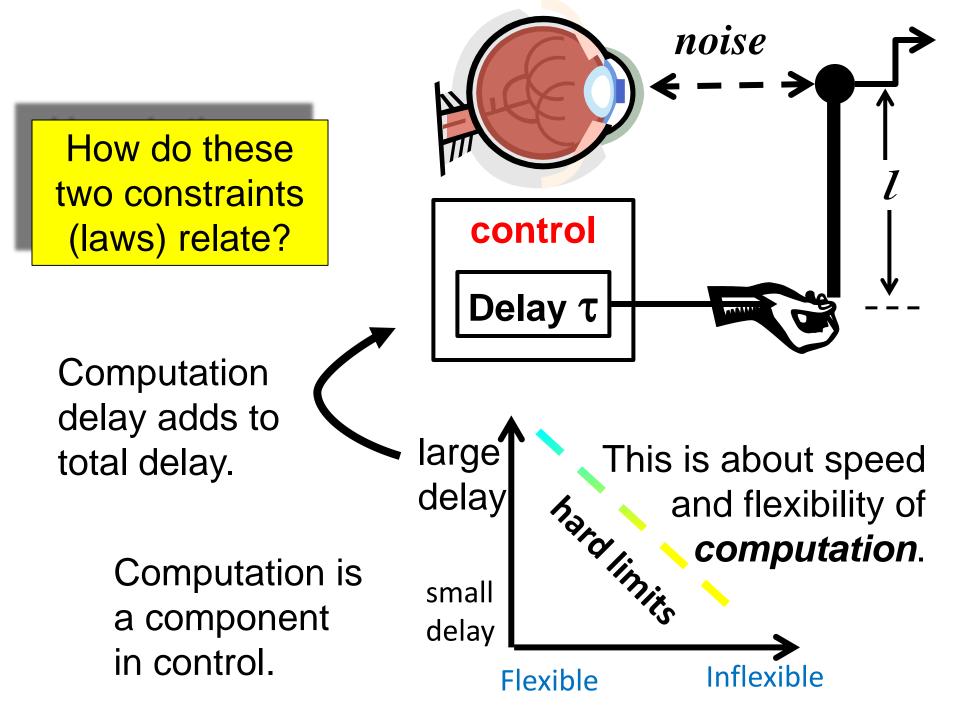
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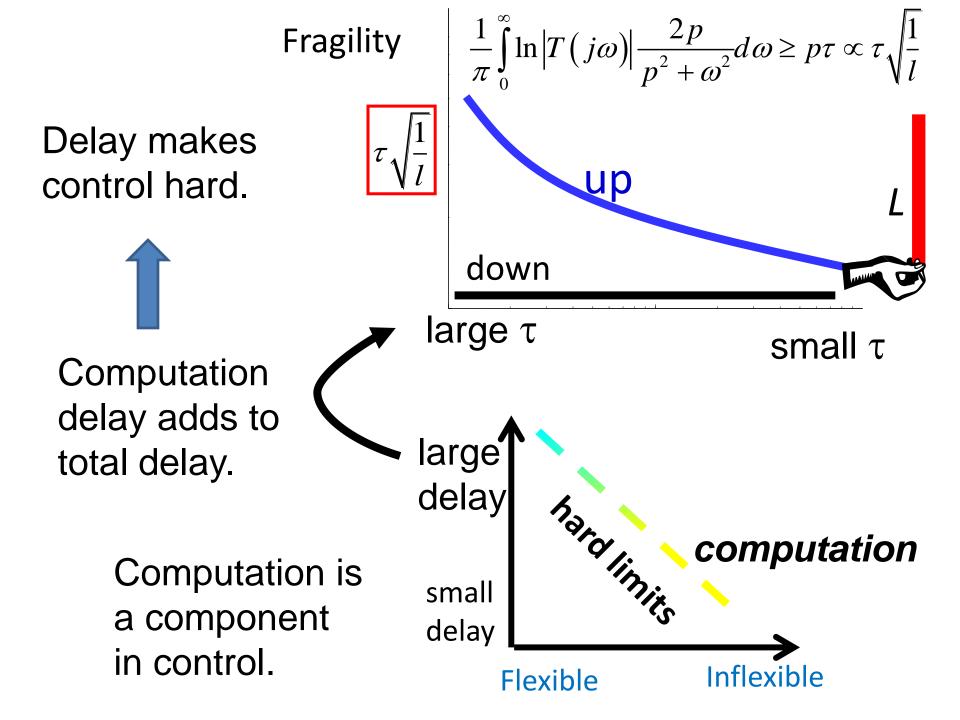


Delay comes from sensing, communications, computing, and actuation. Delay limits robust performance.

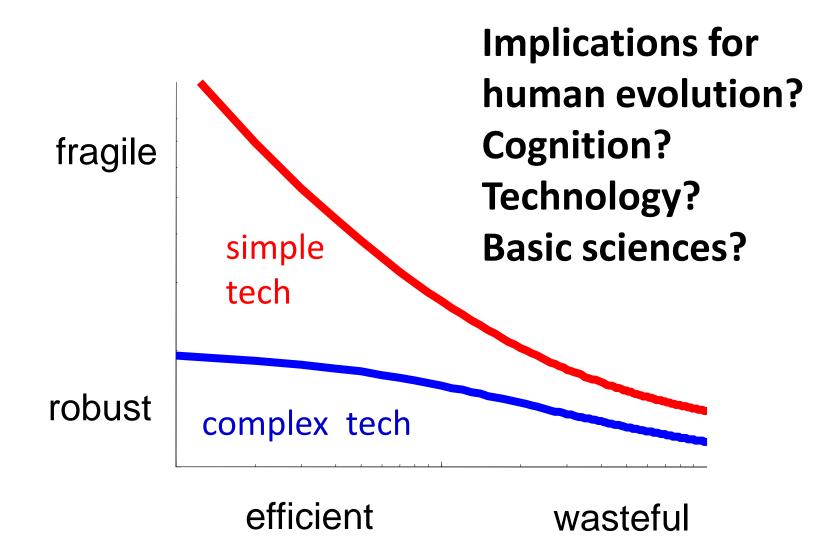


$$\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega\geq\ln\left|T_{mp}(p)\right|=p\tau\propto\tau\sqrt{\frac{1}{l}}$$





How general is this picture?



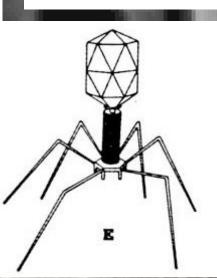
Viruses' Life History: Towards a Mechanistic Basis of a Trade-Off between Survival and Reproduction among Phages

Marianne De Paepe, François Taddei^{*}

Laboratoire de Genetique Moleculaire, Evolutive et Medicale, University of Paris 5, INSERM, Paris, France

July 2006 | Volume 4 | Issue 7 | e193

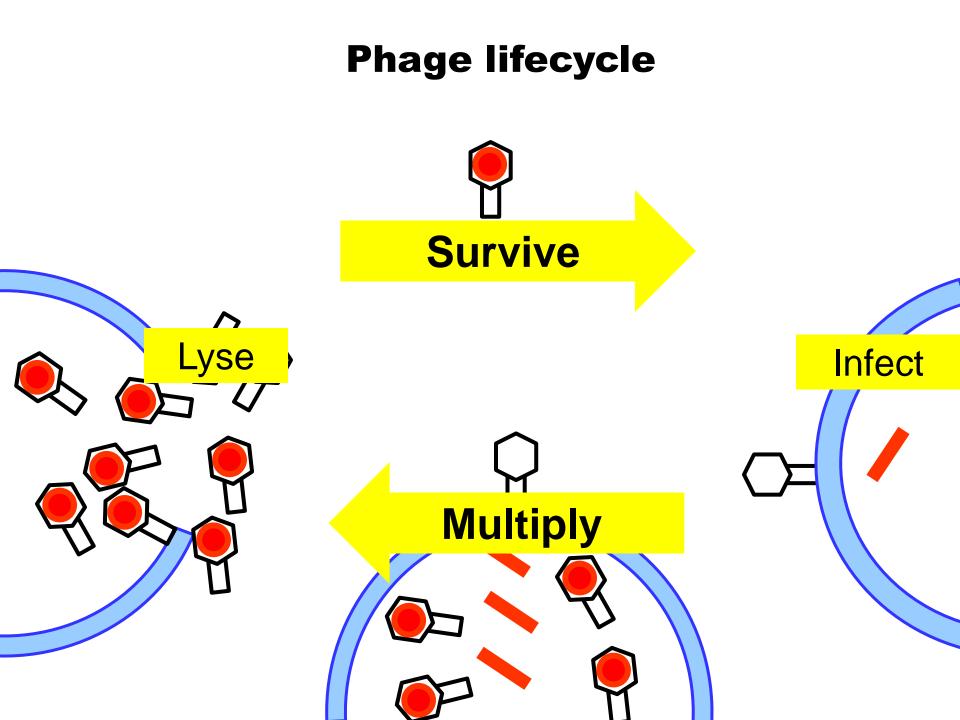
I recently found this paper, a rare example of exploring an explicit tradeoff between robustness and efficiency. This seems like an important paper but it is rarely cited.

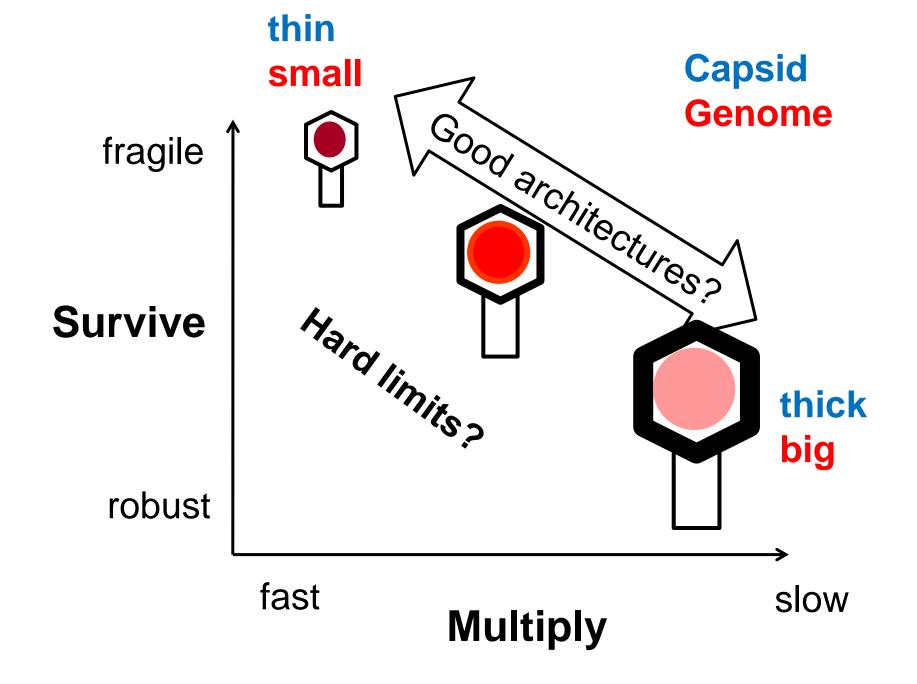


Phage

1μm

Bacteria





Glycolytic Oscillations and Limits on Robust Efficiency

Fiona A. Chandra,¹* Gentian Buzi,² John C. Doyle²

Both engineering and evolution are constrained by trade-offs between efficiency and robustness, but theory that formalizes this fact is limited. For a simple two-state model of glycolysis, we explicitly derive analytic equations for hard trade-offs between robustness and efficiency with oscillations as an inevitable side effect. The model describes how the trade-offs arise from individual parameters, including the interplay of feedback control with autocatalysis of network products necessary to power and catalyze intermediate reactions. We then use control theory to prove that the essential features of these hard trade-off "laws" are universal and fundamental, in that they depend minimally on the details of this system and generalize to the robust efficiency of any autocatalytic network. The theory also suggests worst-case conditions that are consistent with initial experiments.

Chandra, Buzi, and Doyle

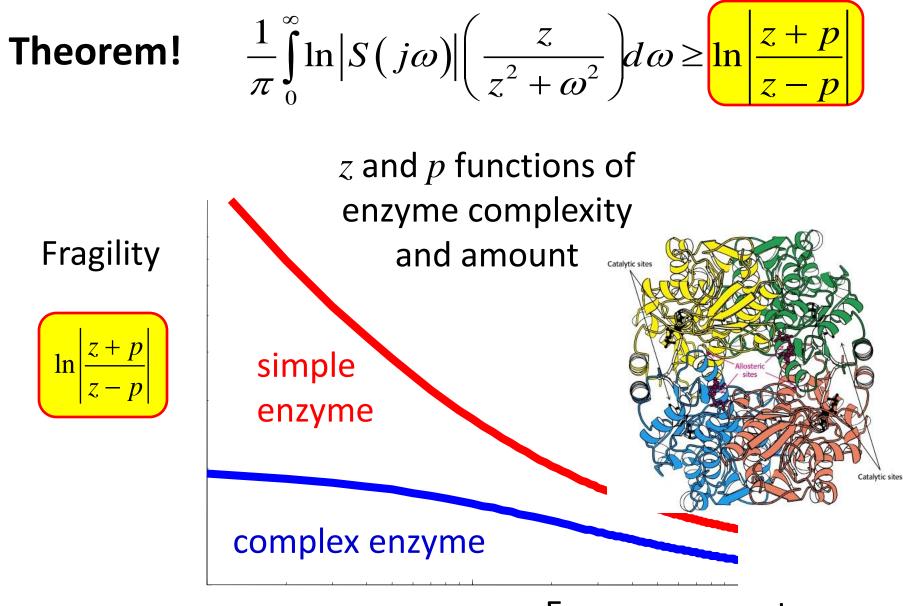
Most important paper so far.

UG biochem, math, control theory

the cen's use of ATF. In giveorysis, two ATP molecules are consumed upstream and four are produced downstream, which normalizes to q = 1(each y molecule produces two downstream) with kinetic exponent a = 1. To highlight essential trade-offs with the simplest possible analysis, we normalize the concentration such that the unperturbed ($\delta = 0$) steady states are $\overline{y} = 1$ and $\overline{x} = 1/k$ [the system can have one additional steady state, which is unstable when (1, 1/k) is stable]. [See the supporting online material (SOM) part I]. The basal rate of the PFK reaction and the consumption rate have been normalized to 1 (the 2 in the numerator and feedback coefficients of the reactions come from these normalizations). Our results hold for more general systems as discussed below and in SOM, but the analysis

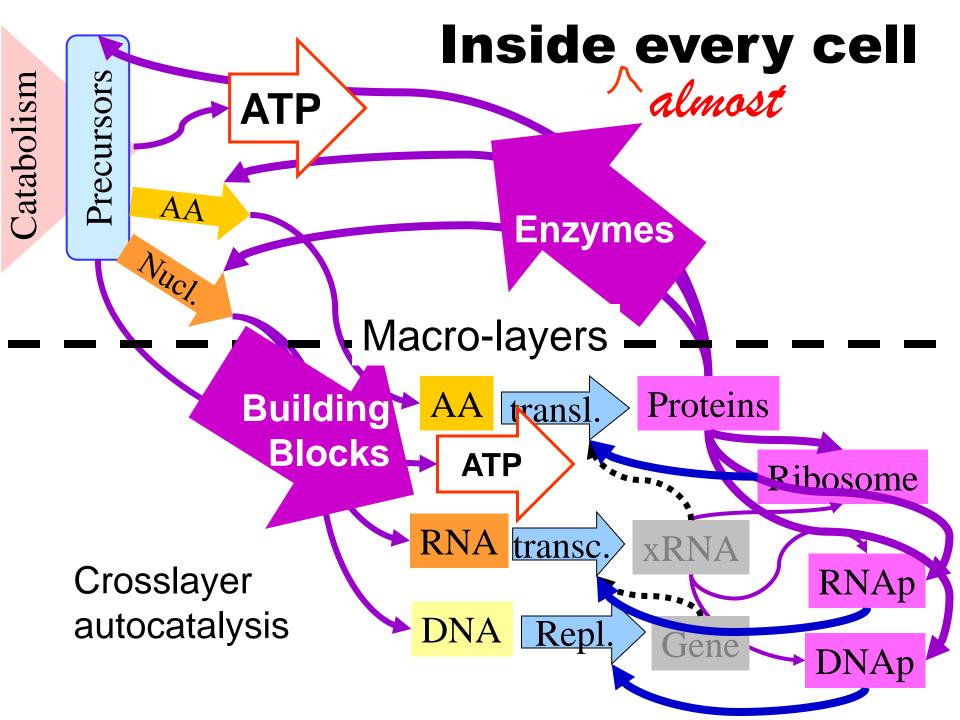


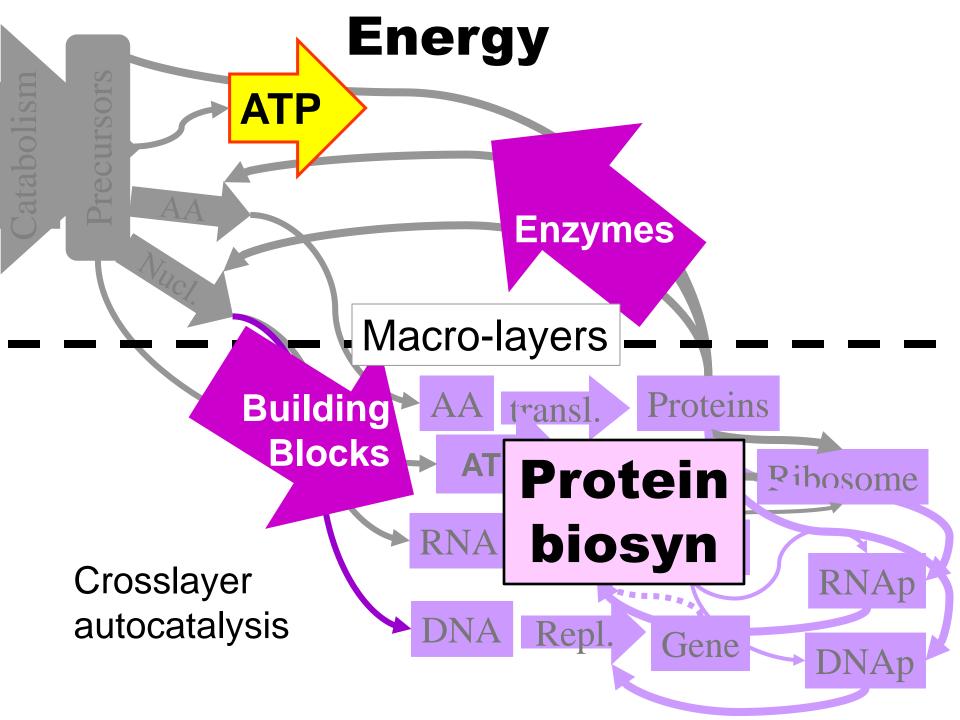
www.sciencemag.org SCIENCE VOL 333 8 JULY 2011

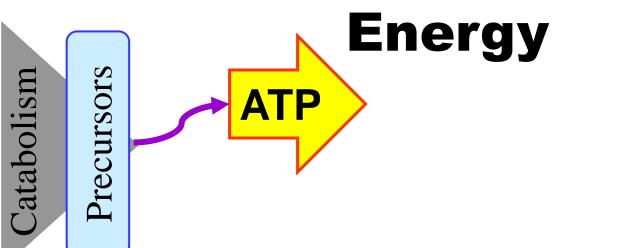


Enzyme amount

Savageaumics



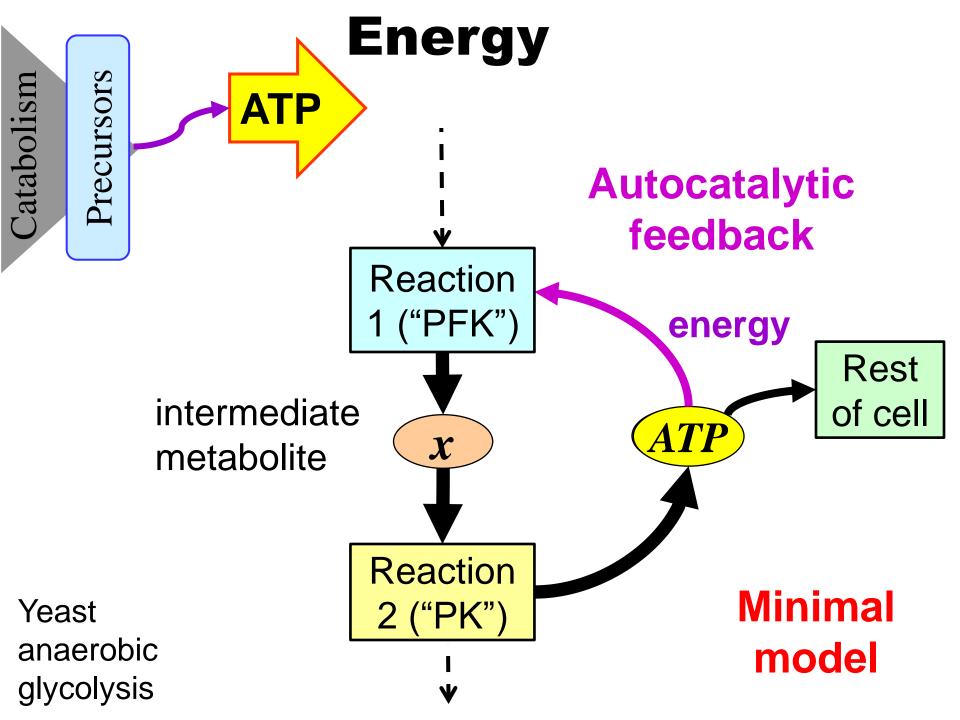




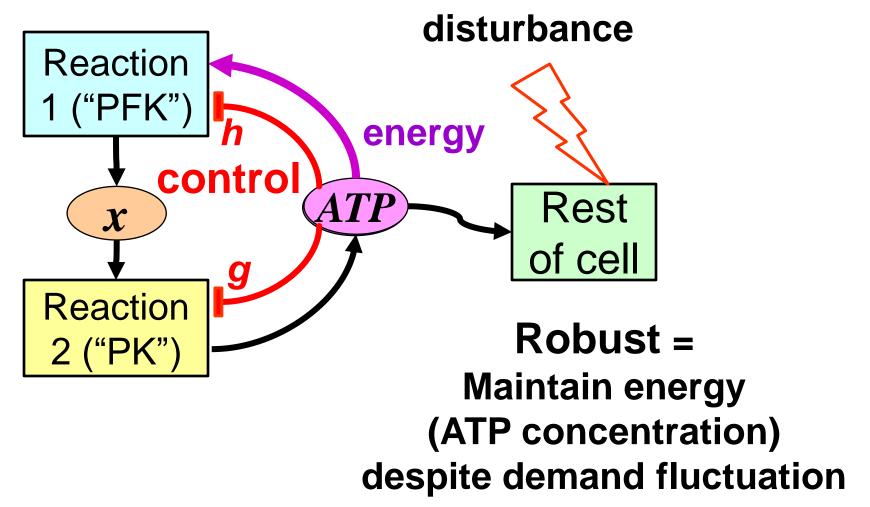
energy Rest of cell

Yeast anaerobic glycolysis

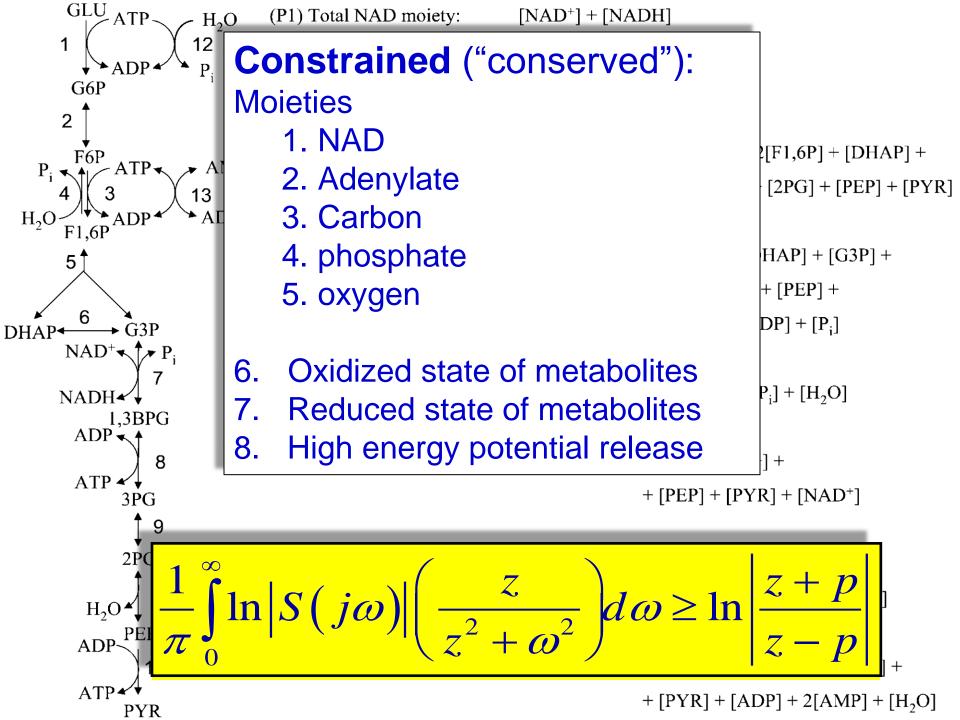




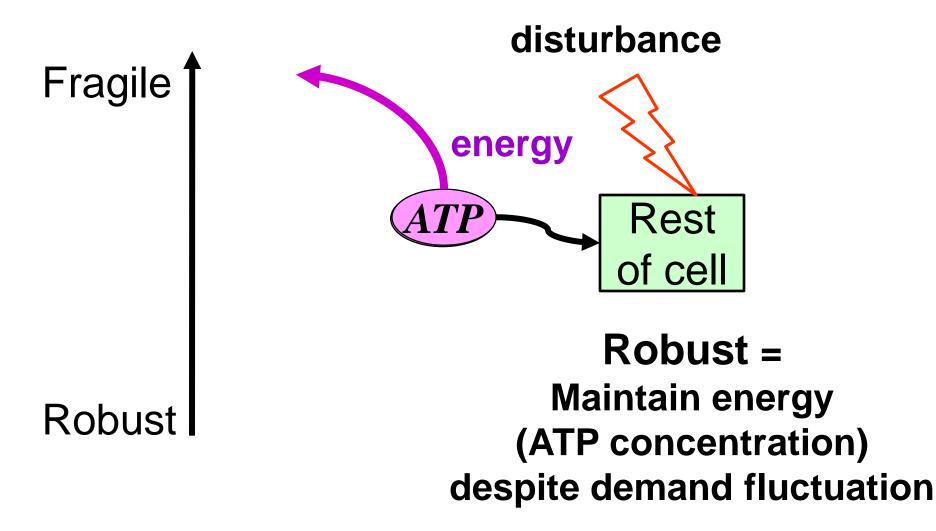
control feedback

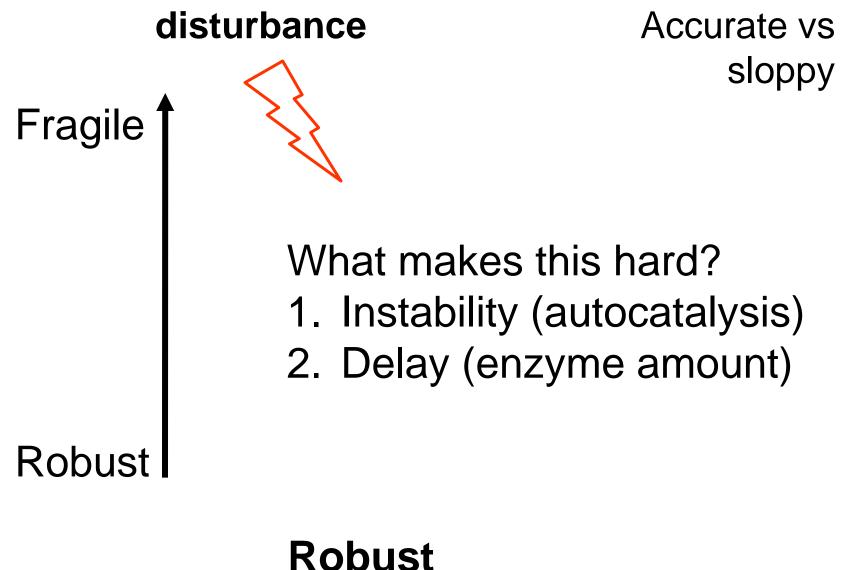


Tight control creates "weak linkage" between power supply and demand



Hard tradeoff in glycolysis





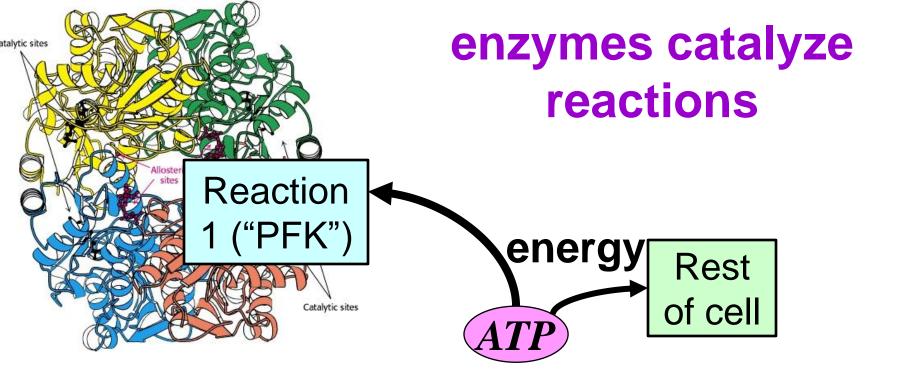


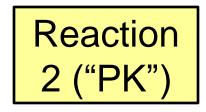
What makes this hard?

- 1. Instability
- 2. Delay

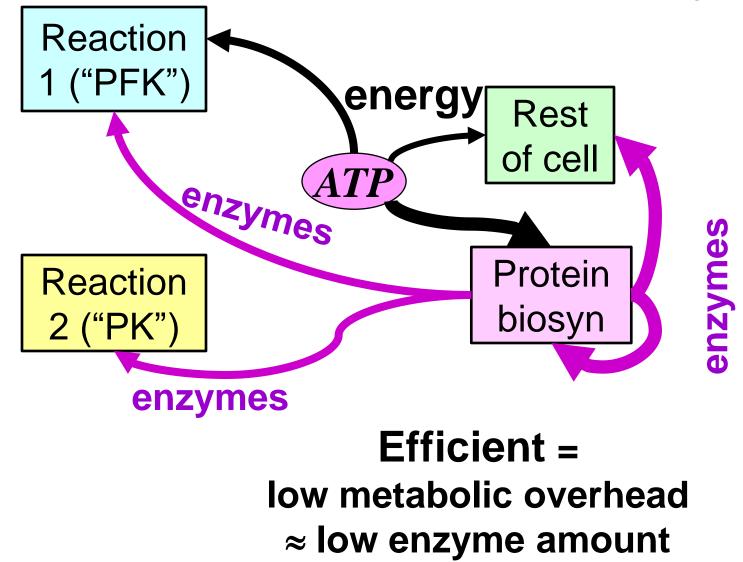
The CNS must cope with both

Today's important point





enzymes catalyze reactions, another source of autocatalysis



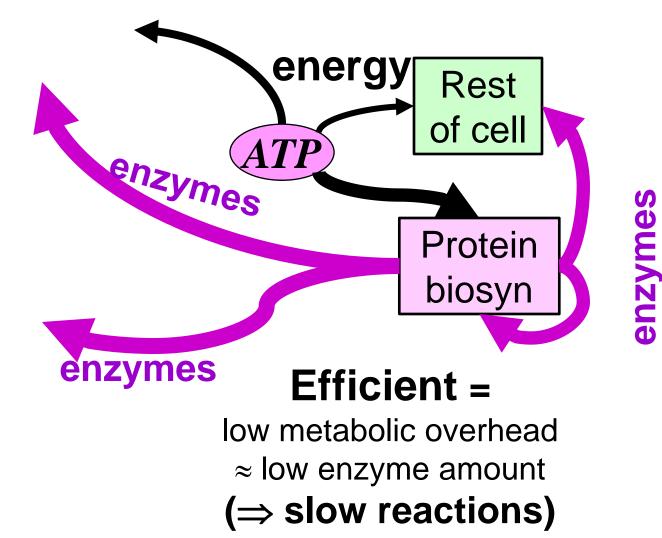
enzymes catalyze reactions, another source of autocatalysis

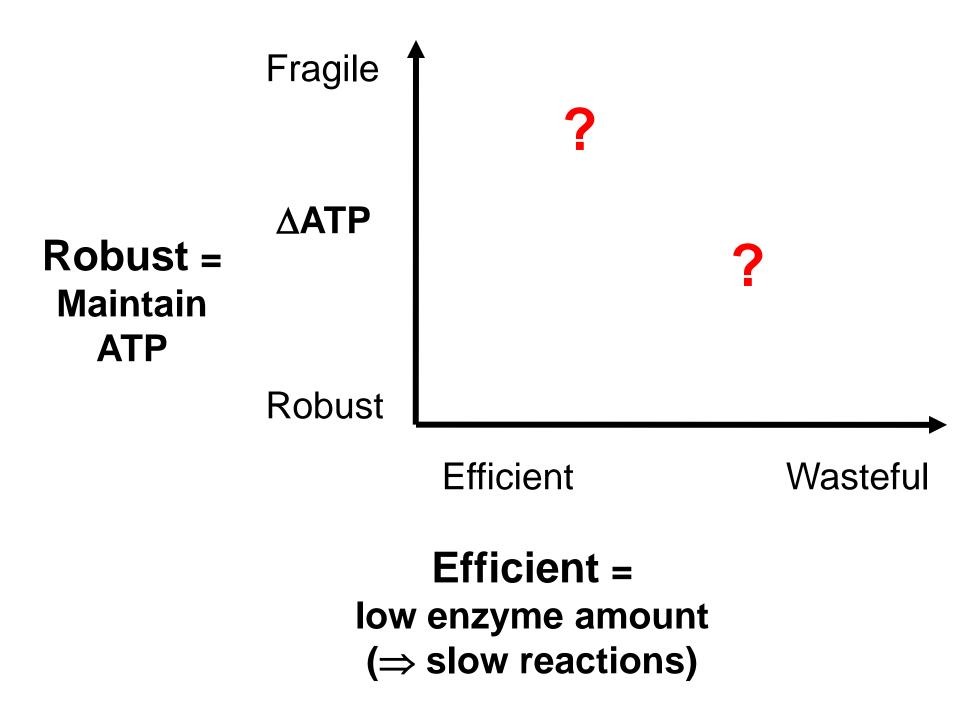
∞ enzyme amount

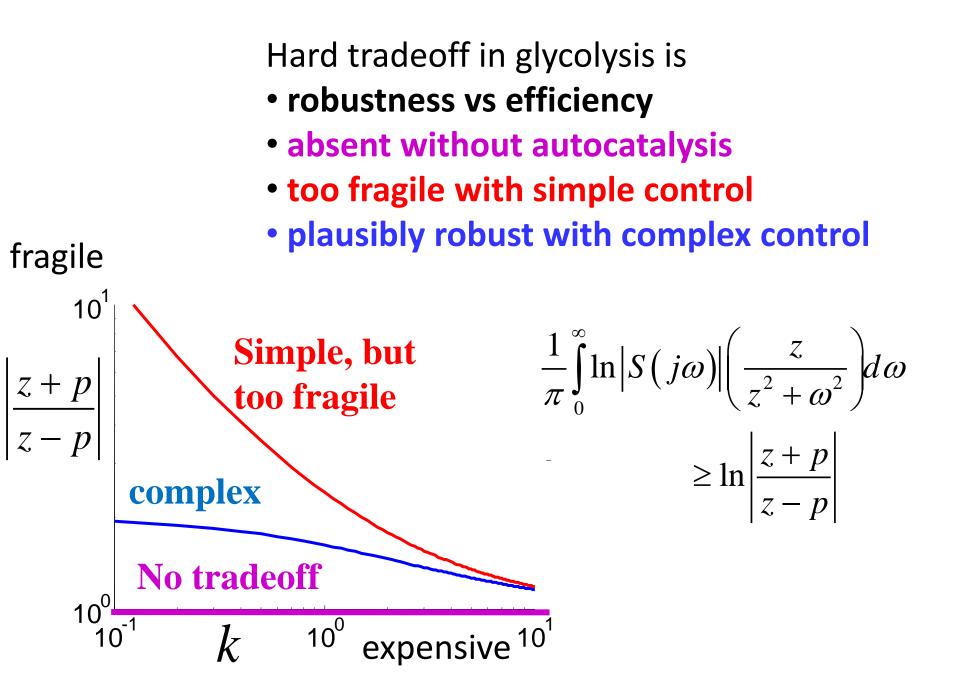
reaction

rates

Can't make too many enzymes here, need to supply rest of the cell.



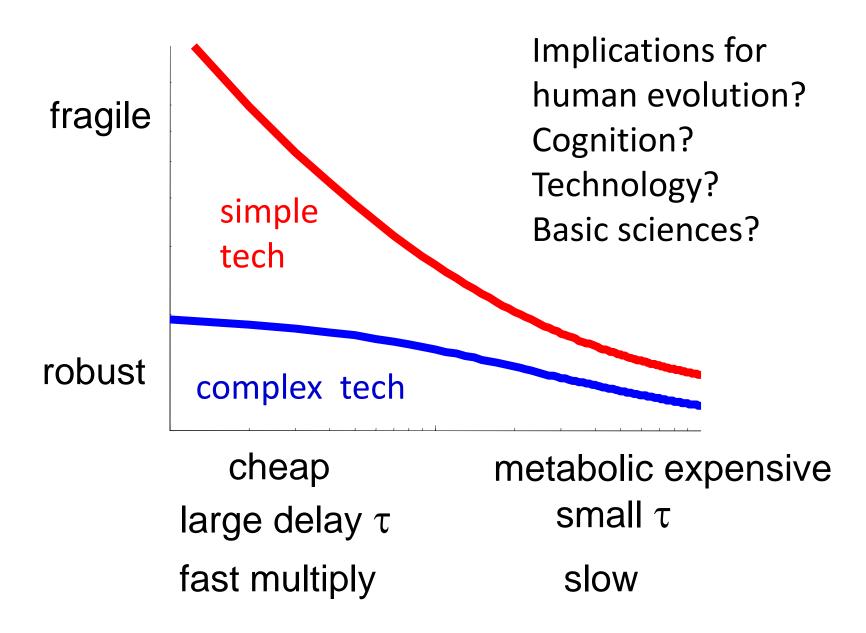




What (some) reviewers say

- "...to establish universality for all biological and physiological systems is simply wrong. It cannot be done...
- ... a mathematical scheme without any real connections to biological or medical...
- ...universality is well justified in physics... for biological and physiological systems ...a dream that will never be realized, due to the vast diversity in such systems.
- ...does not seem to understand or appreciate the vast diversity of biological and physiological systems...
- ...a high degree of abstraction, which ...make[s]
 the model useless ...

This picture is very general

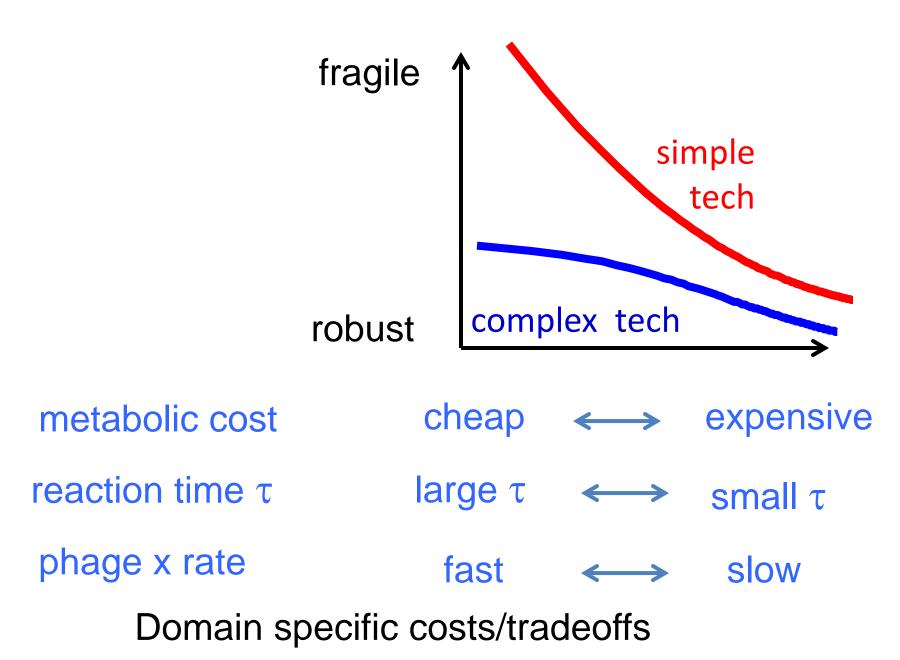


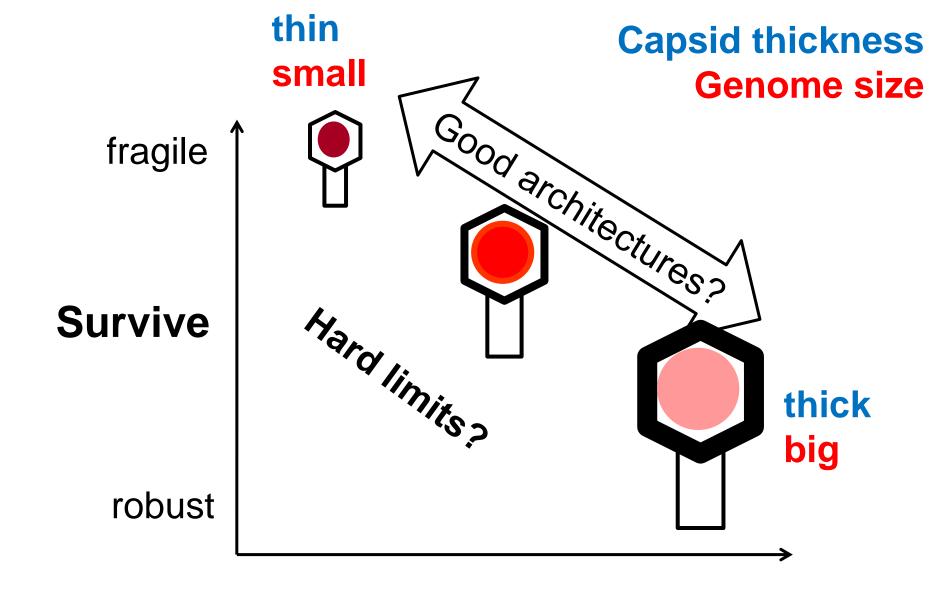
This picture is very general

Domain specific costs/tradeoffs

metabolic overhead	cheap
CNS reaction time τ (delay)	large τ ←→ small τ
phage multiplication rate	fast

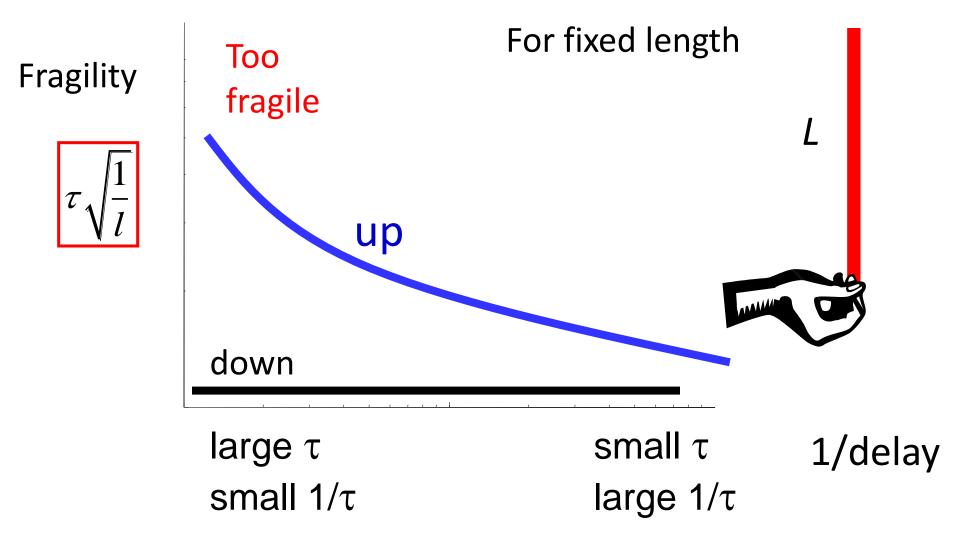
This picture is very general

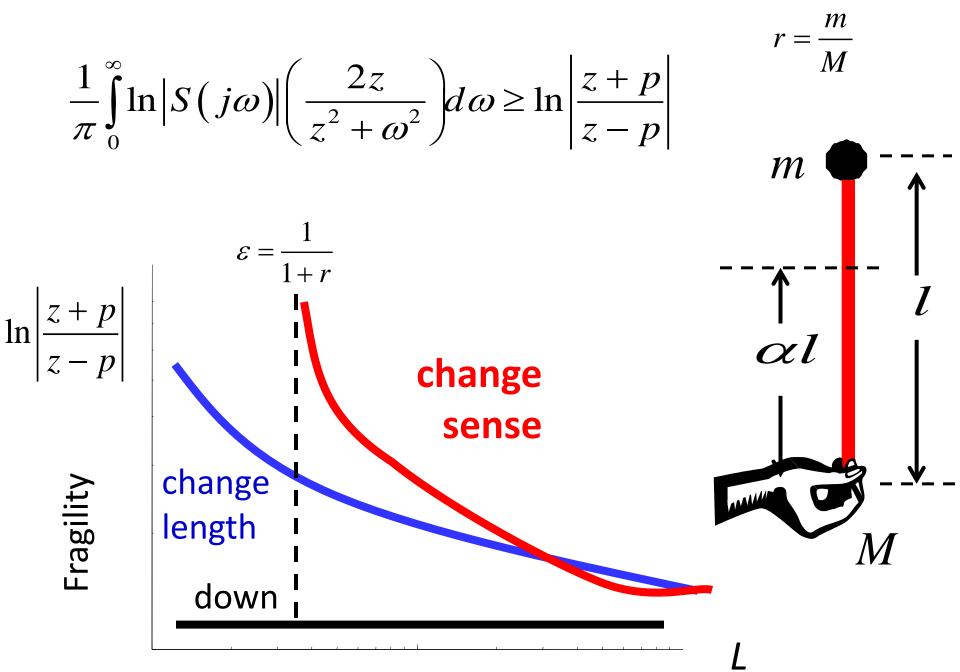




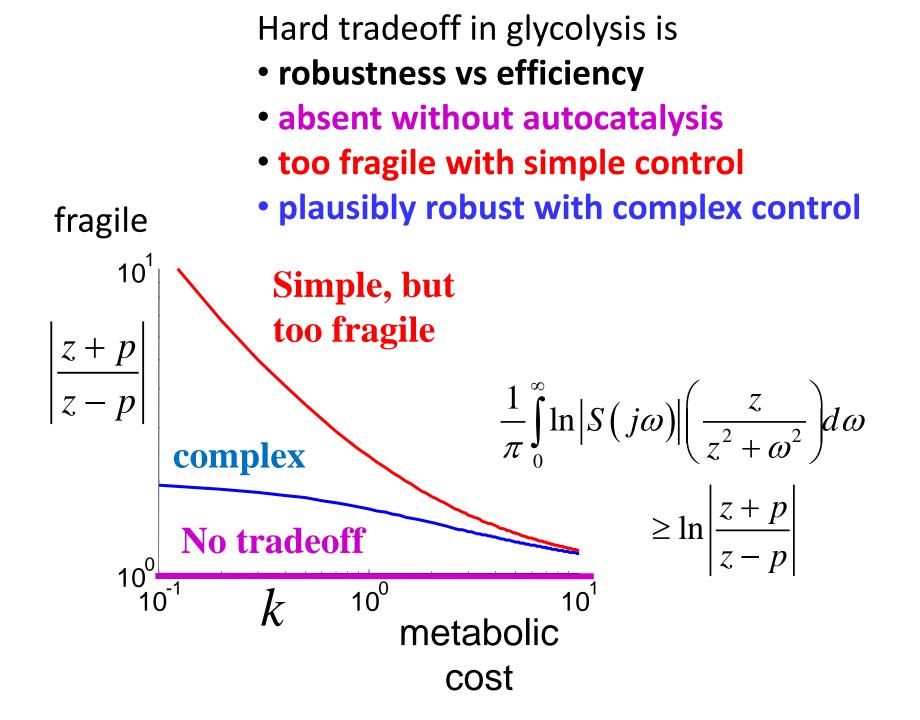
fast multiply slow

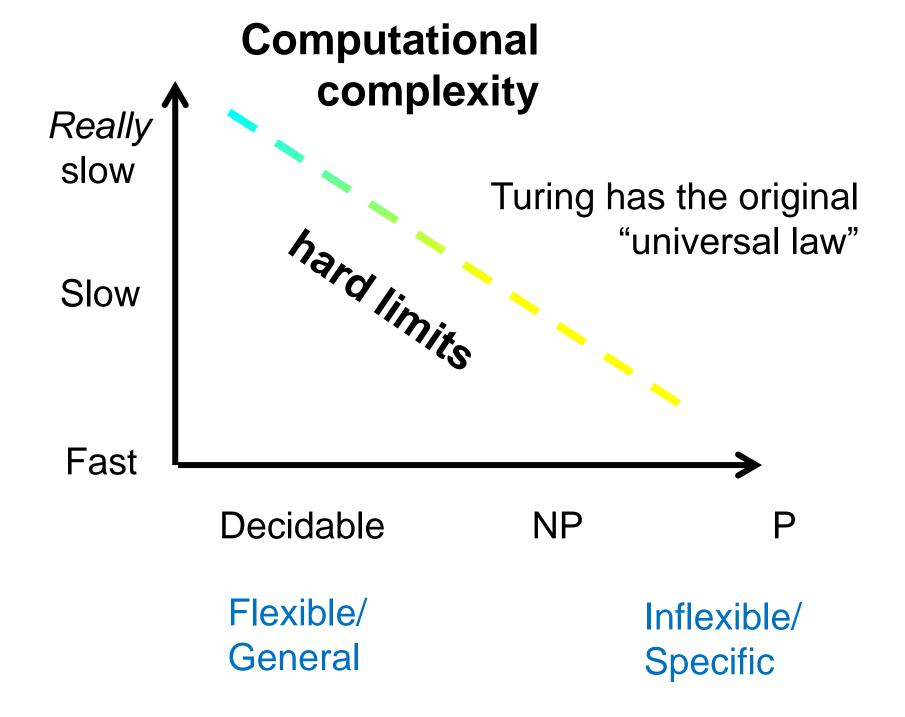
 $\frac{1}{\pi}\int_{0}^{\infty}\ln\left|T\left(j\omega\right)\right|\frac{2p}{p^{2}+\omega^{2}}d\omega \ge p\tau \propto \tau\sqrt{\frac{1}{l}}$

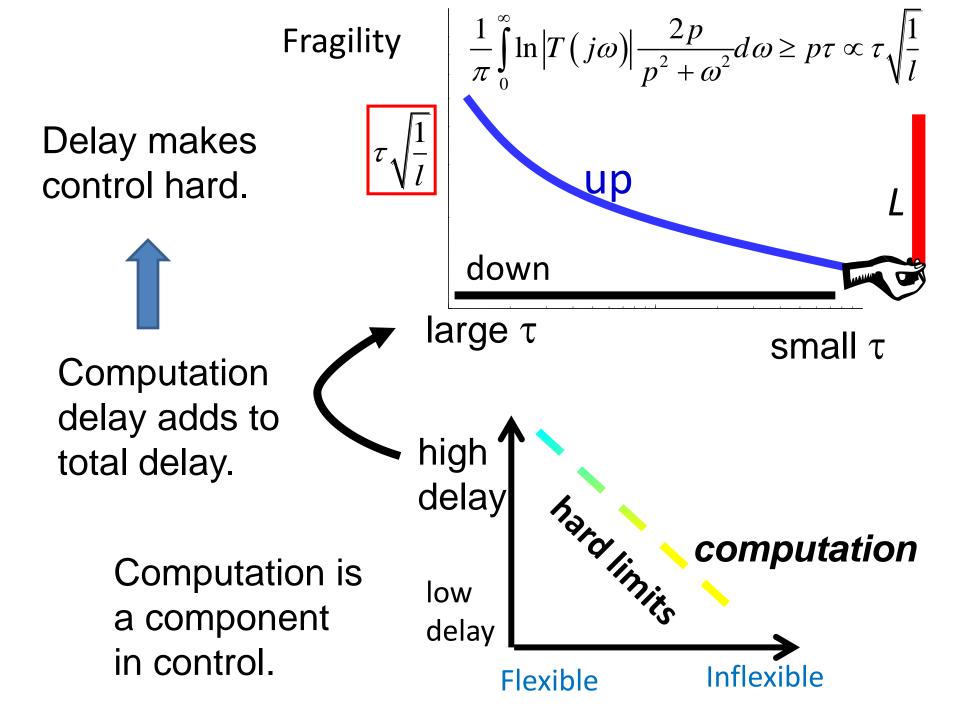


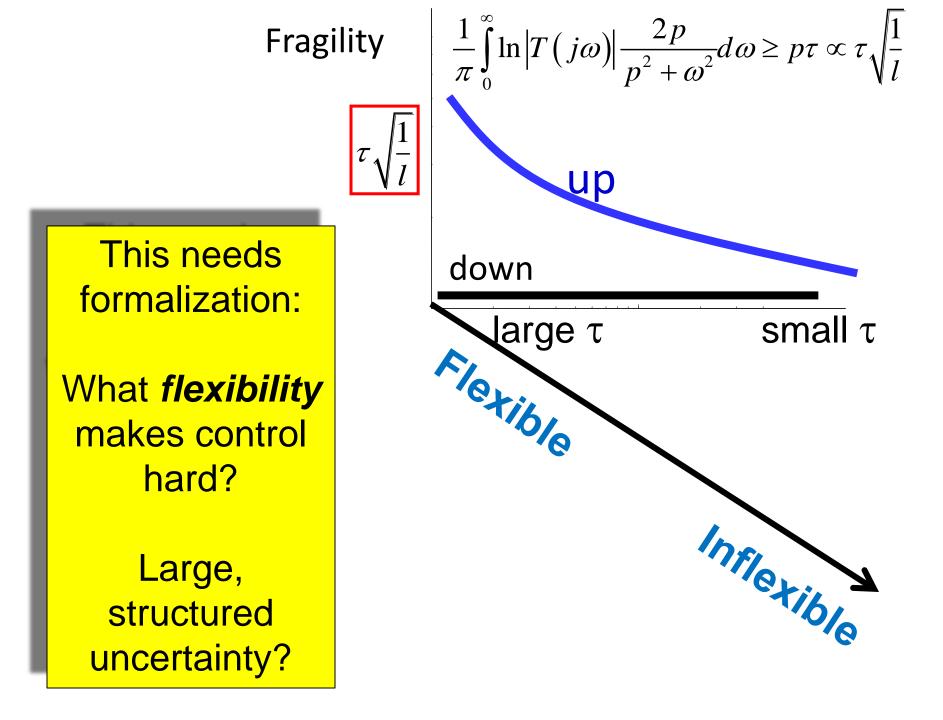


This is a cartoon, but can be made precise.

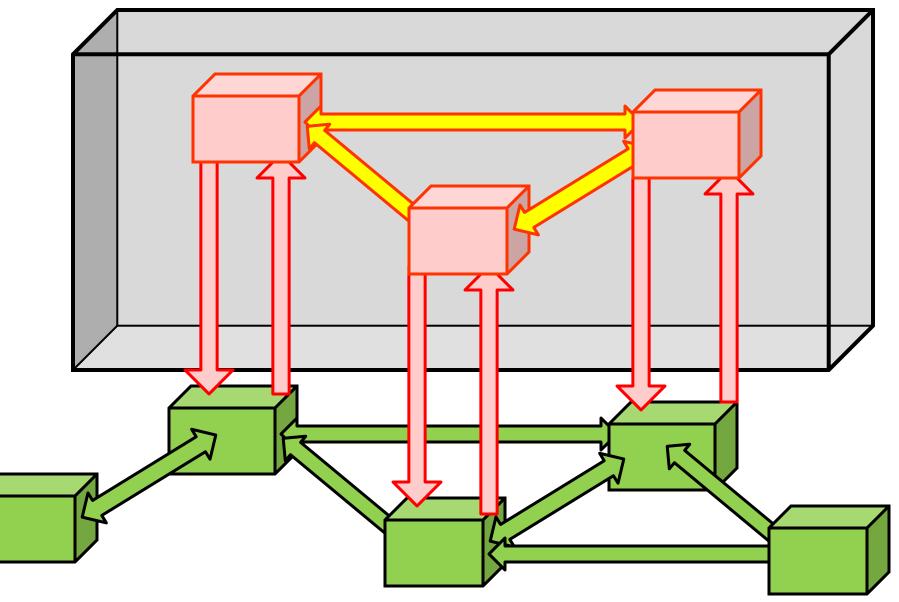


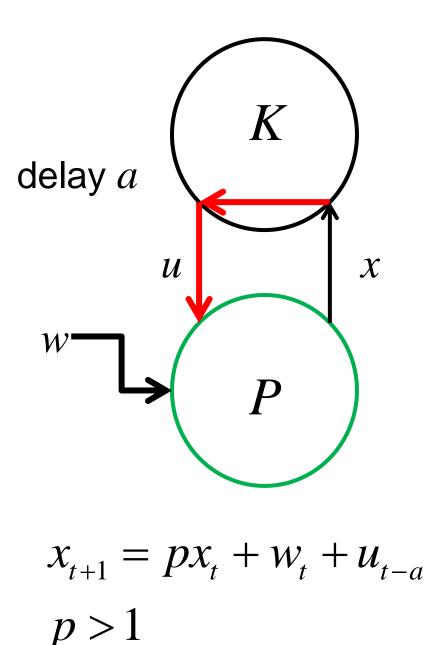






What about: Cyber-physical: decentralized control with internal delays?



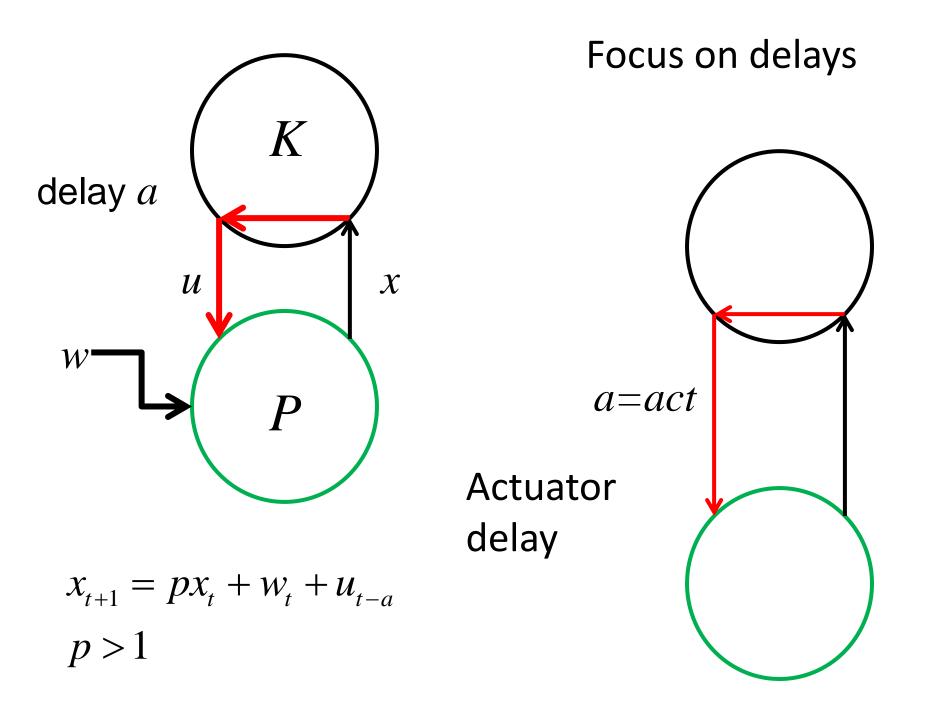


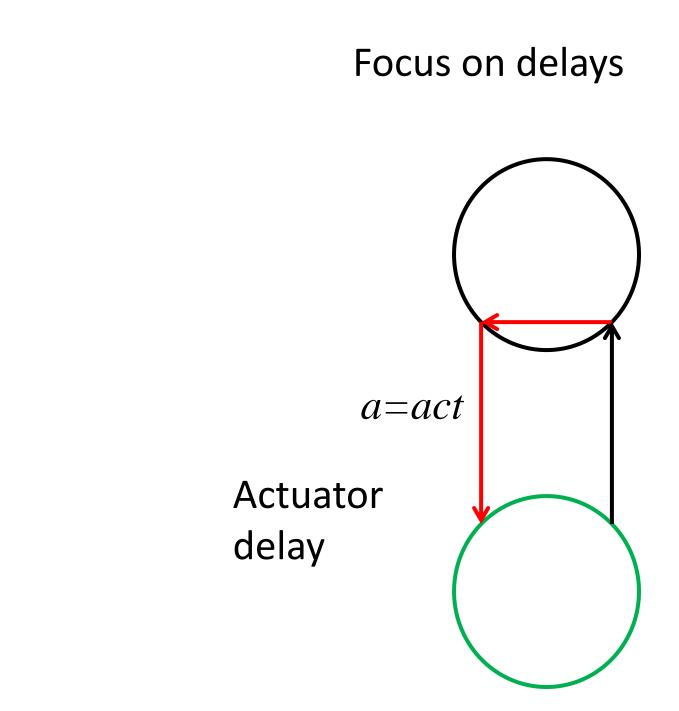
No delay or no uncertainty

$$u_{t-a} = -(px_t + w_t)$$
$$\Rightarrow ||x|| \approx 0 \quad ||u|| \approx ||w||$$

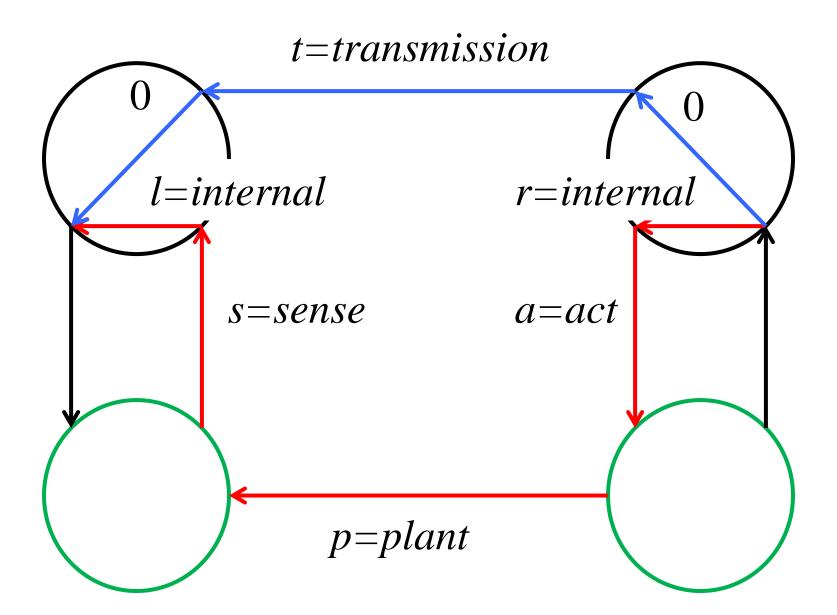
With delay **and** uncertainty

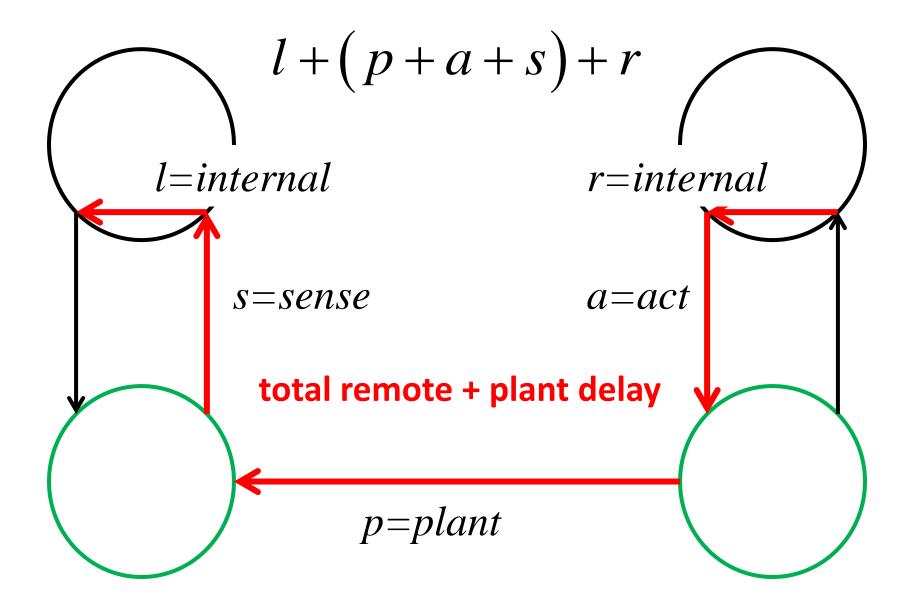
 $\Rightarrow \|x\| \approx \|u\| \approx p^a \|w\|$



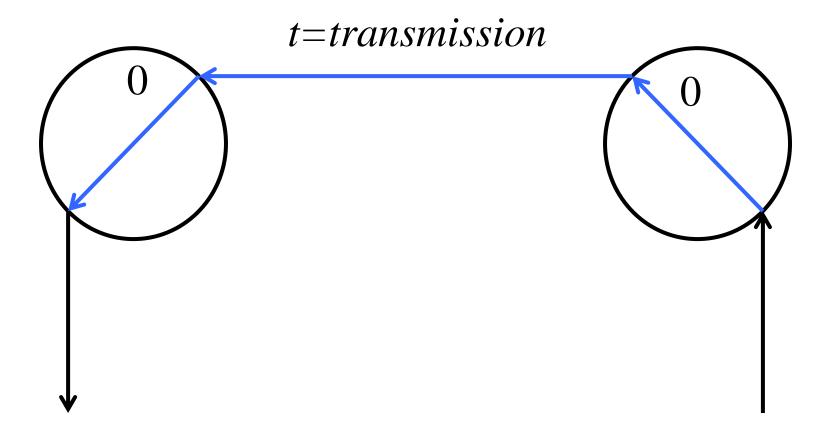


Decentralized control



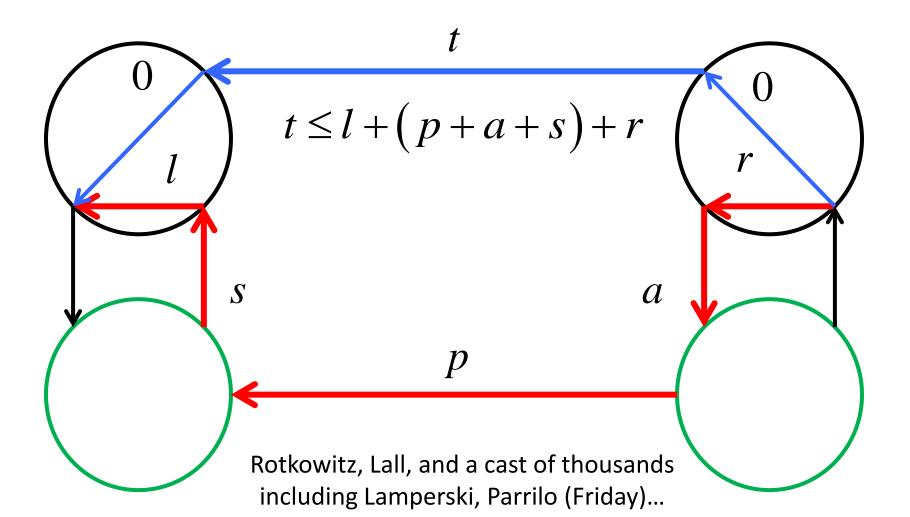


Communications delay

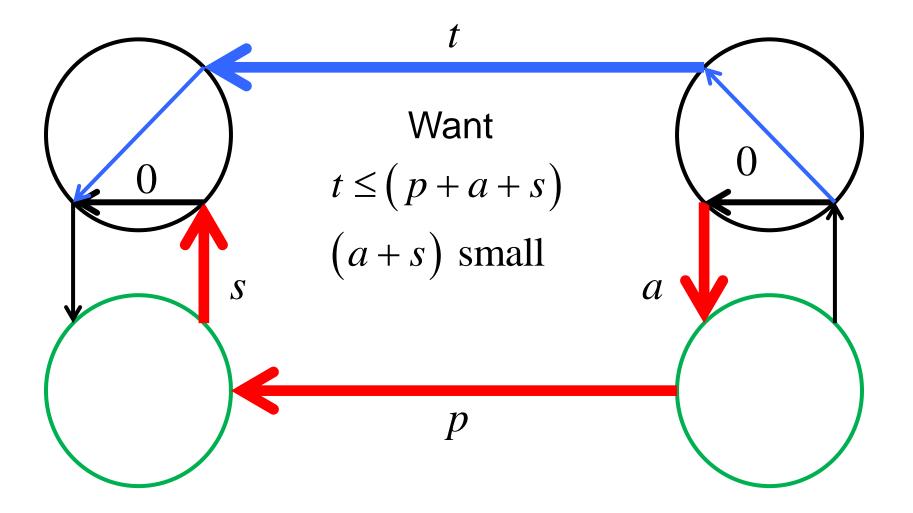


$$t \le l + (p + a + s) + r$$

Then decentralized control design can be made *convex*

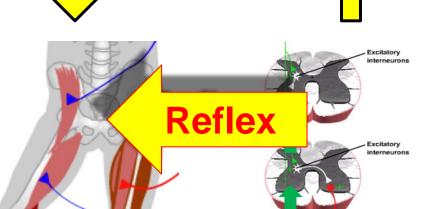


A primary driver of human brain evolution?

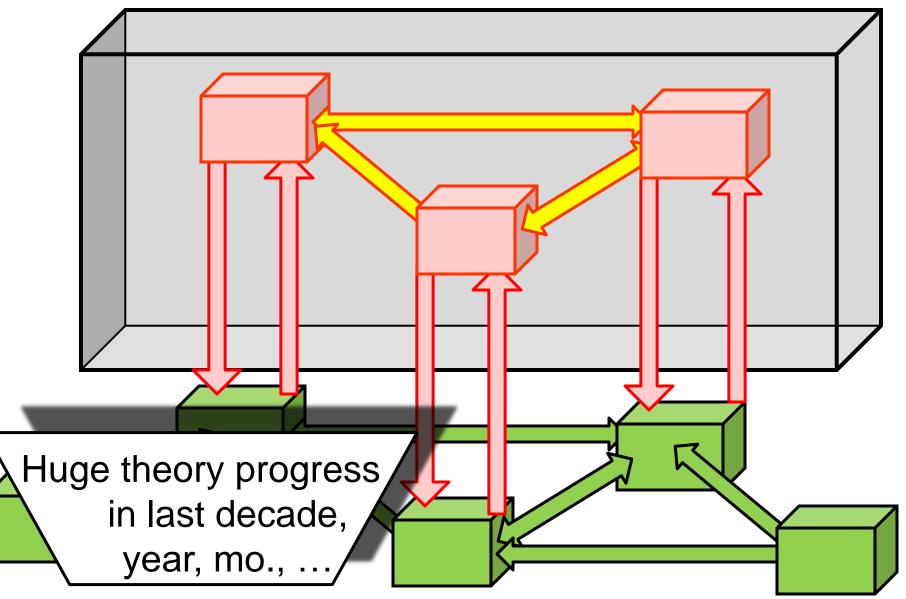


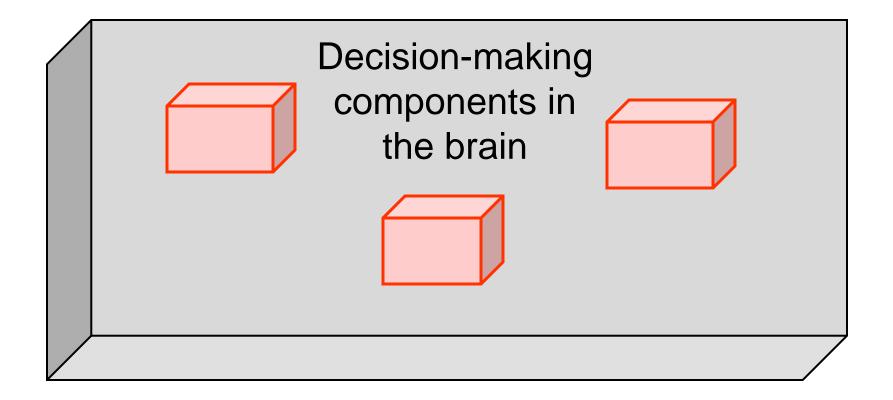


- Acquire
- Translate/ integrate
- Automate

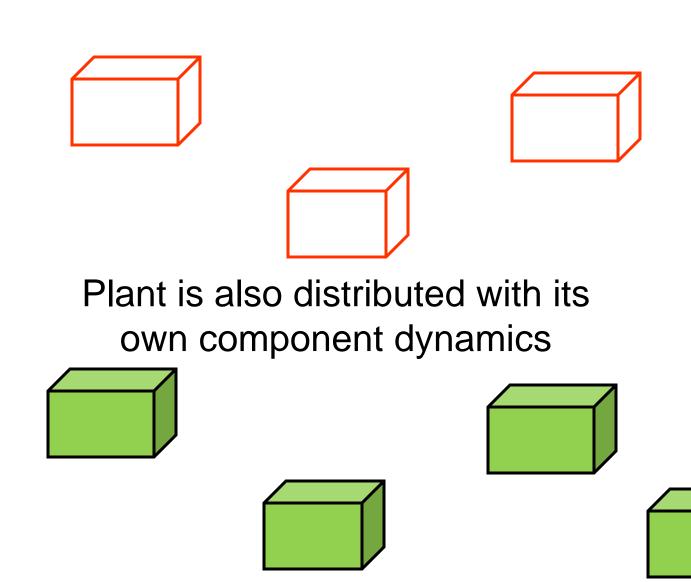


Going beyond black box: control is decentralized with internal delays.

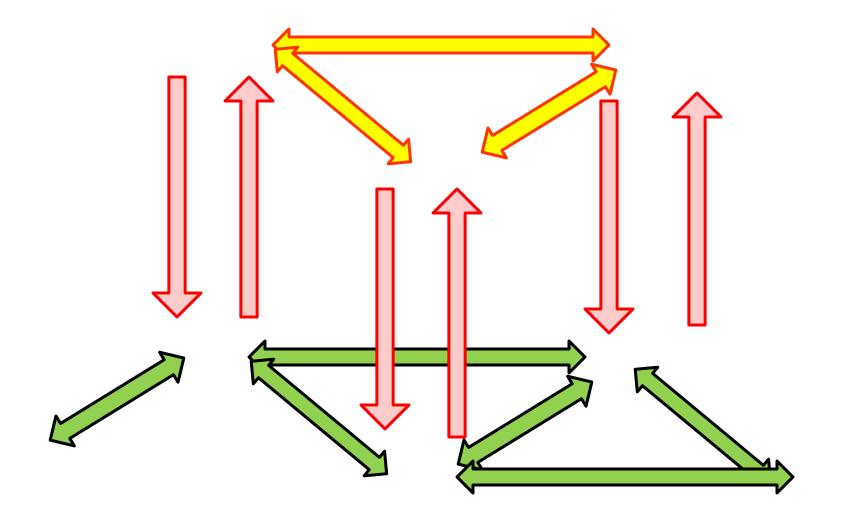




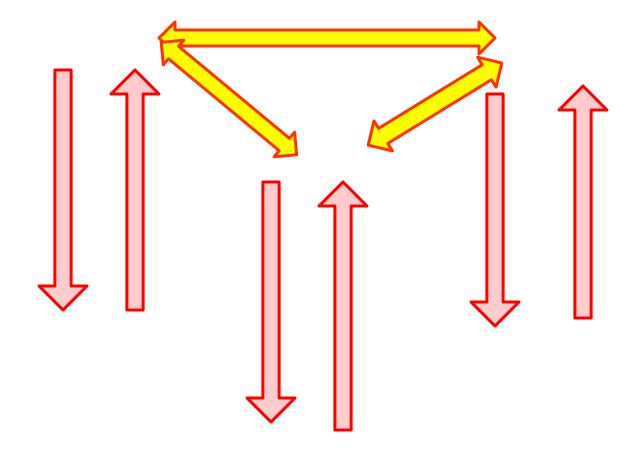
Decentralized, but initially assume computation is fast and memory is abundant.

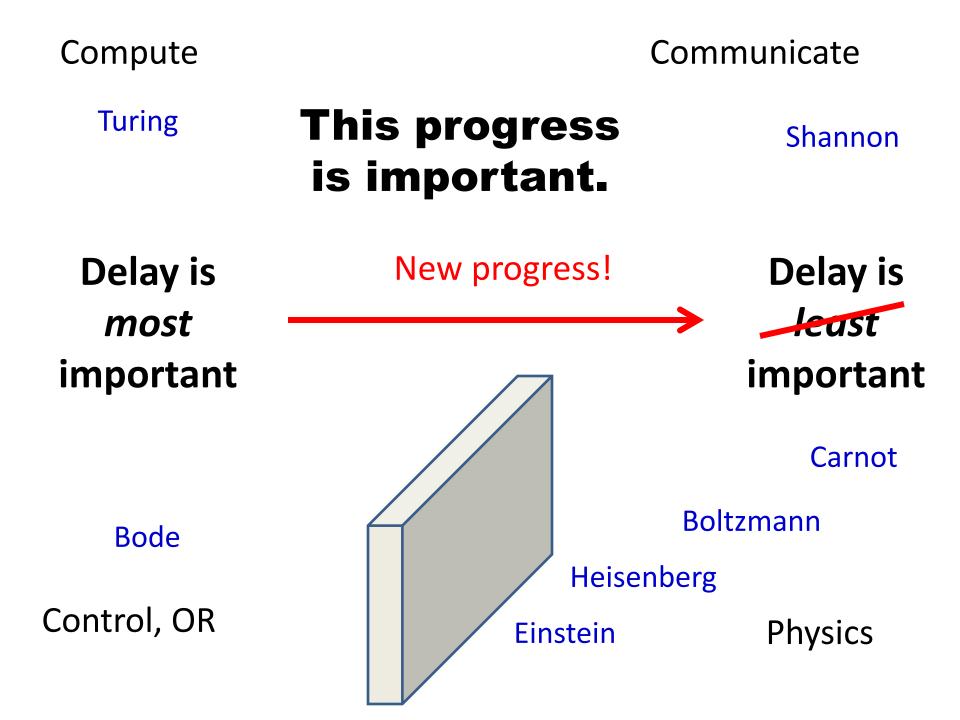


Internal delays between brain components, and their sensor and actuators, and also externally between plant components

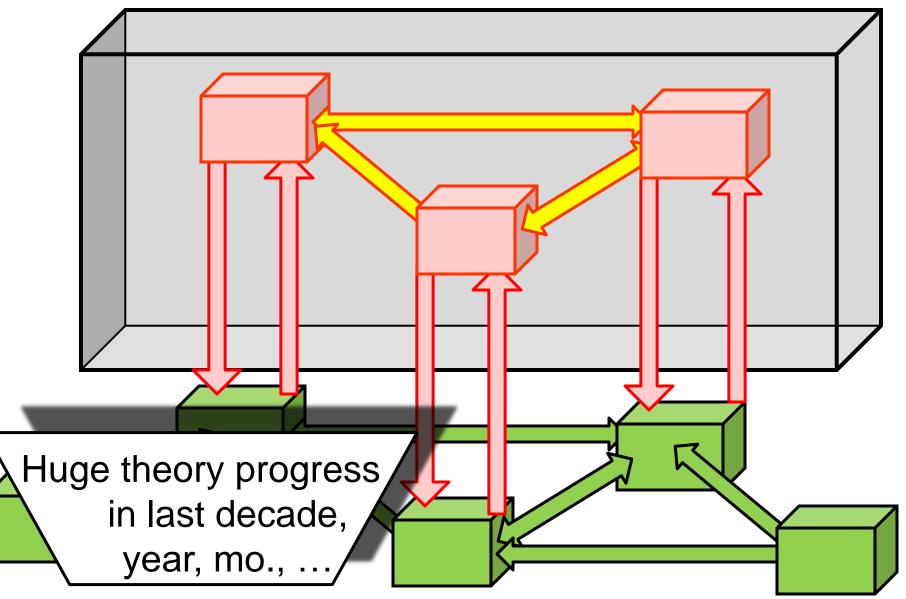


Internal delays involve both computation and communication latencies

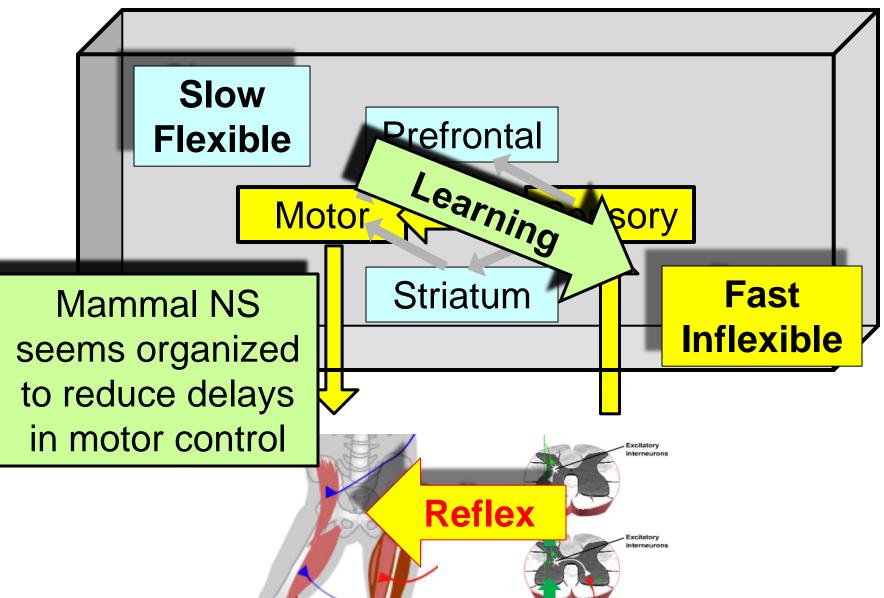




Going beyond black box: control is decentralized with internal delays.



Going beyond black box: control is decentralized with internal delays.

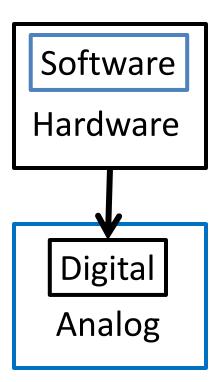


Universal architectures

Implications

(Layered architectures discussed elsewhere)

Turing as "new" starting point?

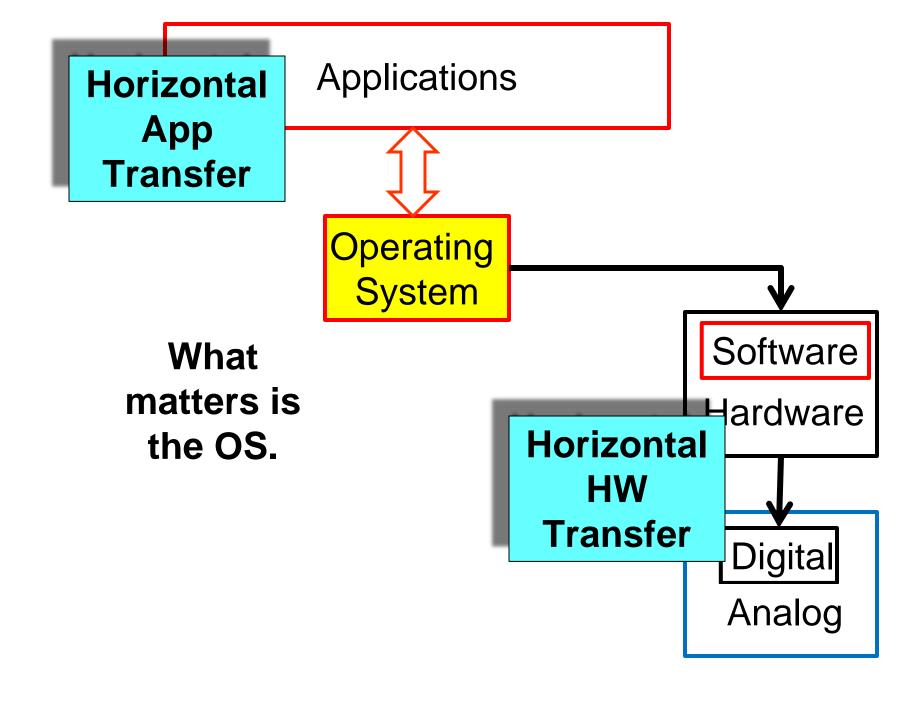


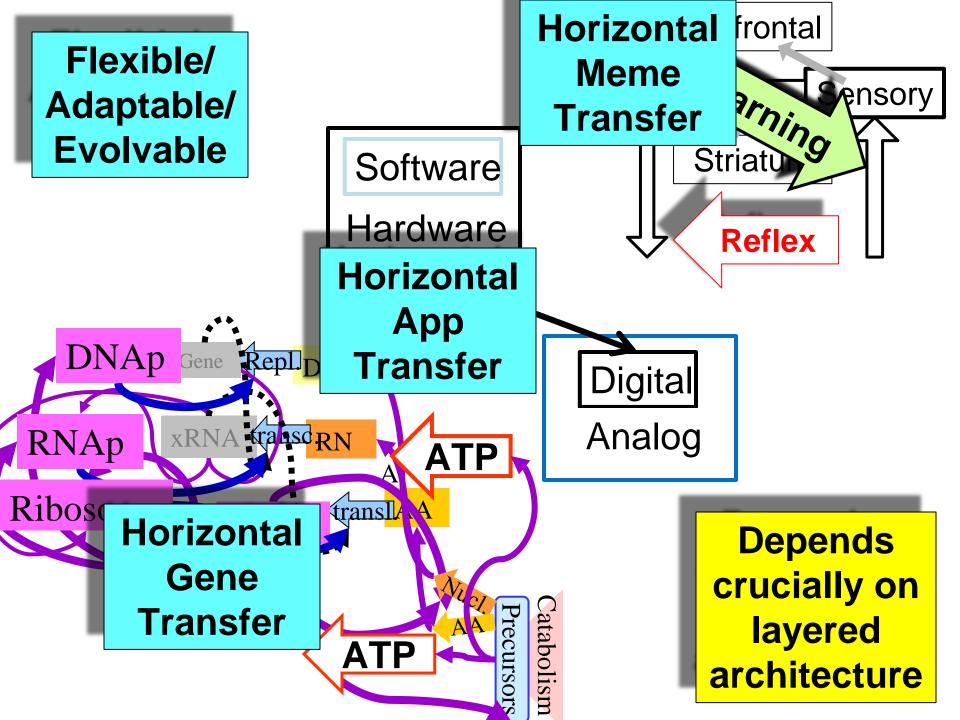
Essentials:

- 0. Model
- 1. Universal laws
- 2. Universal architecture
- 3. Practical implementation

Turing's 3 step research:

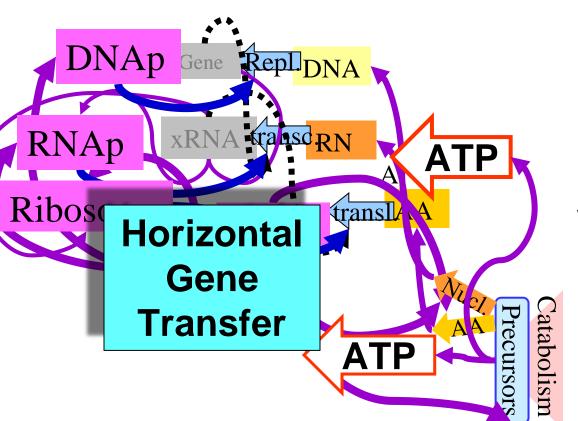
- 0. Virtual (TM) machines
- 1. hard limits, (un)decidability using standard model (TM)
- 2. Universal architecture achieving hard limits (UTM)
- 3. Practical implementation in digital electronics (biology?)



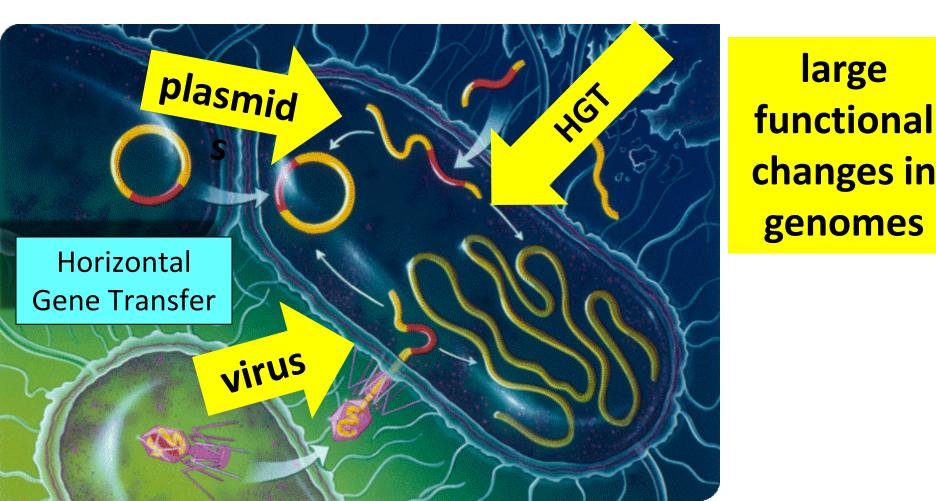


Sequence ~100 E Coli (not chosen randomly)

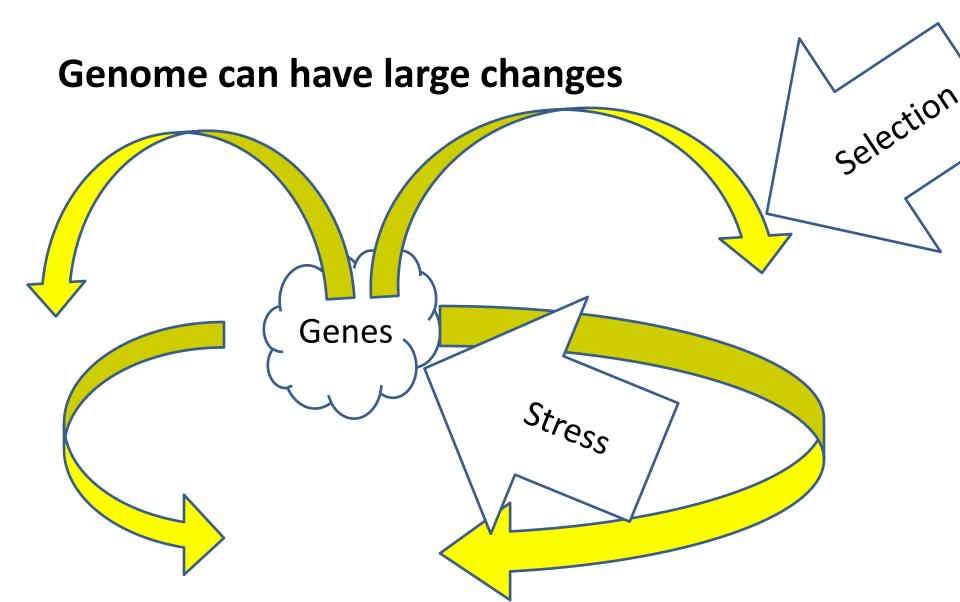
- ~ 4K genes per cell
- ~20K different genes in total
- ~ 1K universally shared genes

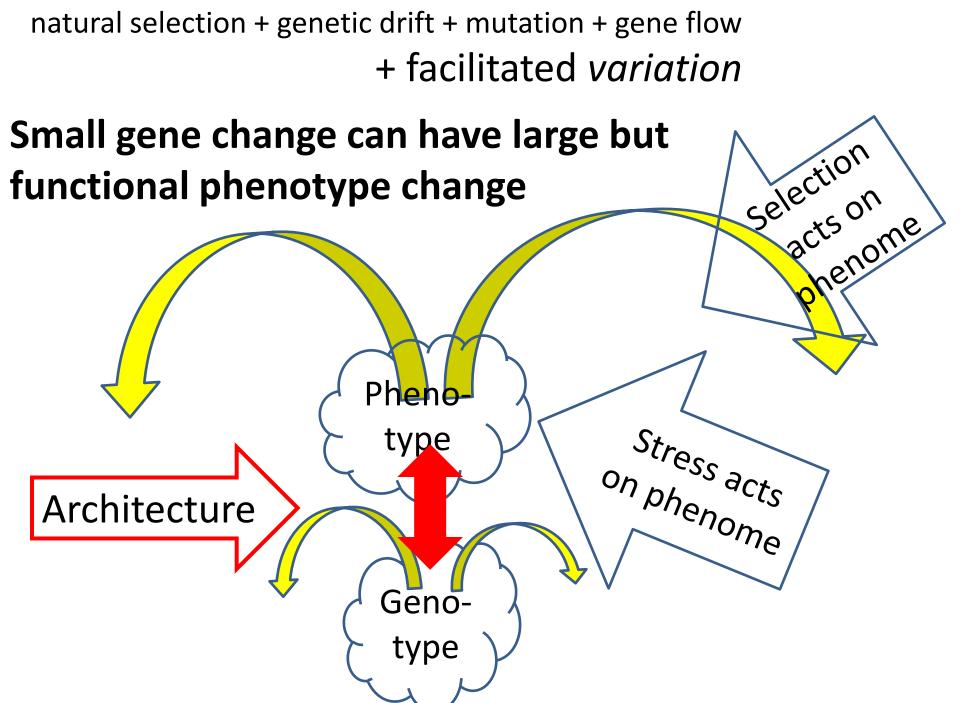


See slides on microbial biosphere laws and architectures.

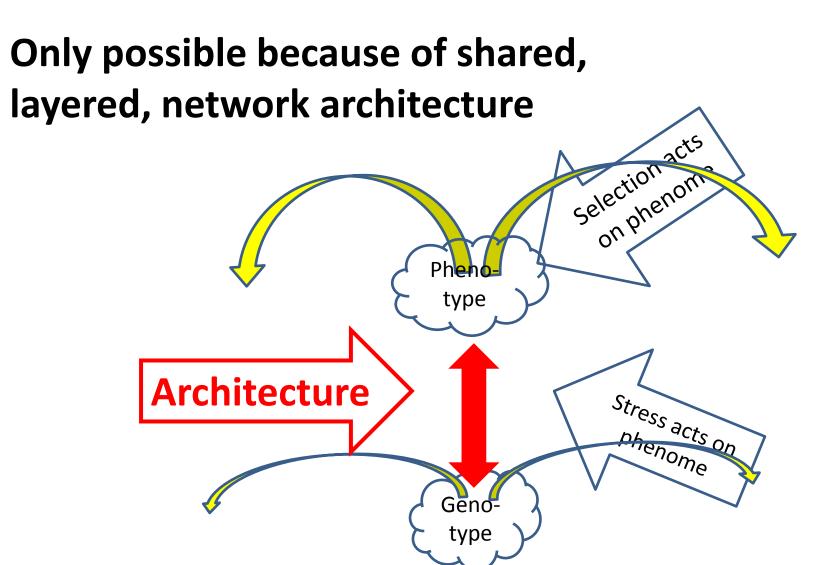


natural selection + genetic drift + mutation + gene flow + facilitated variation



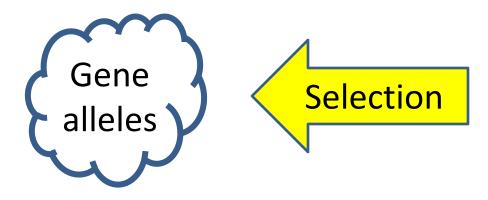


natural selection + genetic drift + mutation + gene flow + facilitated variation



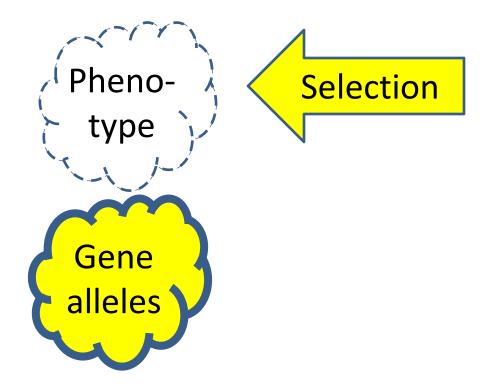
Standard theory: natural selection + genetic drift + mutation + gene flow

Greatly abridged cartoon here

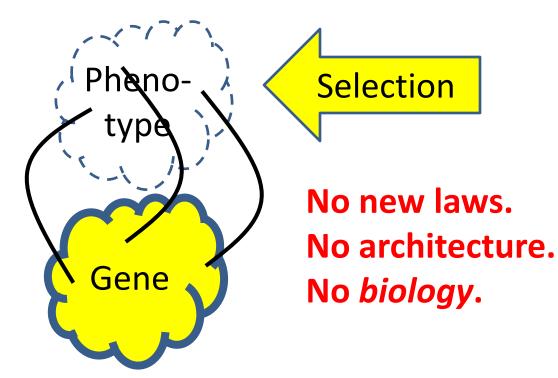


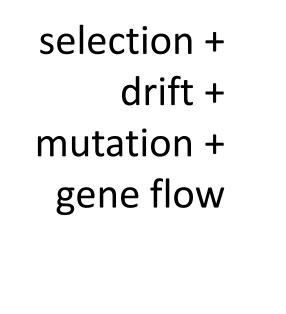
Shapiro explains well what this is and why it's incomplete (but Koonin is more mainstream)

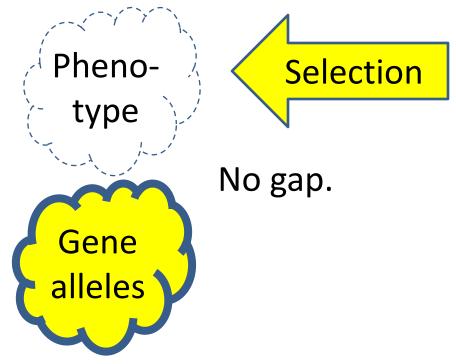
Standard theory: selection + drift + mutation + gene flow



Standard theory: selection + drift + mutation + gene flow







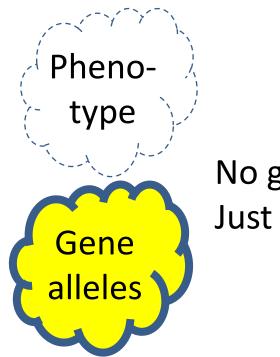
All complexity is emergent from random ensembles with minimal tuning.

No new laws.

No architecture.

The battleground





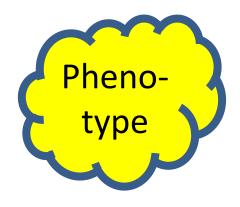
No gap. Just physics. Huge gap. Need supernatural

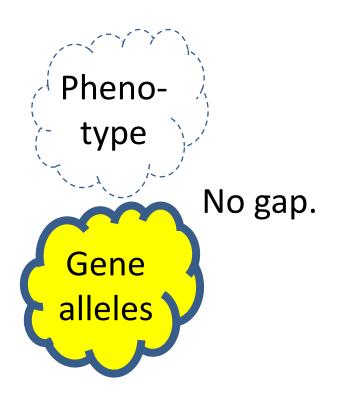


Genes?

What they agree on

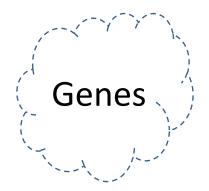
No new laws. No architecture. No biology.

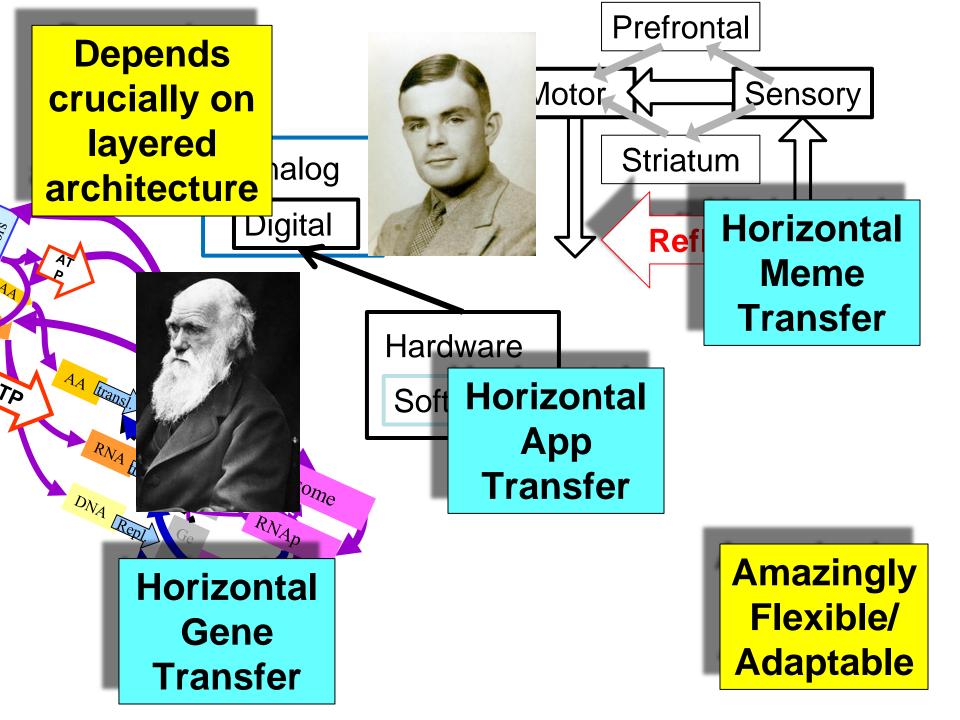




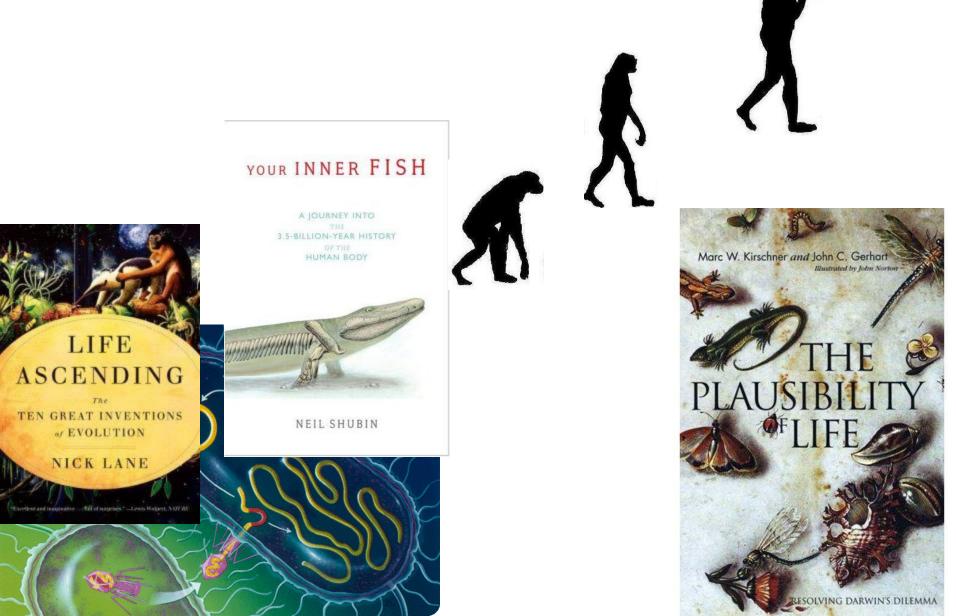
Huge gap.





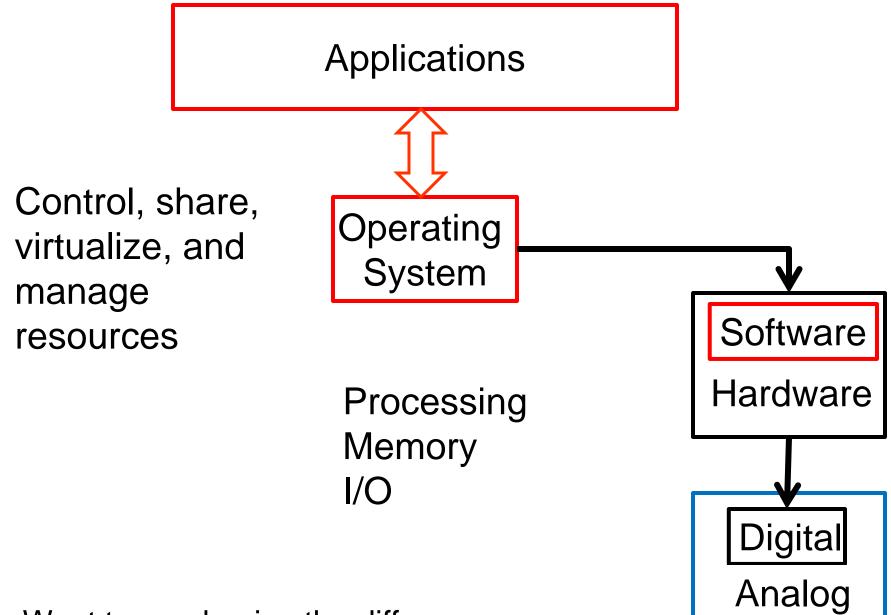


Putting biology back into evolution

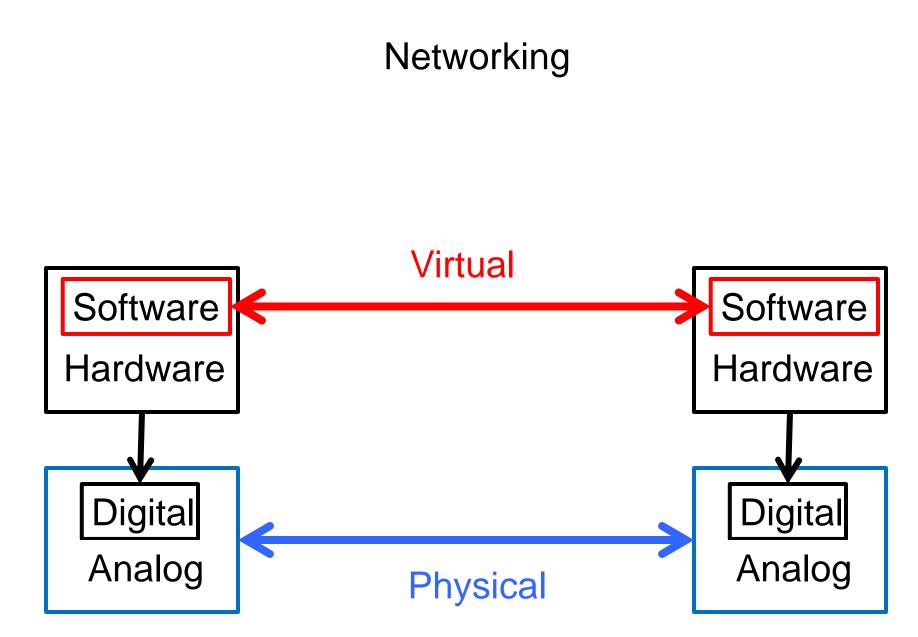


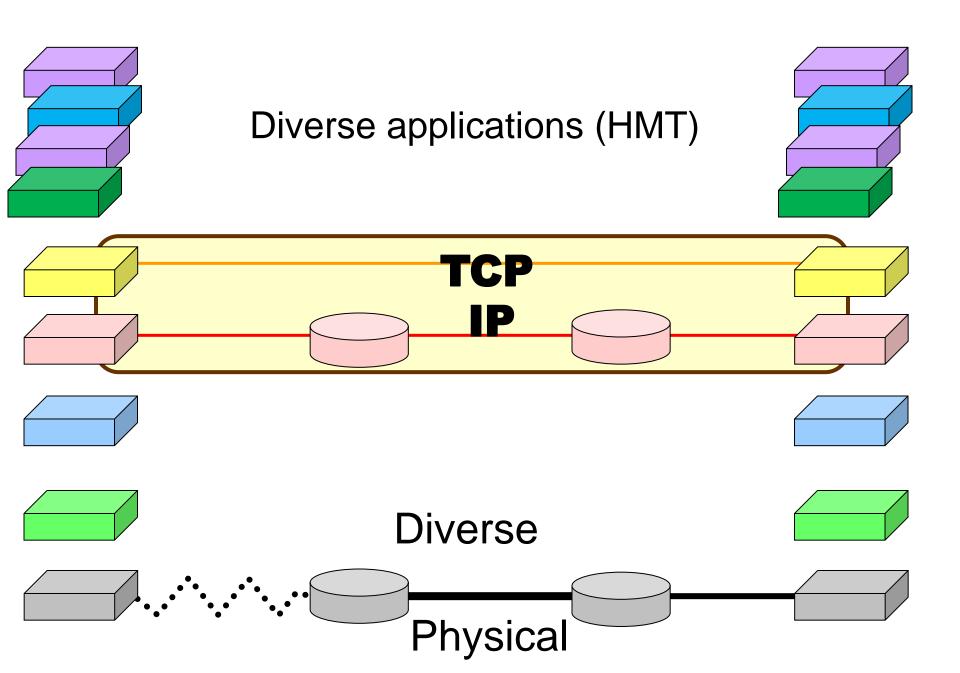
Universal architectures

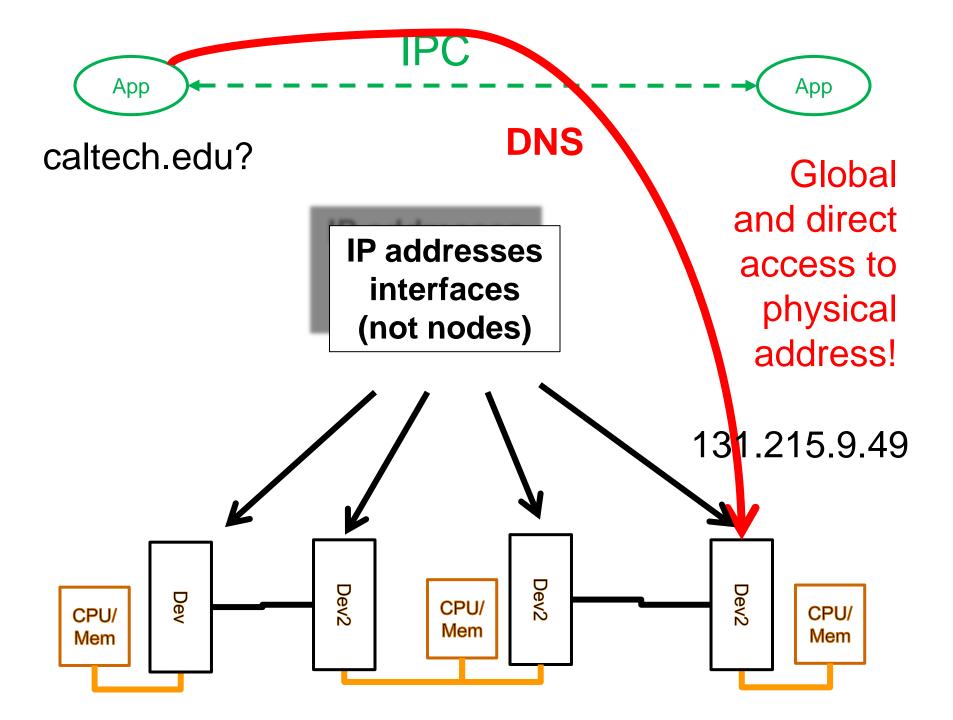
What can go wrong?

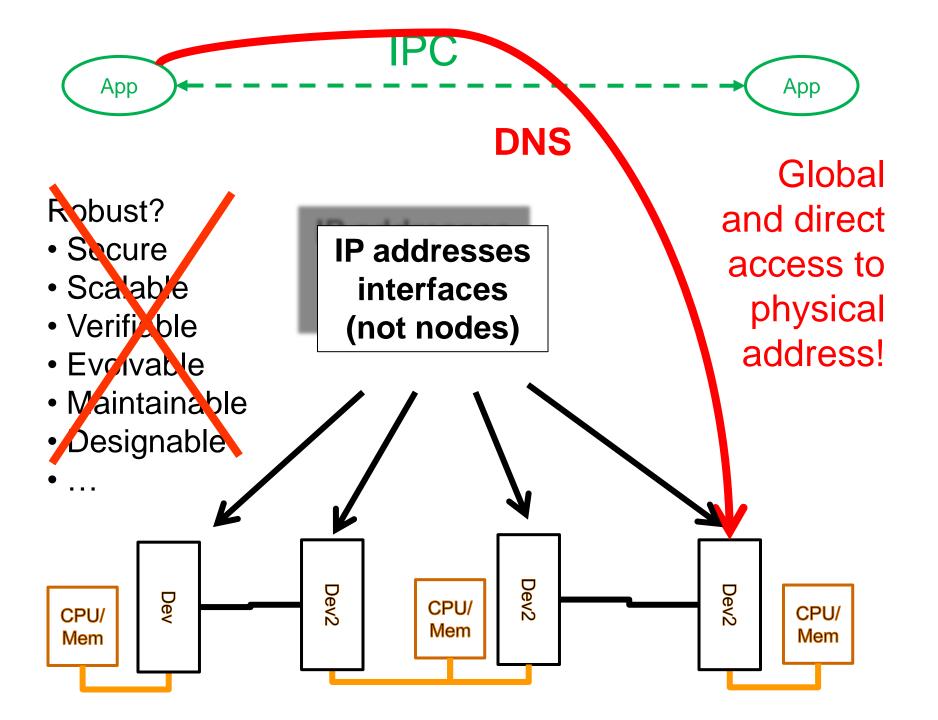


Want to emphasize the differences between these two types of layering.







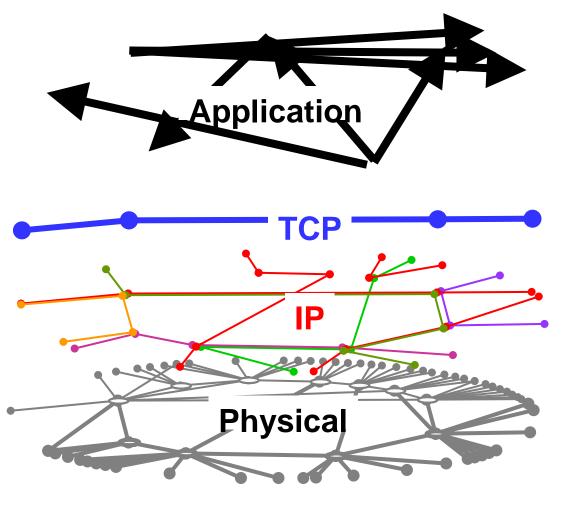


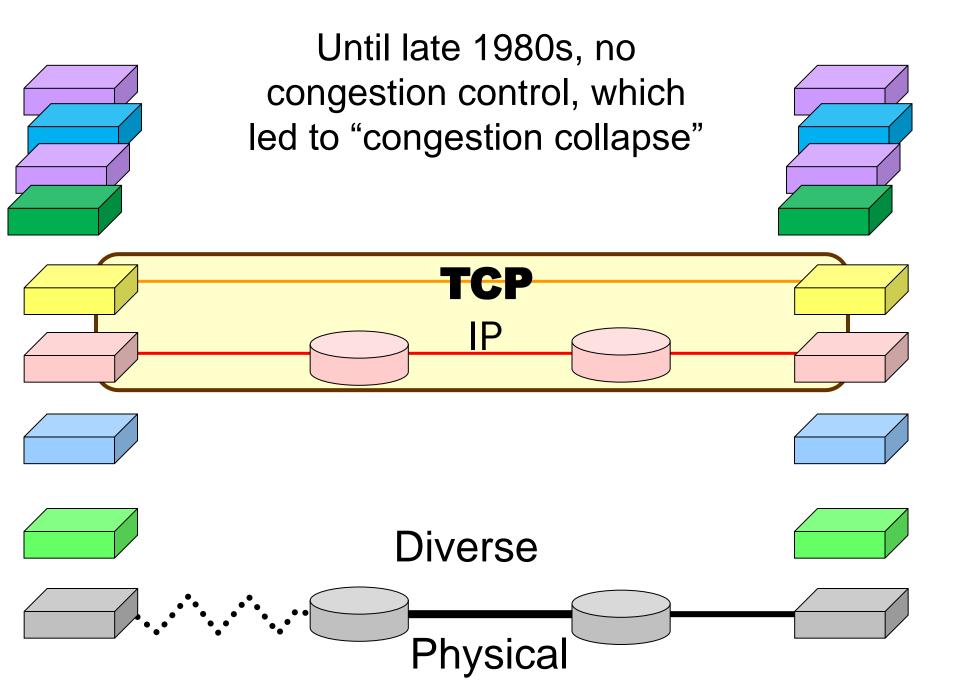
Naming and addressing need to be

- resolved within layer
- translated between layers
- not exposed outside of layer

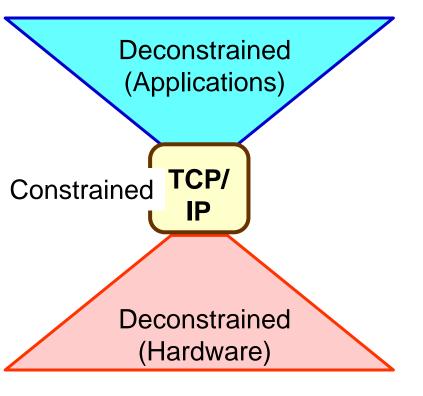
Related "issues"

- VPNs
- NATS
- Firewalls
- Multihoming
- Mobility
- Routing table size
- Overlays





Original design challenge?

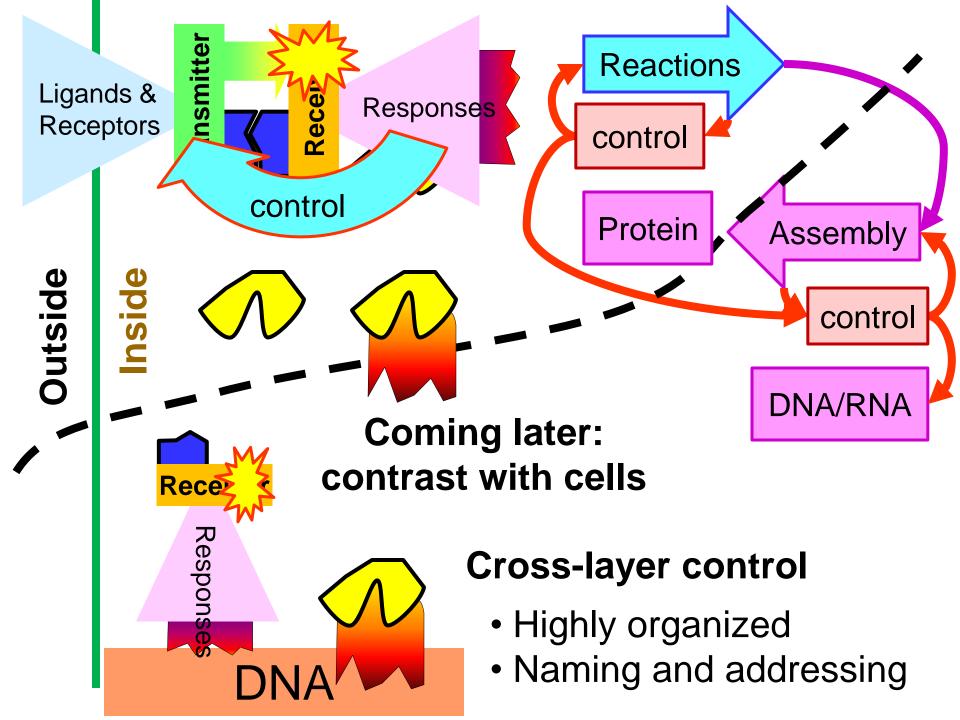


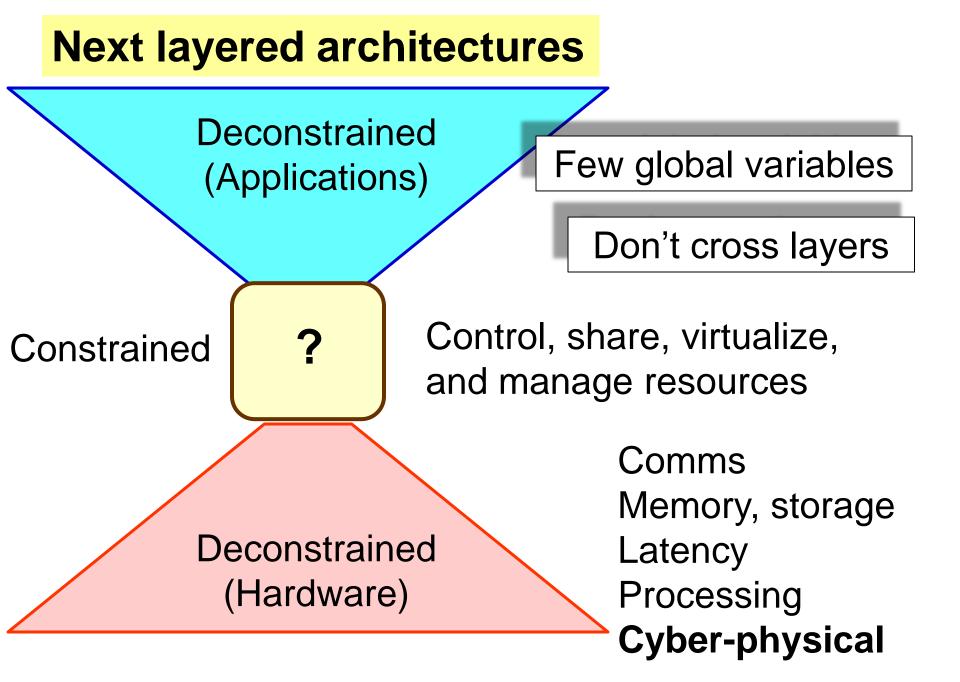
Networked OS

- Expensive mainframes
- Trusted end systems
- Homogeneous
- Sender centric
- Unreliable comms

Facilitated wild evolution Created

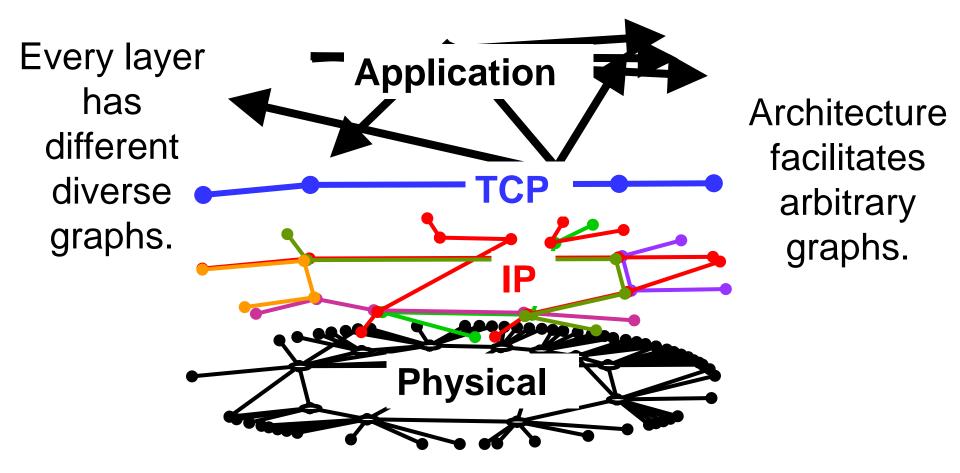
- whole new ecosystem
- completely opposite





Persistent errors and confusion ("network science")

Architecture is *least* graph topology.



The "robust yet fragile" nature of the Internet

John C. Doyle^{*†}, David L. Alderson^{*}, Lun Li^{*}, Steven Low^{*}, Matthew Roughan[‡], Stanislav Shalunov[§], Reiko Tanaka[¶], and Walter Willinger^{||}

*Engineering and Applied Sciences Division, California Institute of Technology, Pasadena, CA 91125; [‡]Applied Mathematics, University of Adelaide, South Australia 5005, Australia; [§]Internet2, 3025 Boardwalk Drive, Suite 200, Ann Arbor, MI 48108; [¶]Bio-Mimetic Control Research Center, Institute of Physical and Chemical Research, Nagoya 463-0003, Japan; and [¶]AT&T Labs–Research, Florham Park, NJ 07932

Edited by Robert M. May, University of Oxford, Oxford, United Kingdom, and approved August 29, 2005 (received for review February 18, 2005)

The search for unifying properties of complex networks is popular, challenging, and important. For modeling approaches that focus on

SVNJ

no self-loops or parallel edges) having the same graph degree We will say that graphs $g \in G(D)$ have scaling-degree sequen

PNAS October 11, 2005 vol. 102 no. 41 14497–14502

Notices of the AMS, 2009

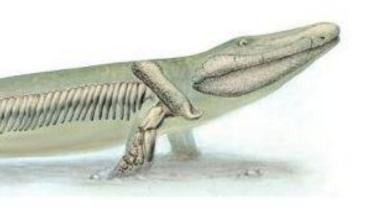
Mathematics and the Internet: A Source of Enormous Confusion and Great Potential

Walter Willinger, David Alderson, and John C. Doyle

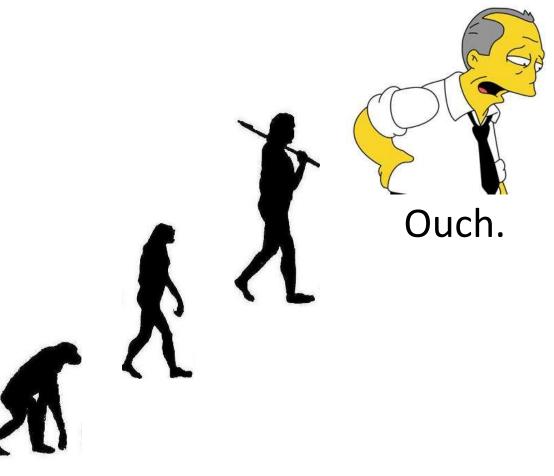
Unfortunately, not intelligent design

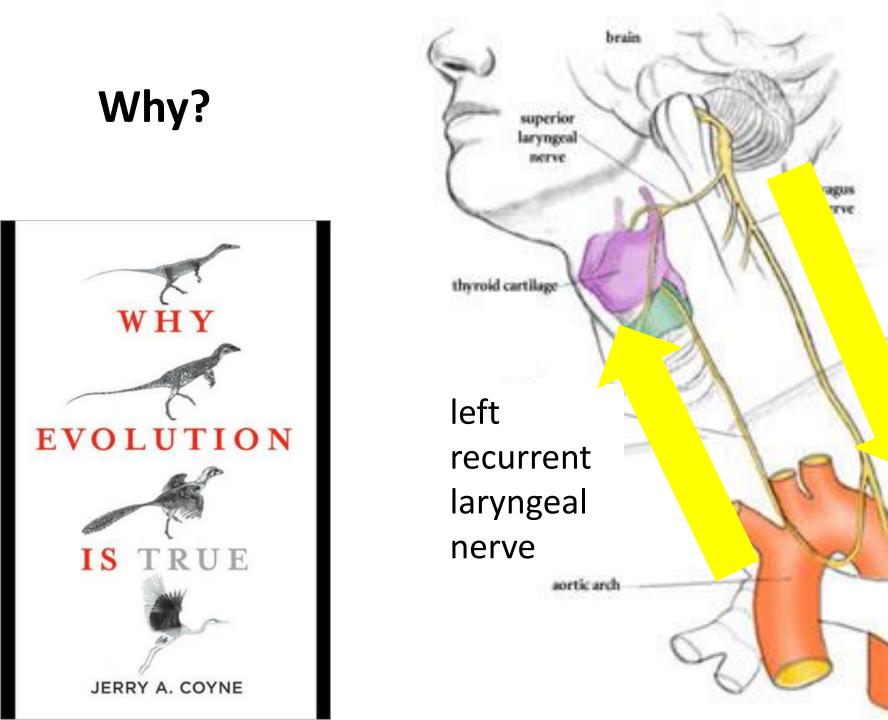
YOUR INNER FISH

A JOURNEY INTO THE 3.5-BILLION-YEAR HISTORY OF THE HUMAN BODY



NEIL SHUBIN





Why? Building humans from fish parts.

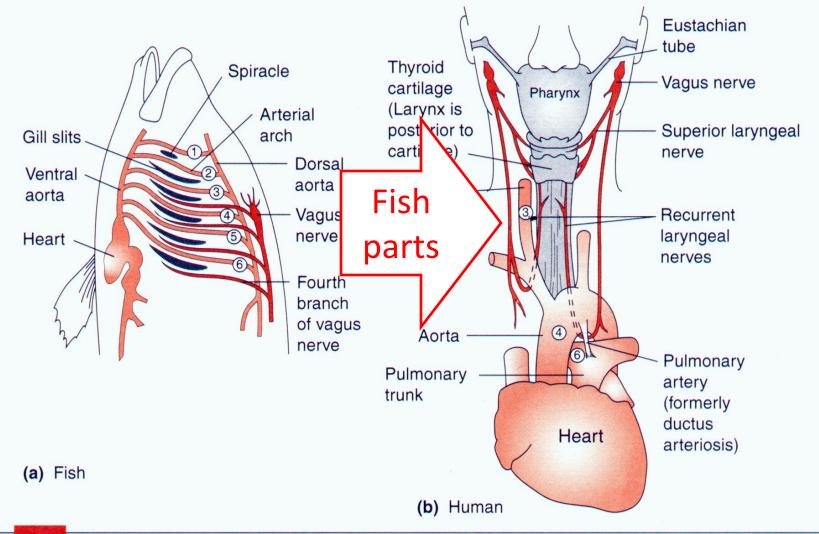
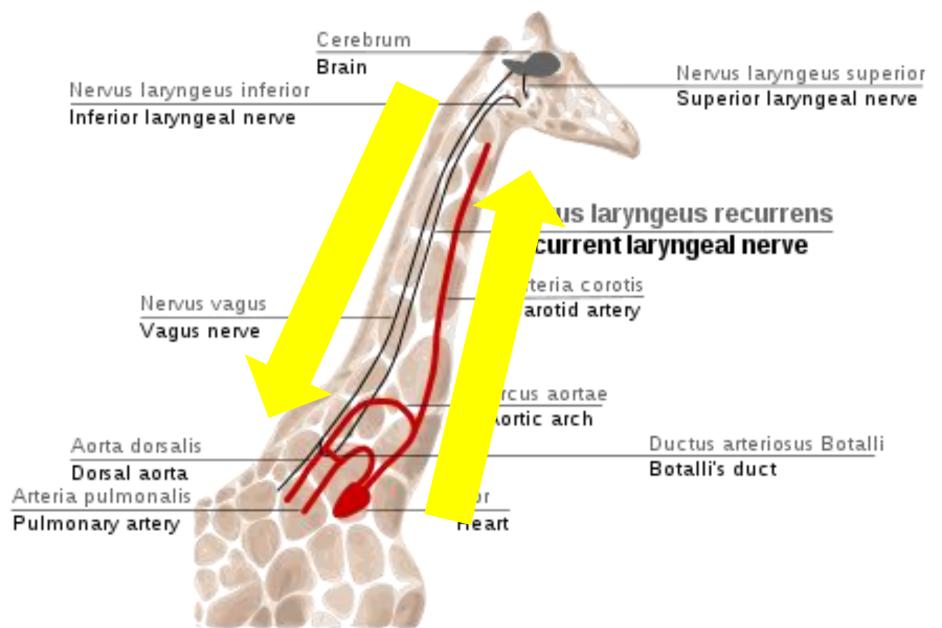
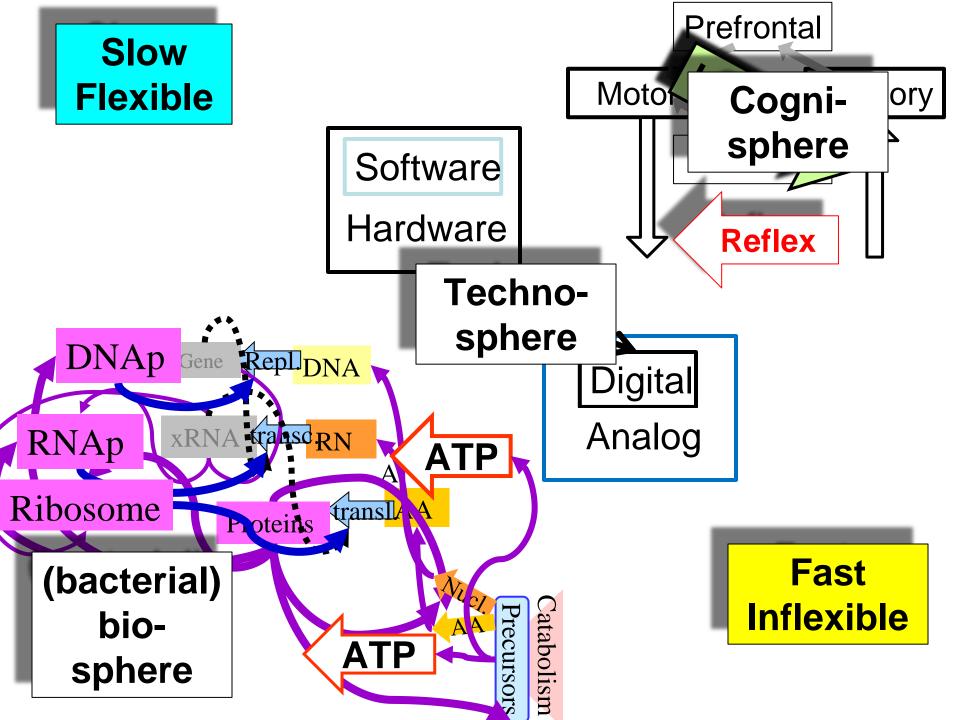
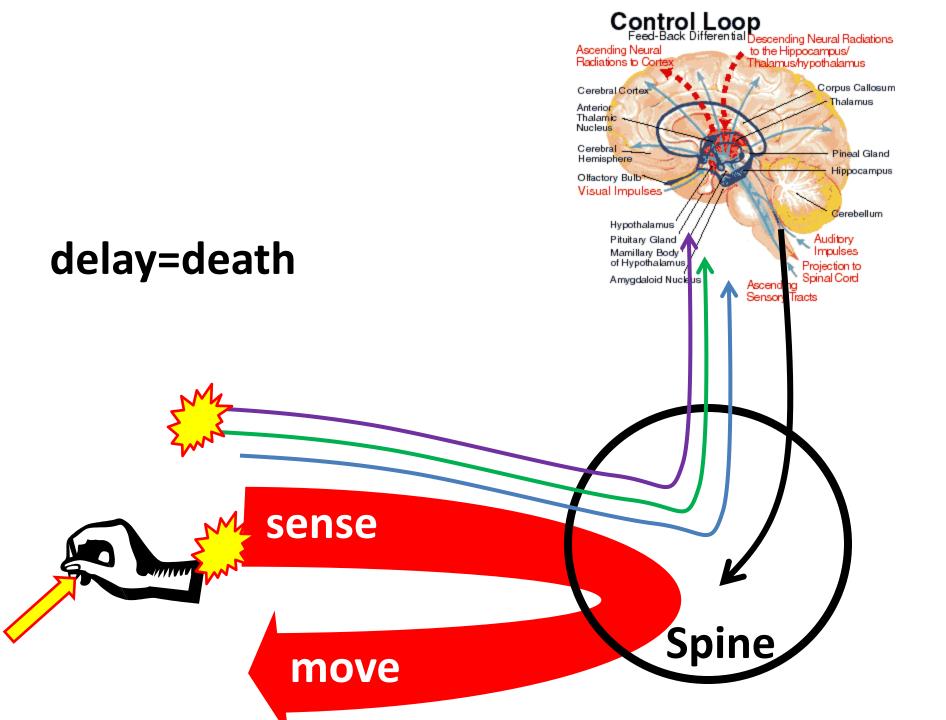


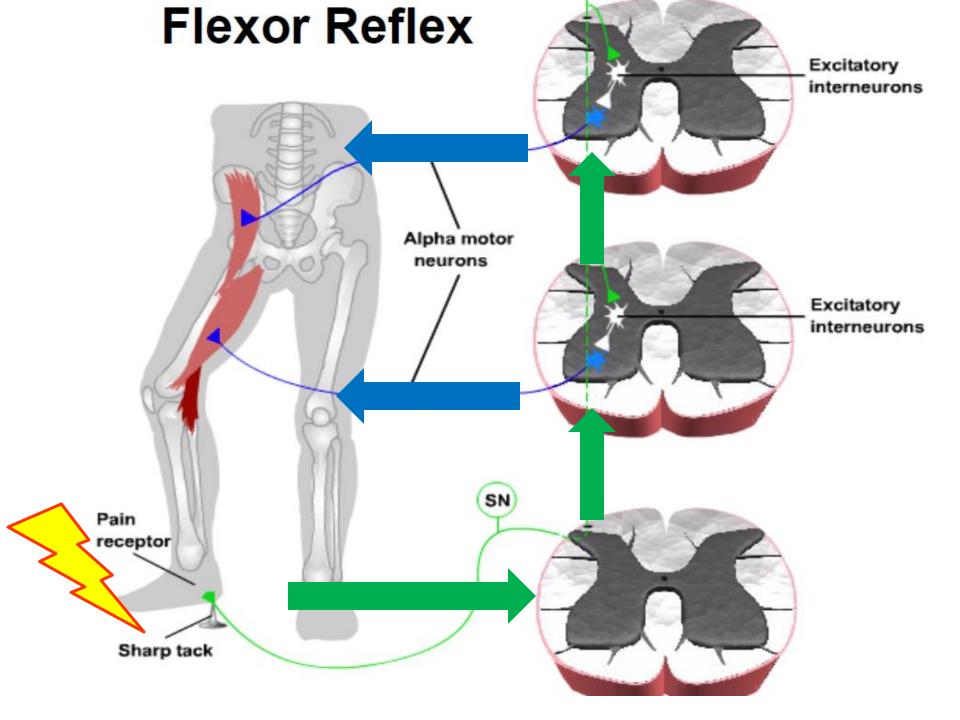
FIGURE 3–11 Schematic diagram showing the relationship between the vagus cranial nerve and the arterial arches in fish (a) and human (b). Only the third, fourth, and part of the sixth arterial arches remain in placental mammals, the sixth acting only during fetal development to carry blood to the placenta. The fourth vagal nerve in mammals (the recurrent laryngeal nerve) loops around the sixth arterial arch just as it did in the original fishlike ancestor, but must now travel a greater distance since the remnant of the sixth arch is in the thorax.

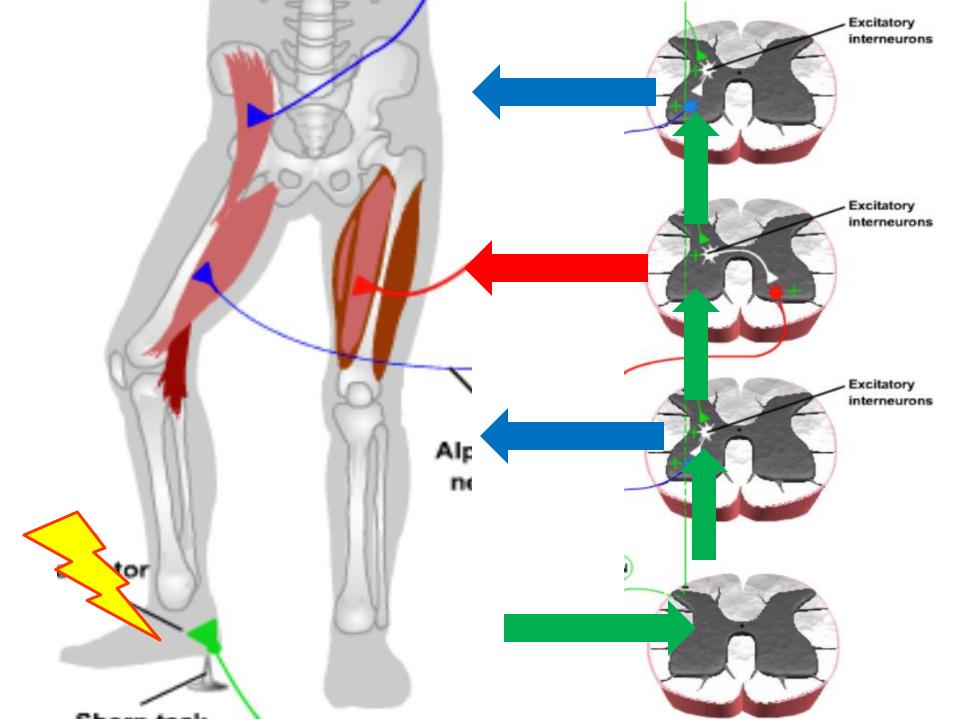
It could be worse.

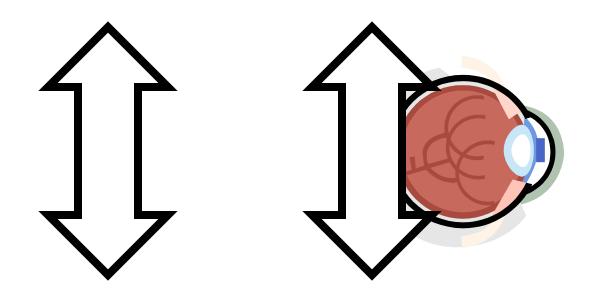


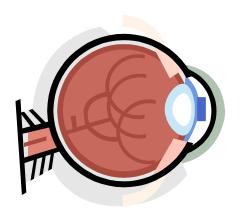


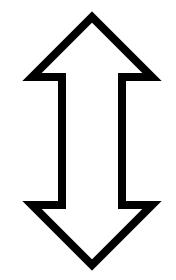


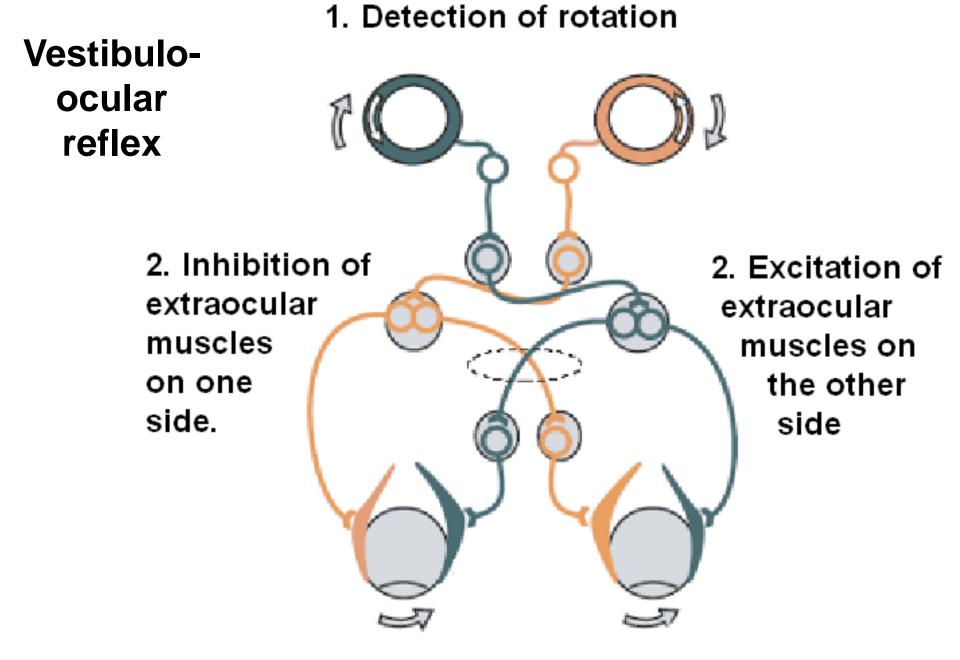




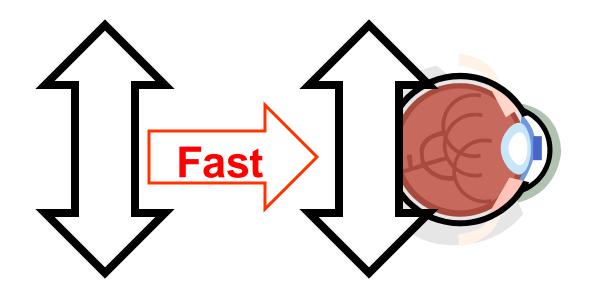


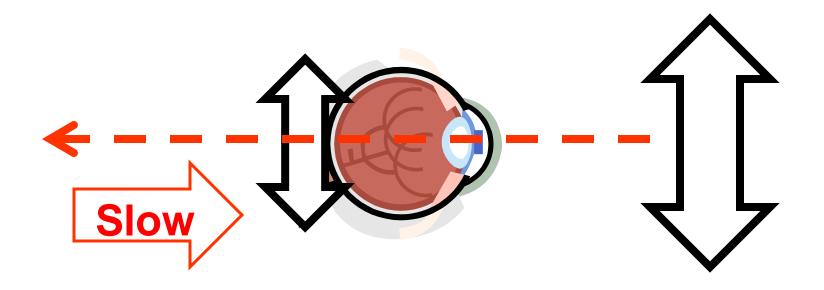


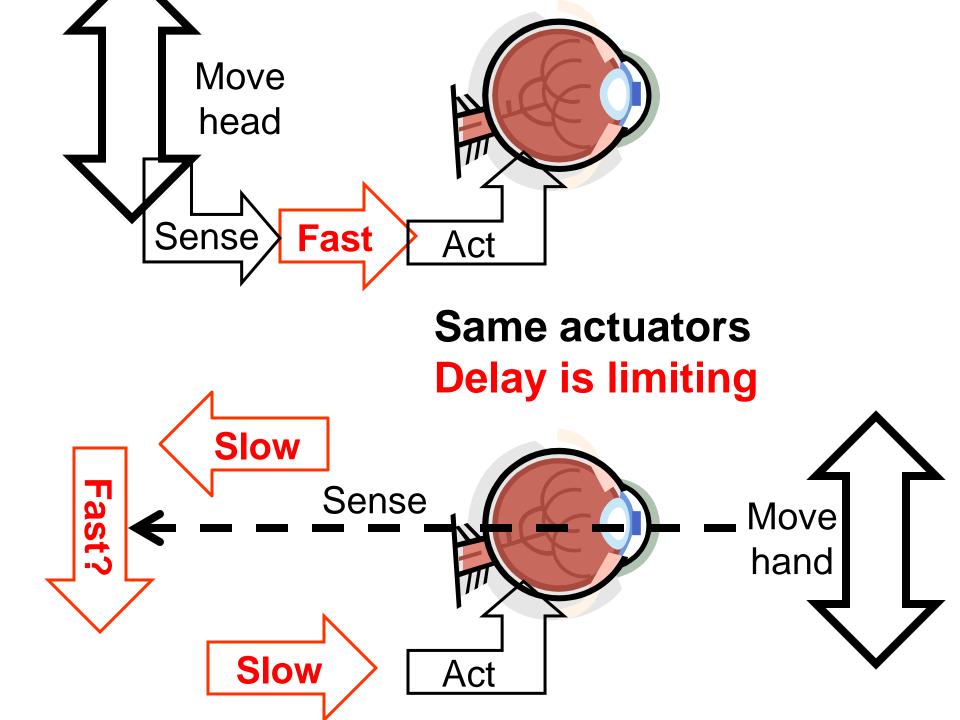


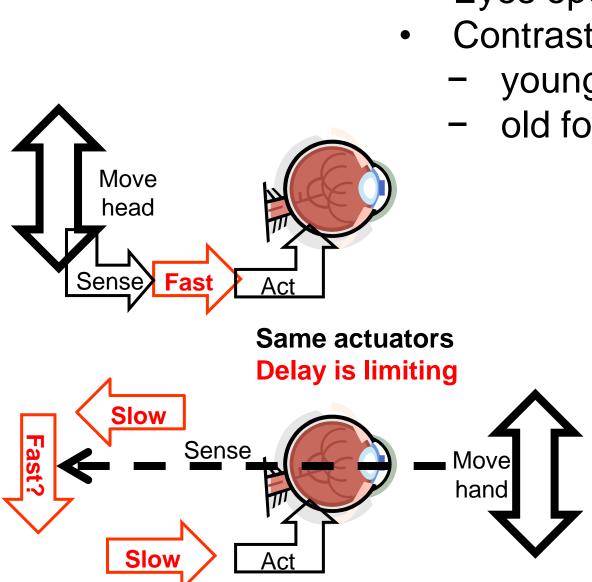


3. Compensating eye movement



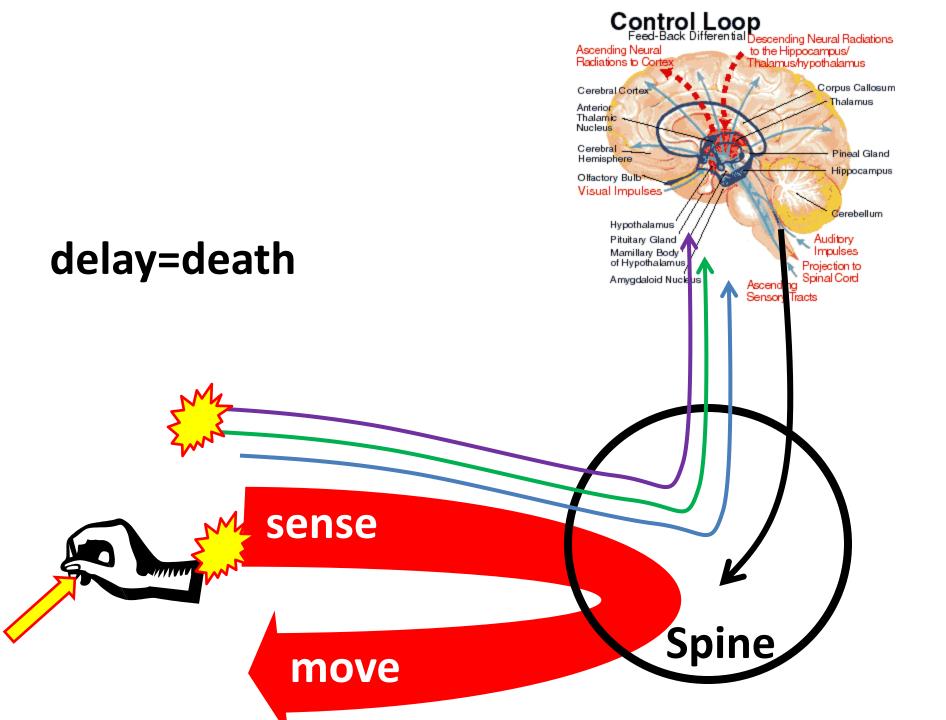






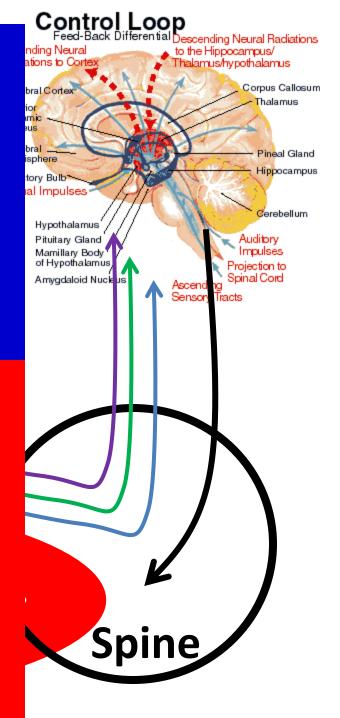
Versus standing on one leg

- Eyes open vs closed
- Contrast
 - young surfers
 - old football players



Reflect







Reflect

Control Loop Feed-Back Differential Descending Neural Radiations to the Hippocampus/ mus/hypothalamus

Corpus Callosum Thalamus

> Pineal Gland Hippocampus

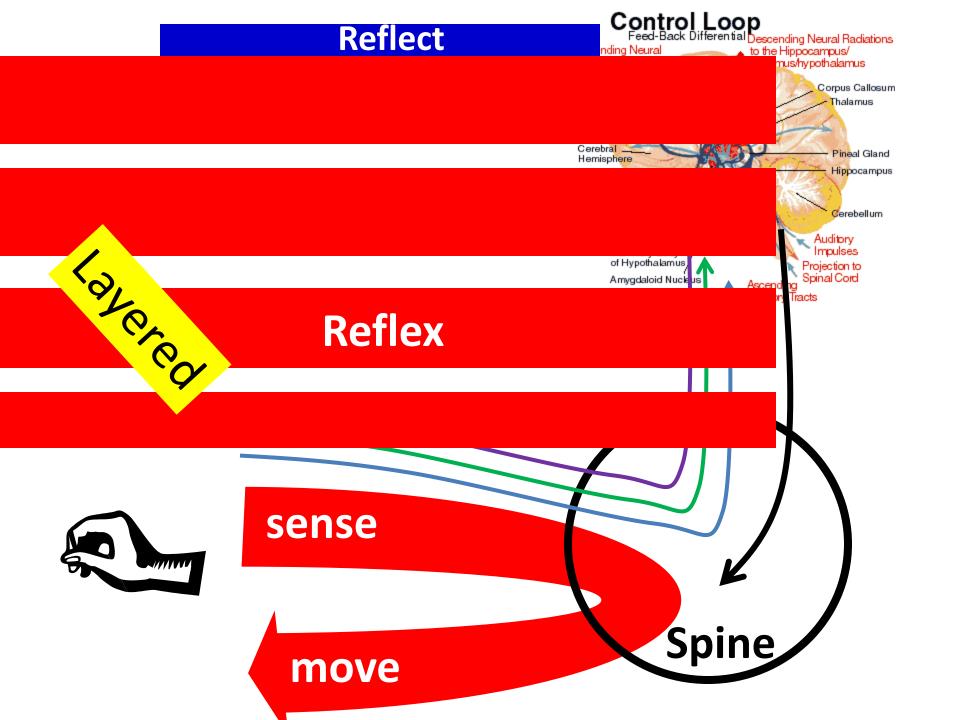
Cerebellum

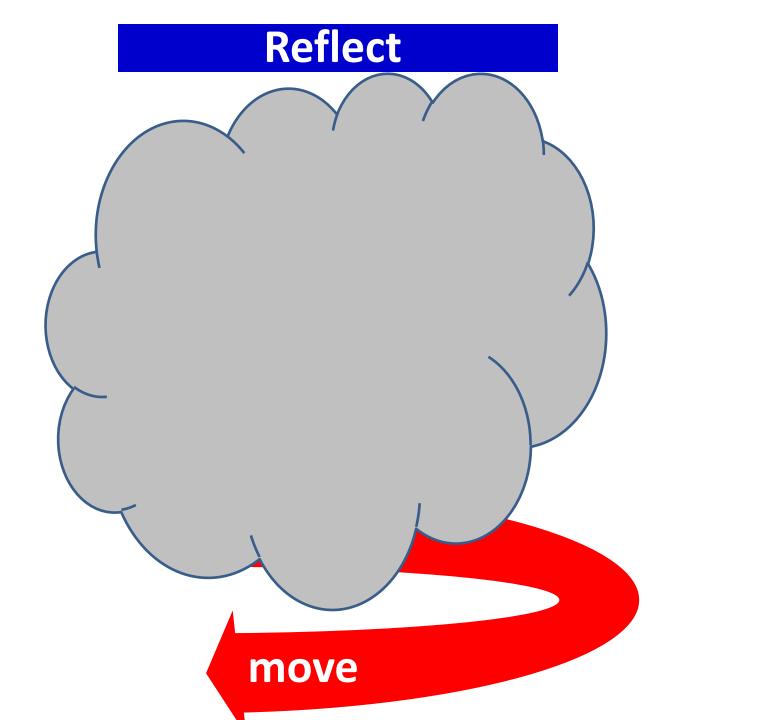
Auditory Impulses

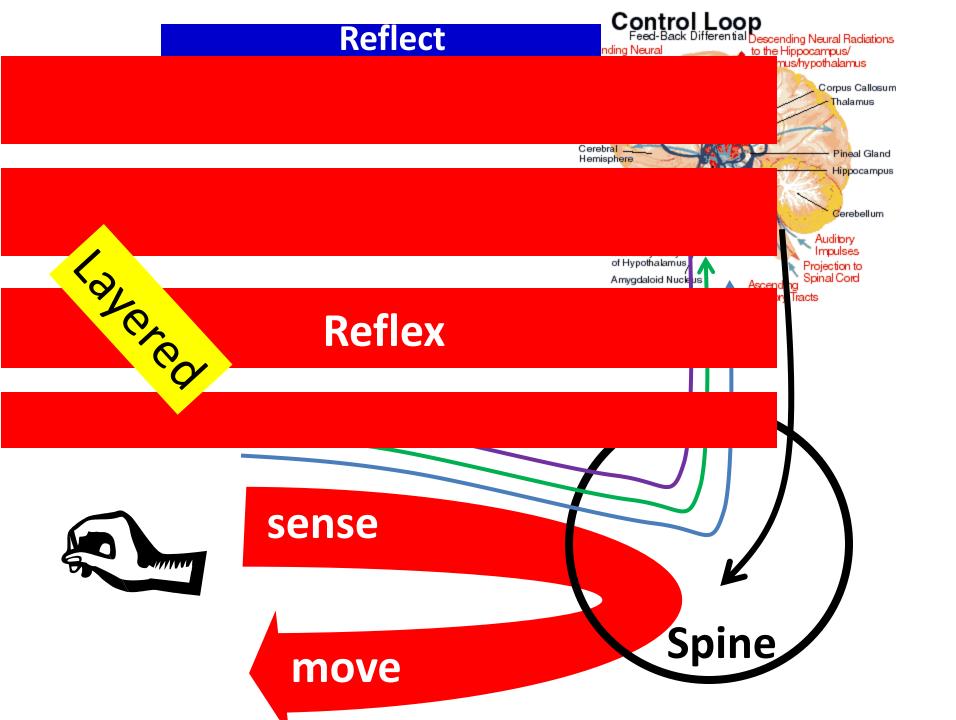
Projection to Spinal Cord

ng Tracts

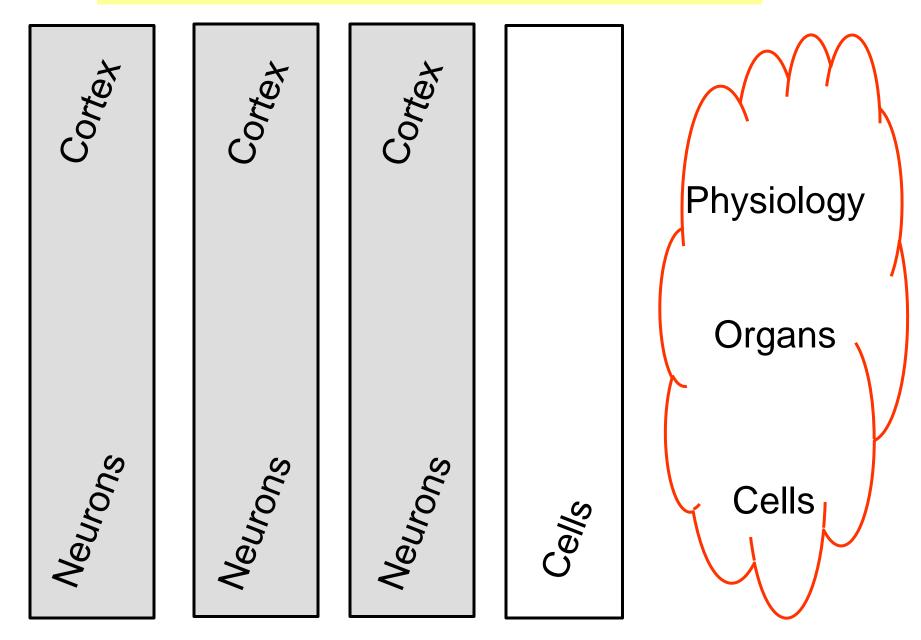
Reflex

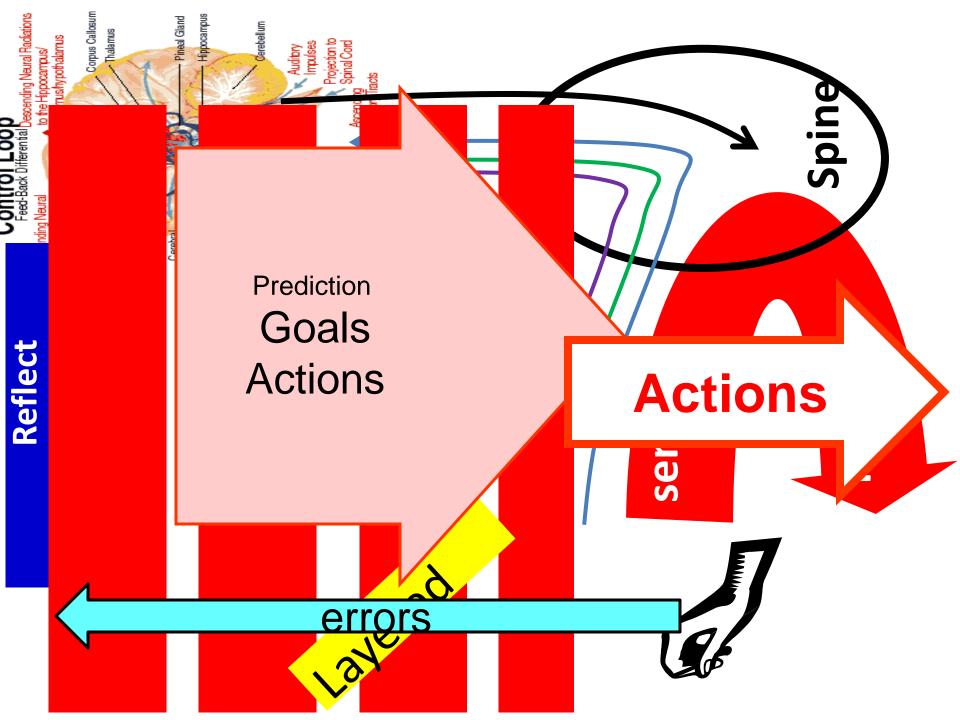


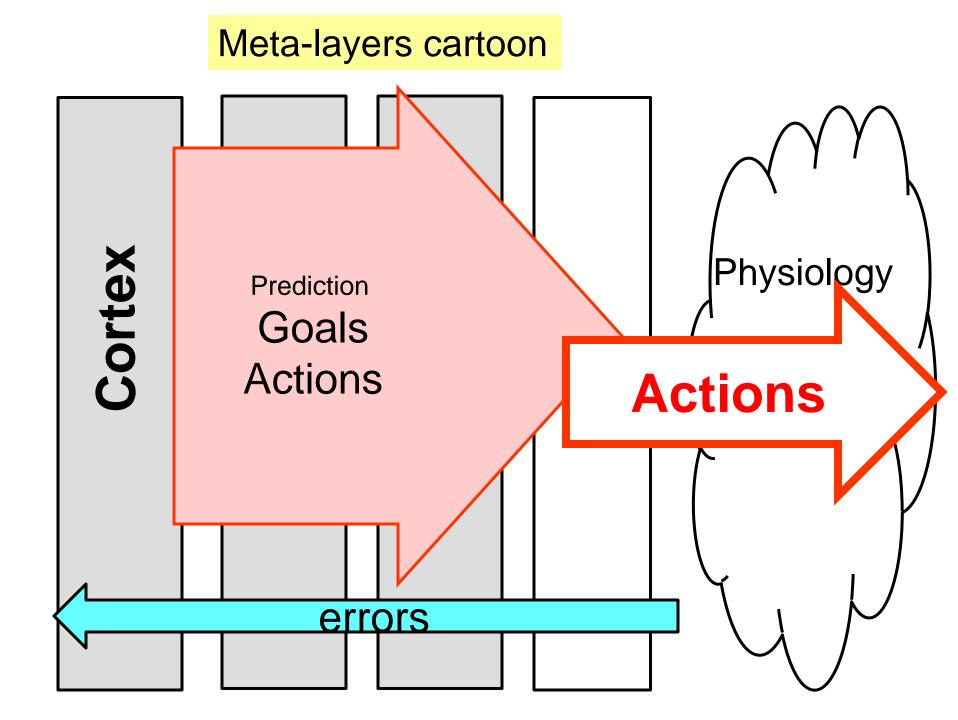


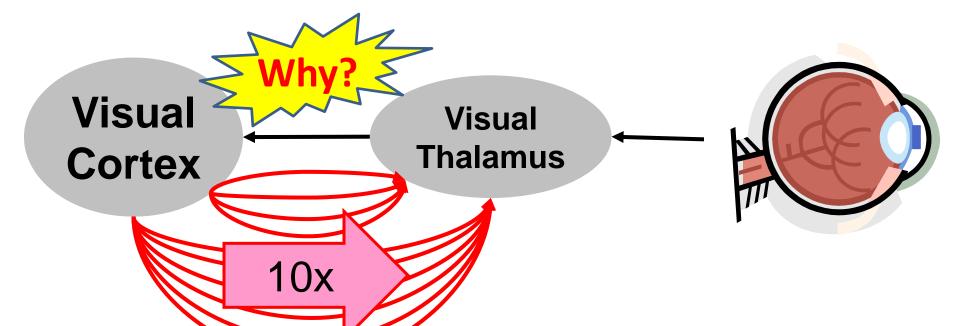


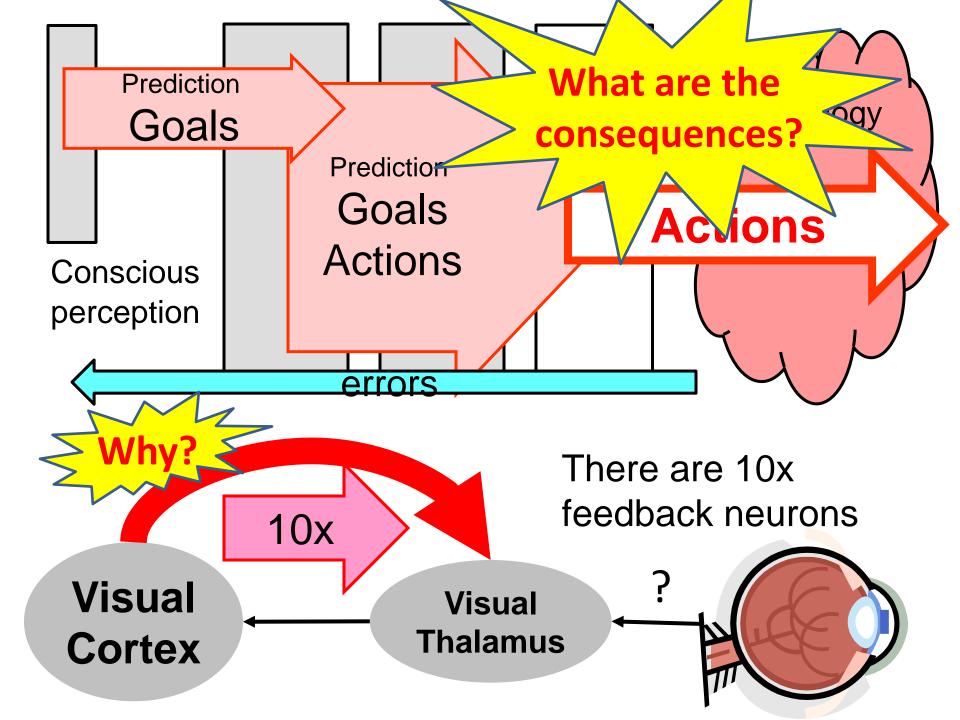
Layered architectures (cartoon)



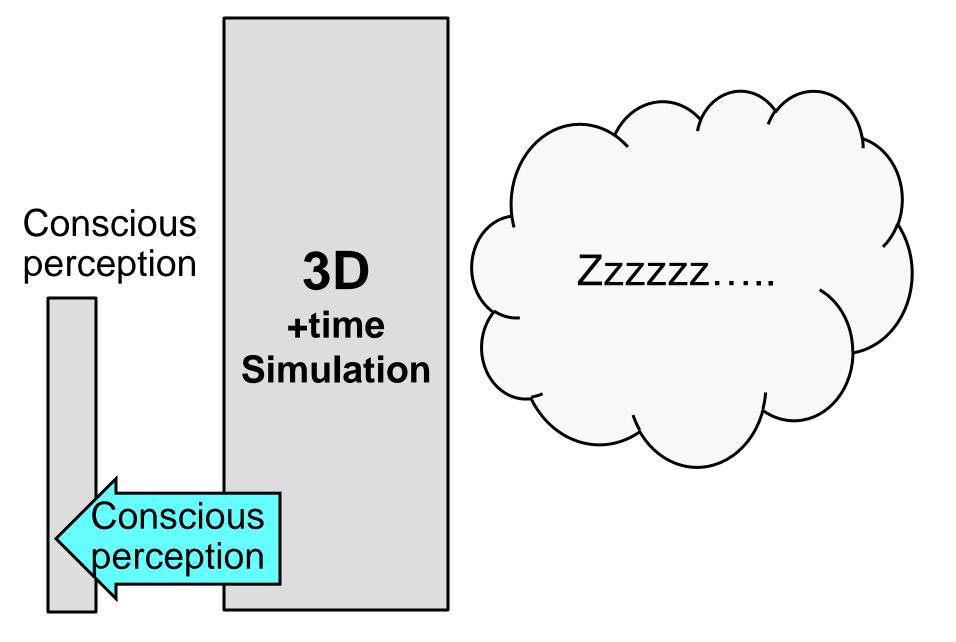


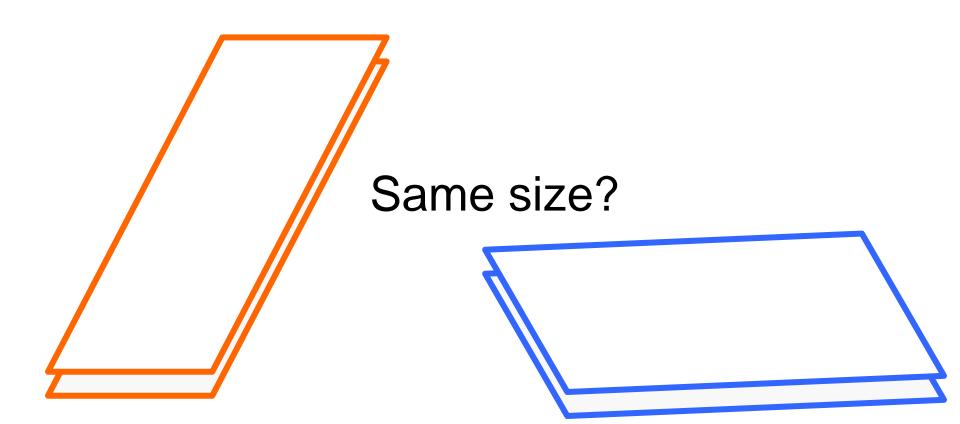


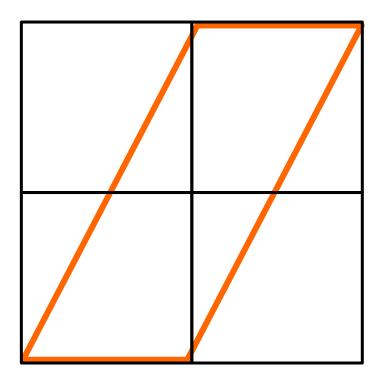


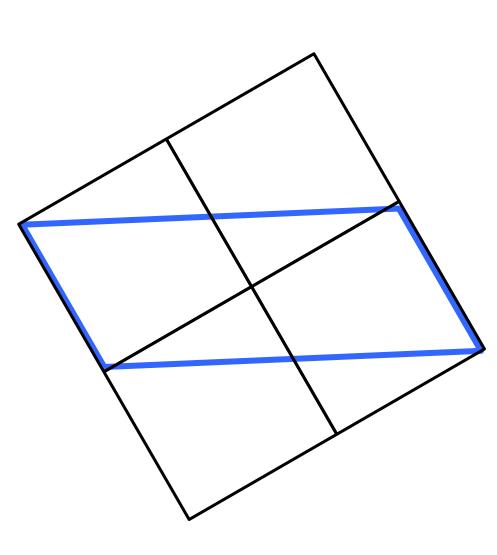


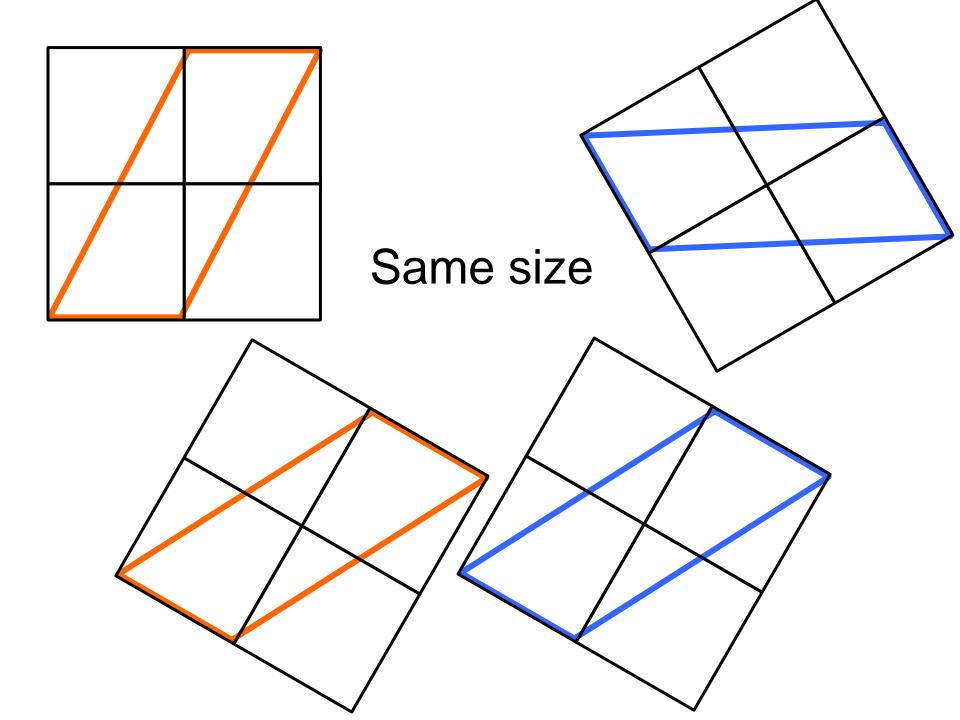
Seeing is *dreaming*

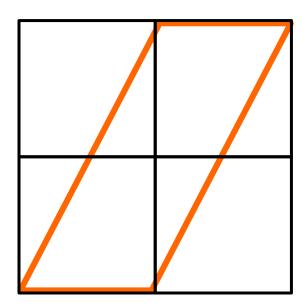


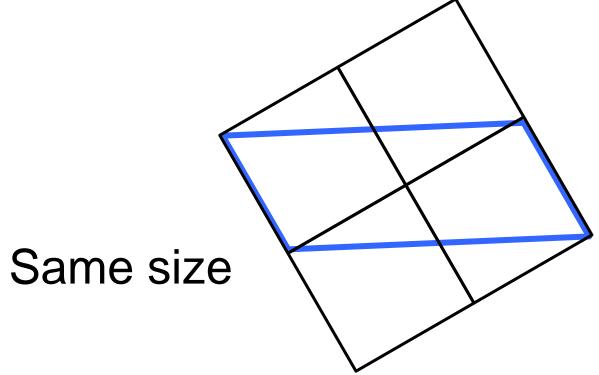


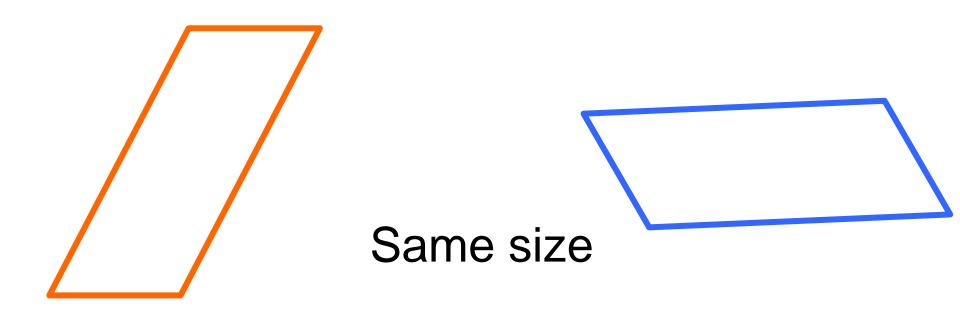












Toggle between this slide and the ones before and after

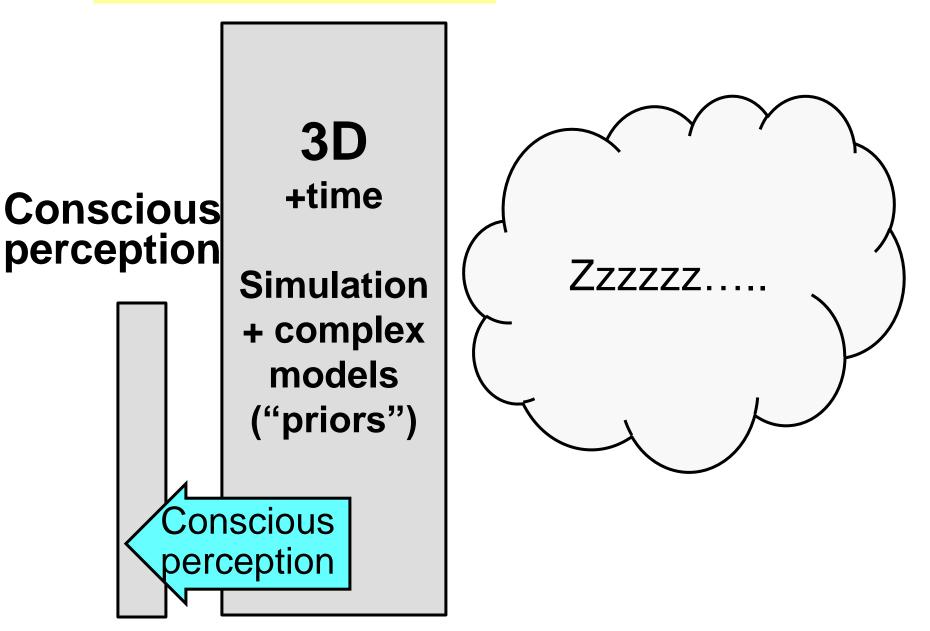
Even when you "know" they are the same, they appear different

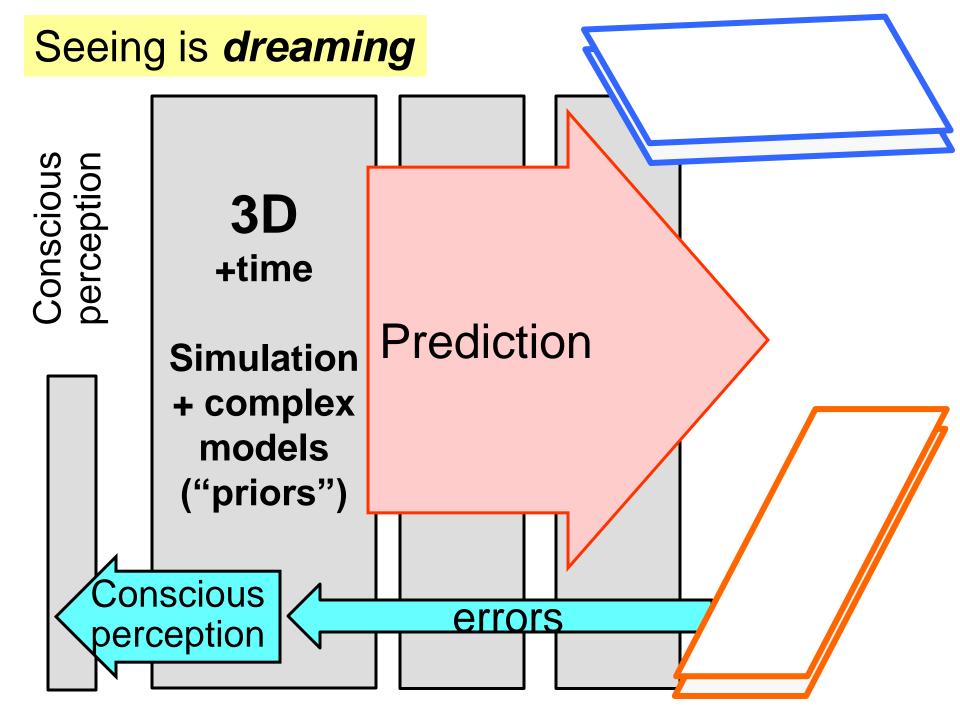


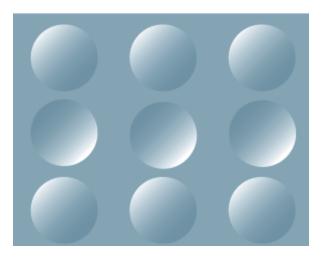
Vision: evolved for complex simulation and control, not 2d static pictures

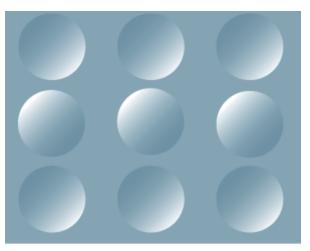
Even when you "know" they are the same, they appear different

Seeing is *dreaming*

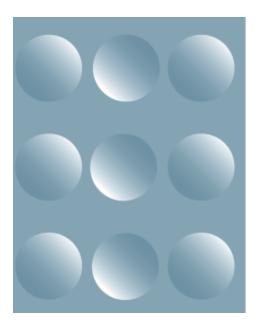


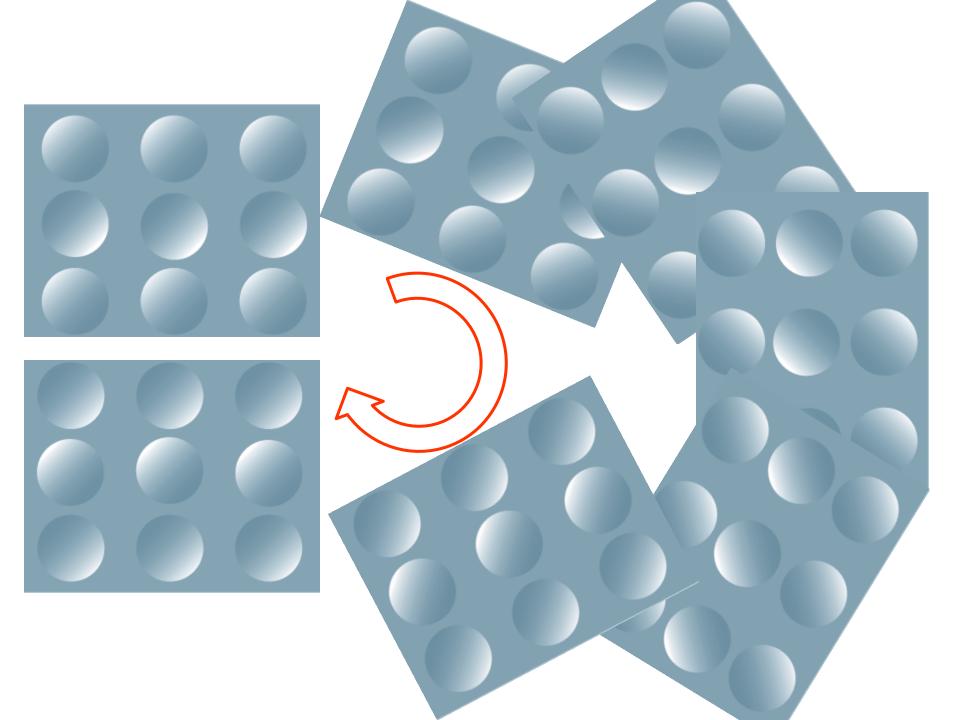


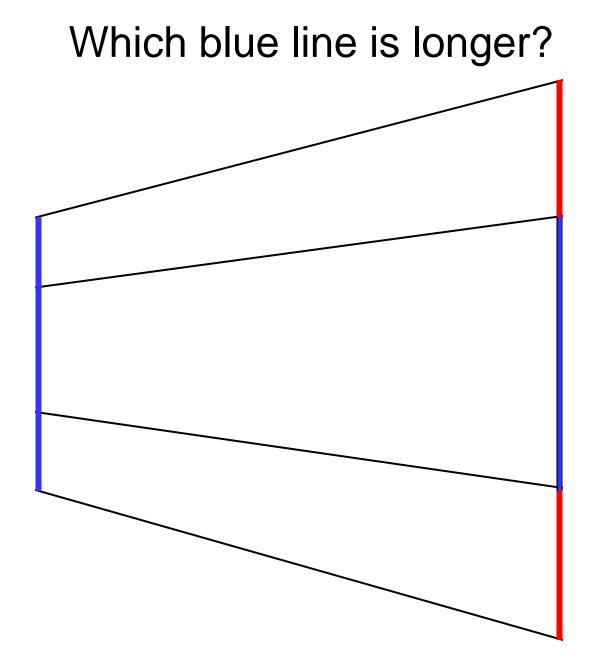


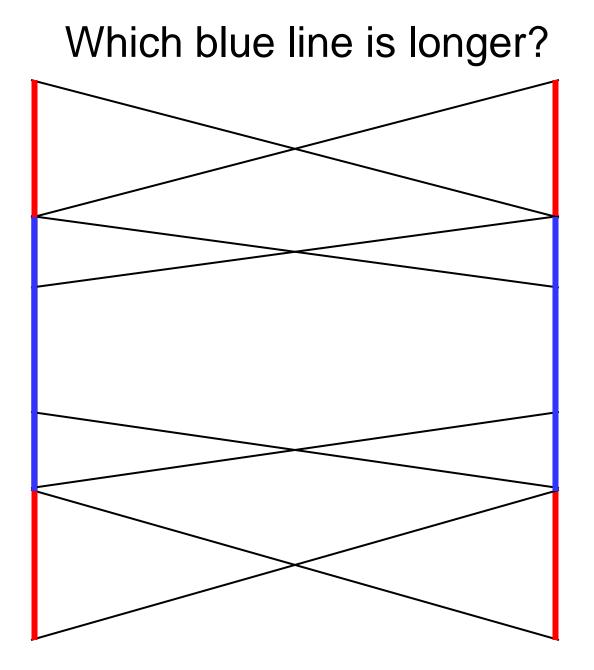


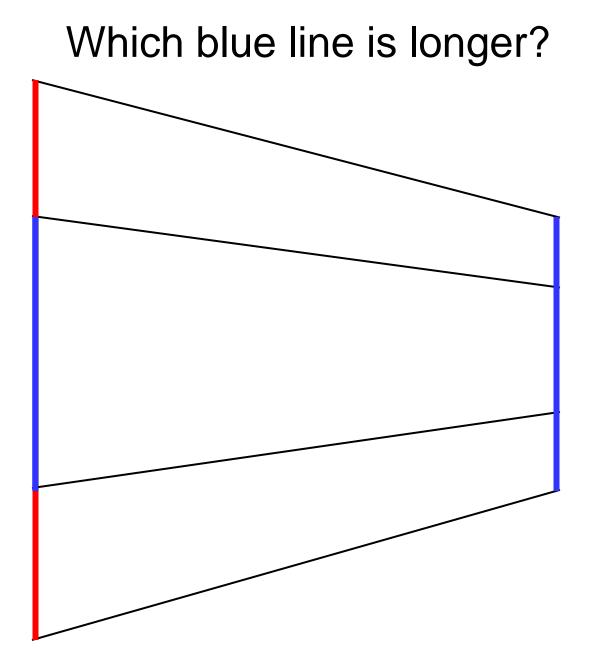
Inferring shape from shading

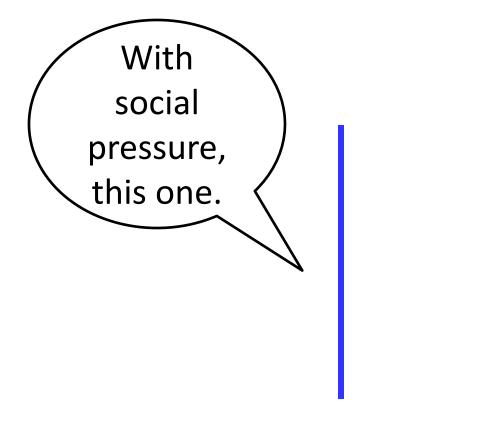












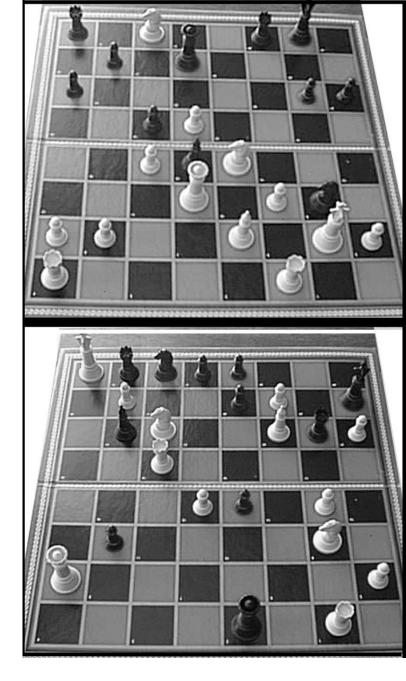
Standard social psychology experiment.



Chess experts

- can reconstruct entire chessboard with < ~ 5s inspection
- can recognize 1e5 distinct patterns
- can play multiple games
 blindfolded and simultaneous
- are no better on random boards

(Simon and Gilmartin, de Groot)



www.psywww.com/intropsych/ch07_cognition/expertise_and_domain_specific_knowledge.html

Specialized Face Learning Is Associated with Individual Recognition in Paper Wasps



Michael J. Sheehan* and Elizabeth A. Tibbetts

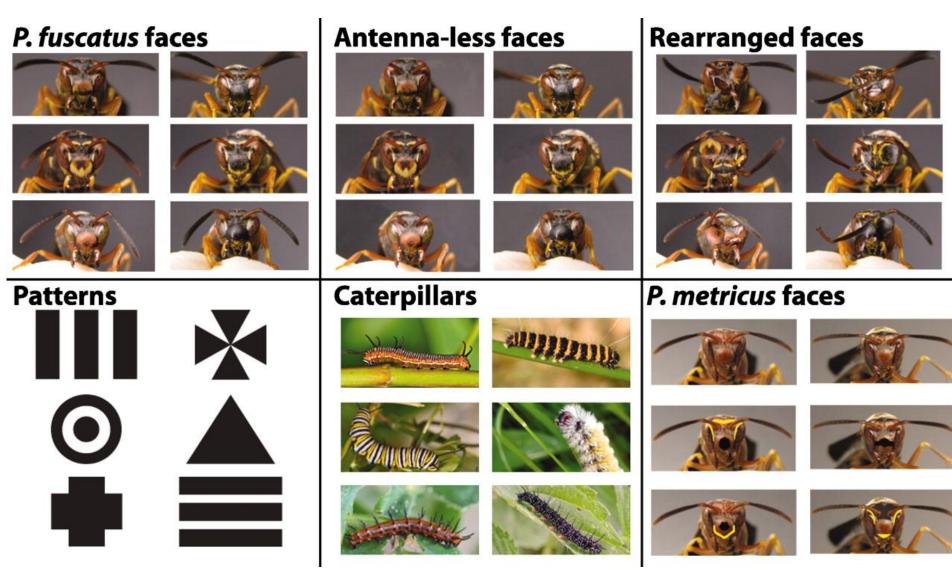
We demonstrate that the evolution of facial recognition in wasps is associated with specialized face-learning abilities. *Polistes fuscatus* can differentiate among normal wasp face images more rapidly and accurately than nonface images or manipulated faces. A close relative lacking facial recognition, *Polistes metricus*, however, lacks specialized face learning. Similar specializations for face learning are found in primates and other mammals, although *P. fuscatus* represents an independent evolution of specialization. Convergence toward face specialization in distant taxa as well as divergence among closely related taxa with different recognition behavior suggests that specialized cognition is surprisingly labile and may be adaptively shaped by species-specific selective pressures such as face recognition.

When needed, even wasps can do it.

2 DECEMBER 2011 VOL 334 SCIENCE www.sciencemag.org

- *Polistes fuscatus* can differentiate among normal wasp face images more rapidly and accurately than nonface images or manipulated faces.
- *Polistes metricus* is a close relative lacking facial recognition and specialized face learning.
- Similar specializations for face learning are found in primates and other mammals, although *P. fuscatus* represents an independent evolution of specialization.
- Convergence toward face specialization in distant taxa as well as divergence among closely related taxa with different recognition behavior suggests that specialized cognition is surprisingly labile and may be adaptively shaped by species-specific selective pressures such as face recognition.

Fig. 1 Images used for training wasps.





M J Sheehan, E A Tibbetts Science 2011;334:1272-1275

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