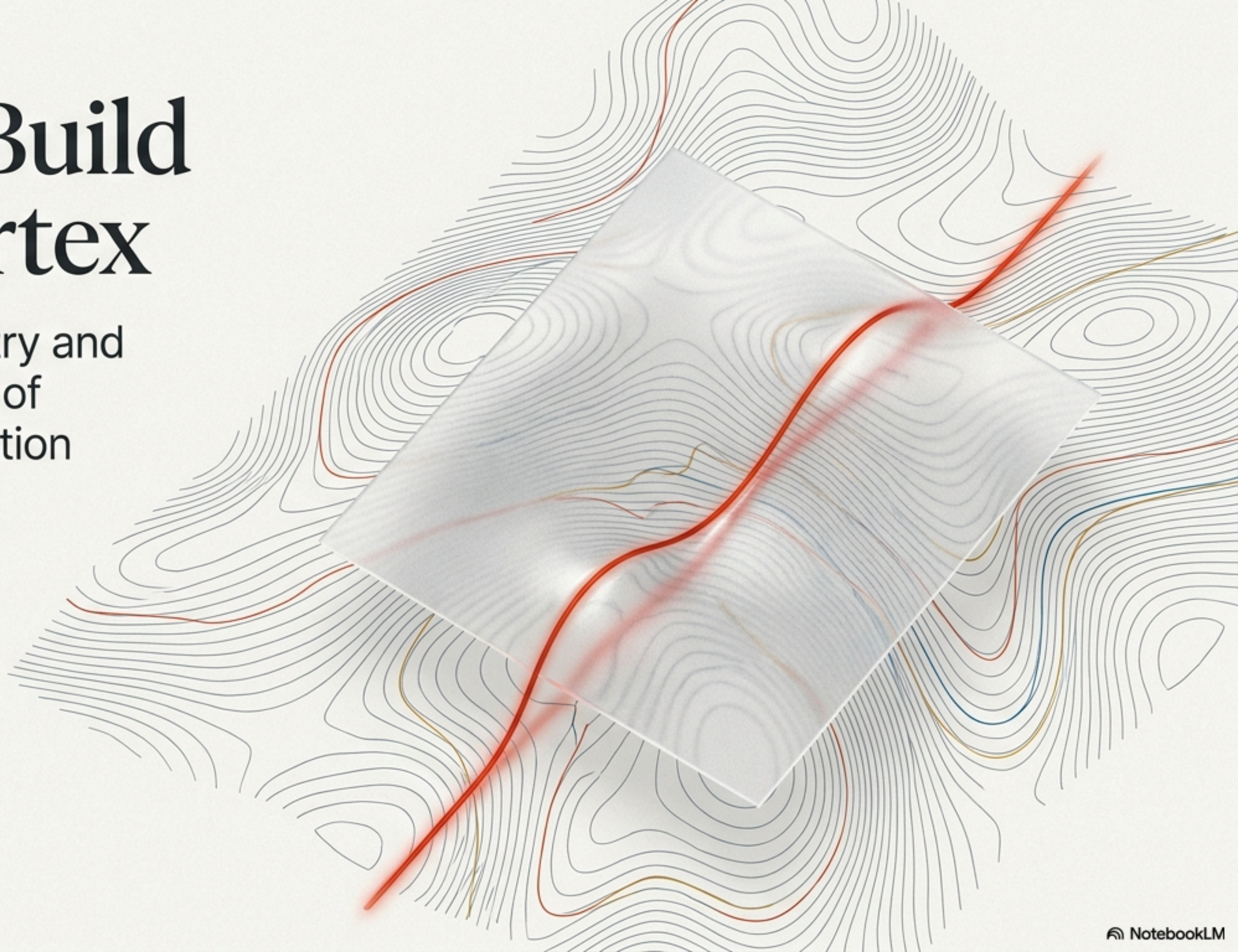


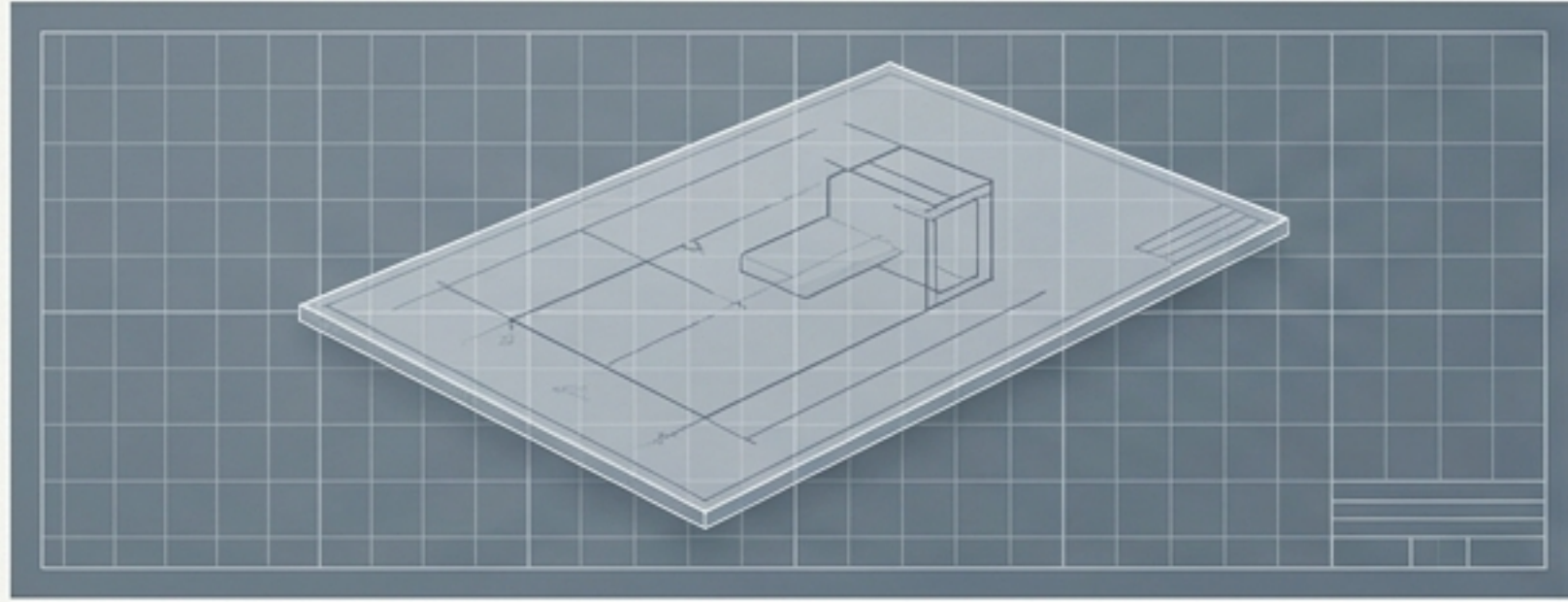
# How to Build a Neocortex

Constraint Geometry and  
the Active Medium of  
Biological Organization

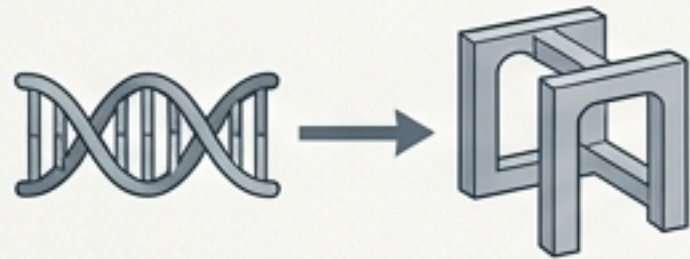


# The Blueprint Model of Biology is Dead

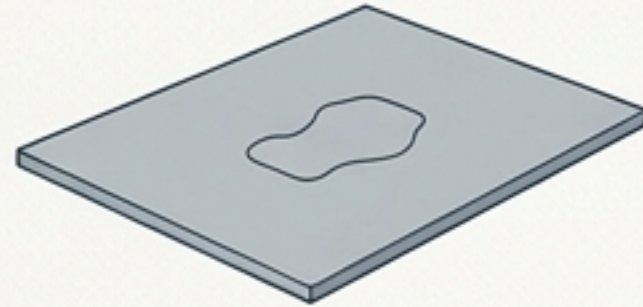
## The Old Paradigm: The Blueprint Model



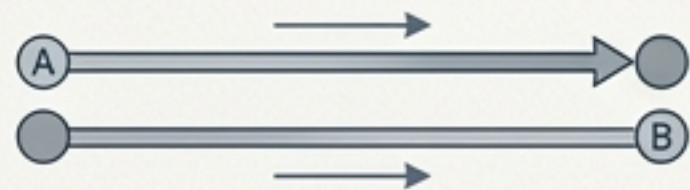
- **Direct Specification:**  
Genes encode exact structures.



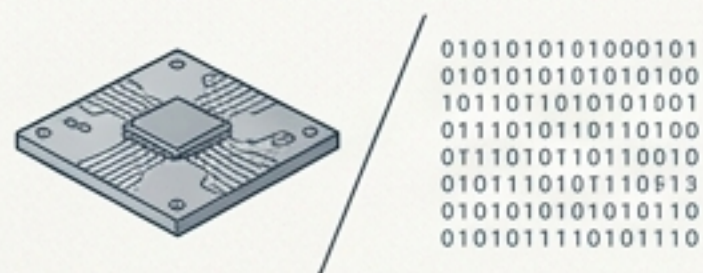
- **Passive Medium:**  
Tissue is a blank canvas.



- **Fixed Channels:**  
Signals flow through static wires.

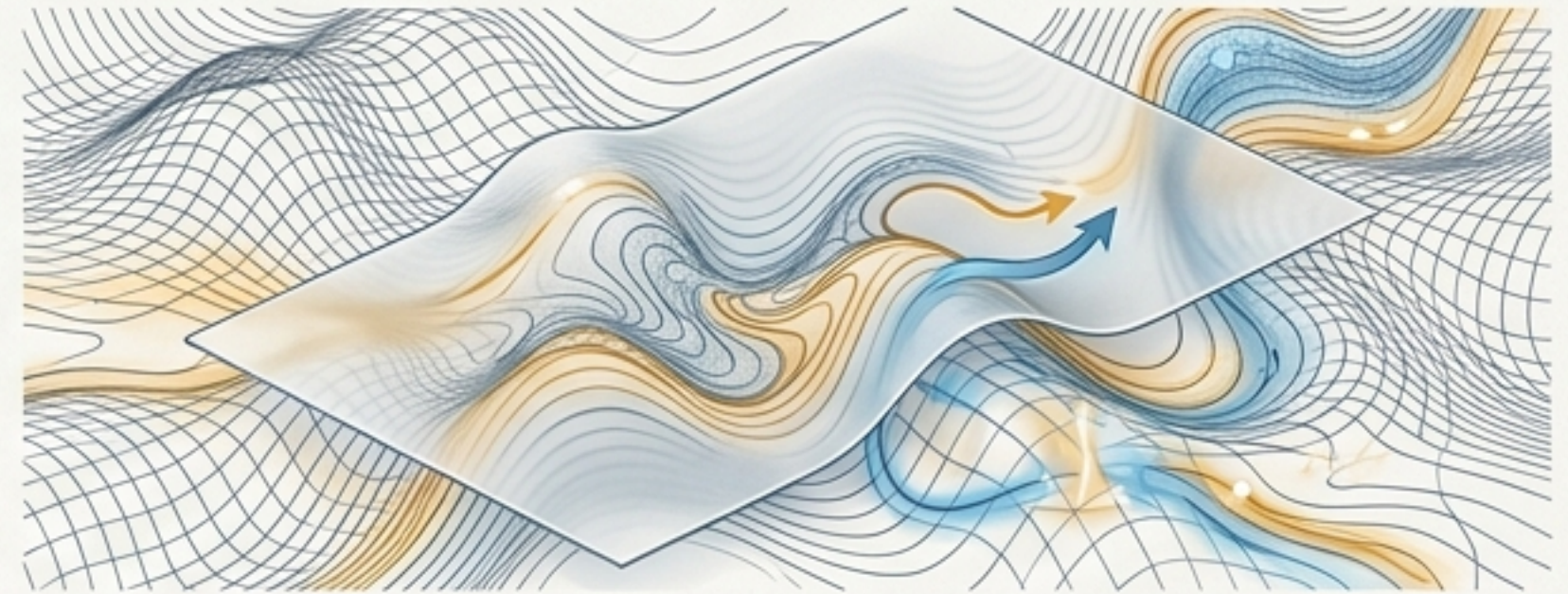


- **Dichotomy:**  
Hardware vs. Software split.

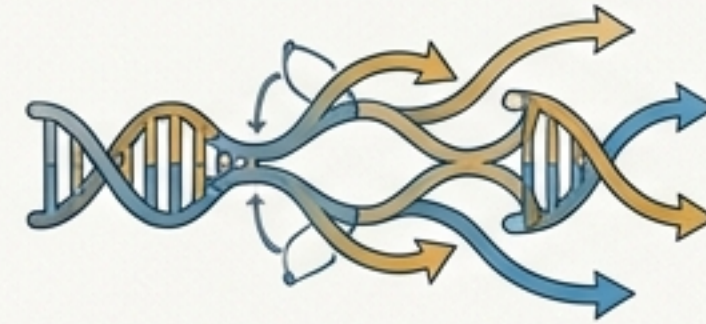


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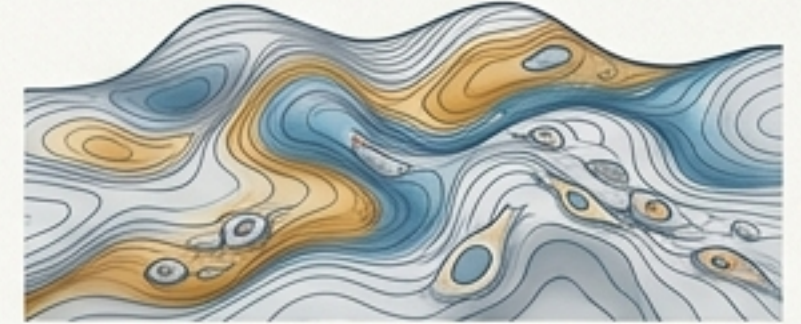
## The New Paradigm: Constraint Geometry



- **Accessibility Modulated:**  
Genes open/close dynamic pathways.



- **Active Medium:**  
Tissue reshapes itself continuously.



- **Self-Modifying:**  
Signals physically alter the wires.



- **Unification:**  
Hardware is Software.



# The Architecture of Biological Explanation

## 1. State Manifold ( $\mathcal{X}$ )

The fixed spatial boundaries of all possible configurations.

## 2. Admissibility Structure ( $\Gamma_{\text{adm}}$ )

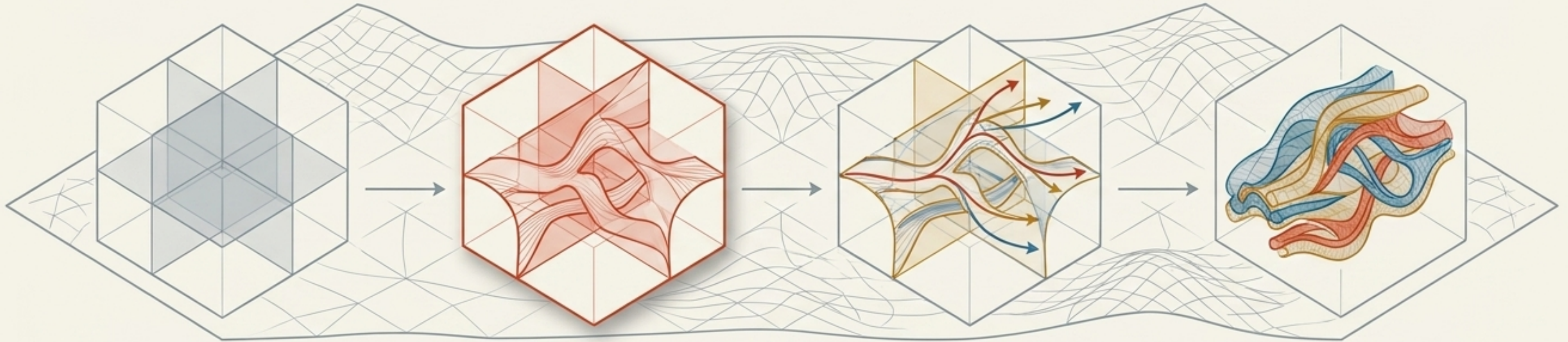
The dynamically evolving operator  $C(t)$  defining which paths are physically open.

## 3. Stochastic Dynamics ( $\mu_t$ )

The probability measure selecting among the available open paths.

## 4. Realized Structure ( $\langle \gamma \rangle$ )

The observable biological outcome (anatomy, gene expression).



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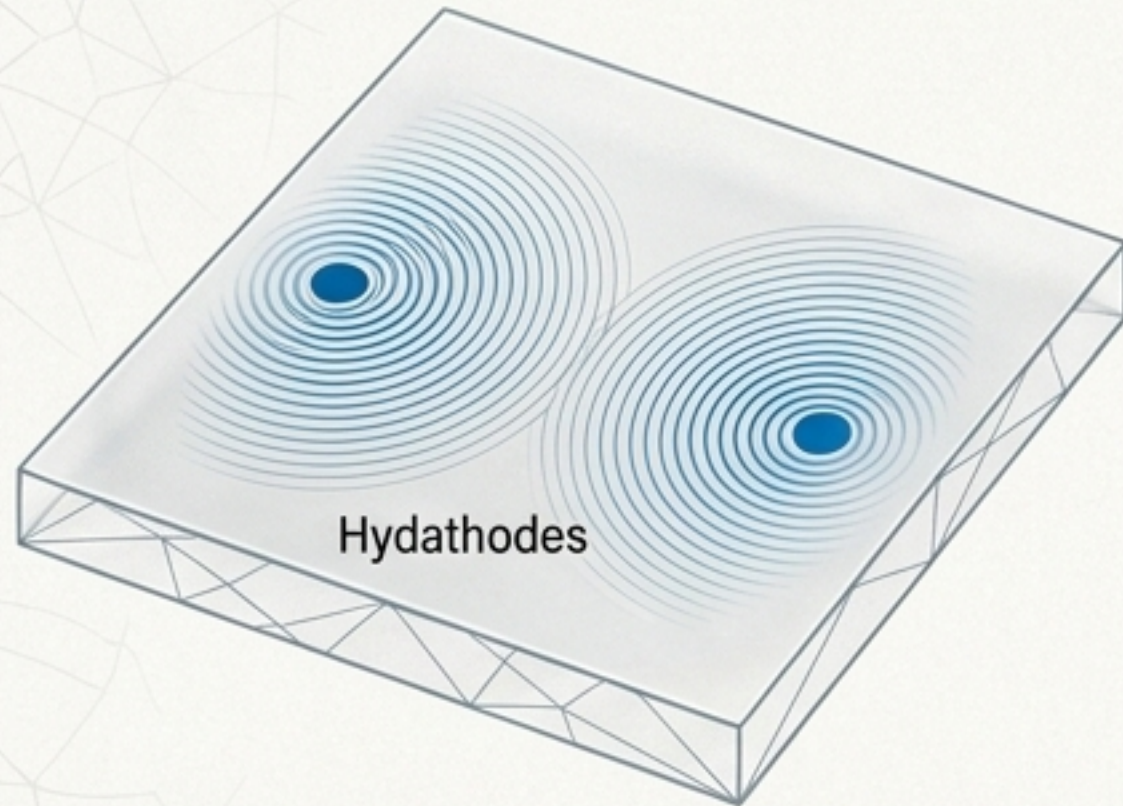
The probability measure selecting among the available open paths.

## 4. Realized Structure ( $\langle \gamma \rangle$ )

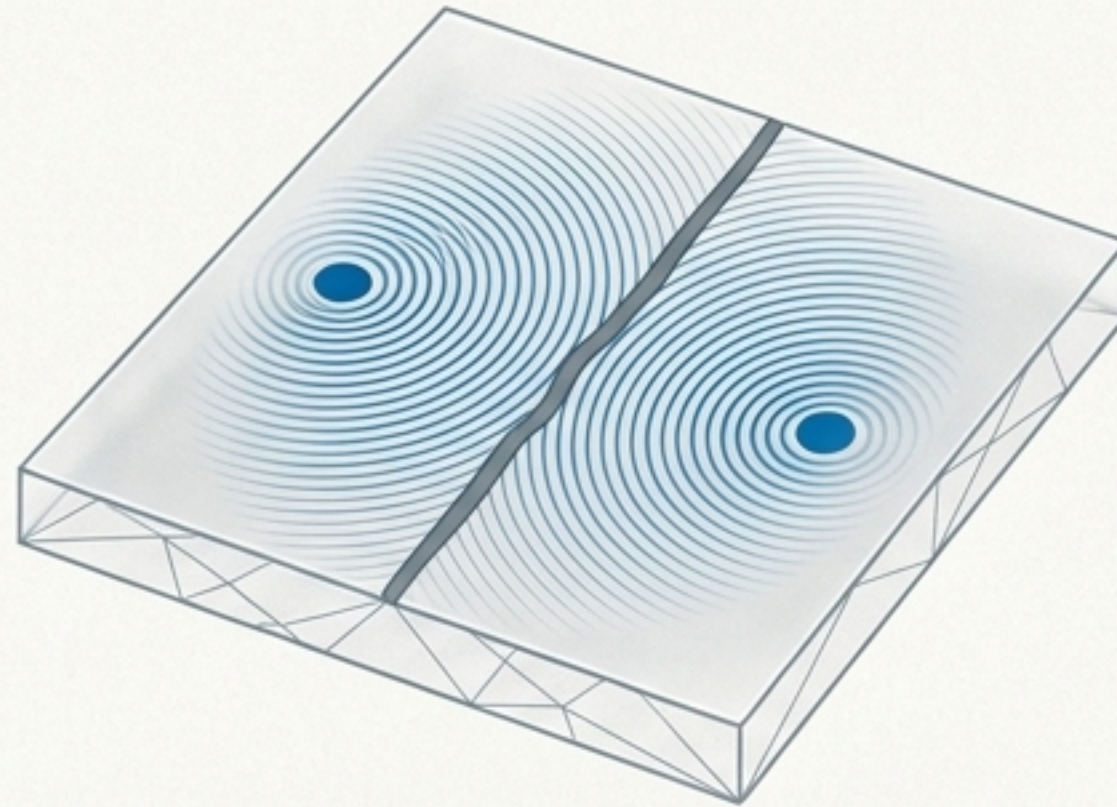
The observable biological outcome (anatomy, gene expression).

Evolution and regulation occur here: in the manipulation of geometric constraints, not the specification of final structures.

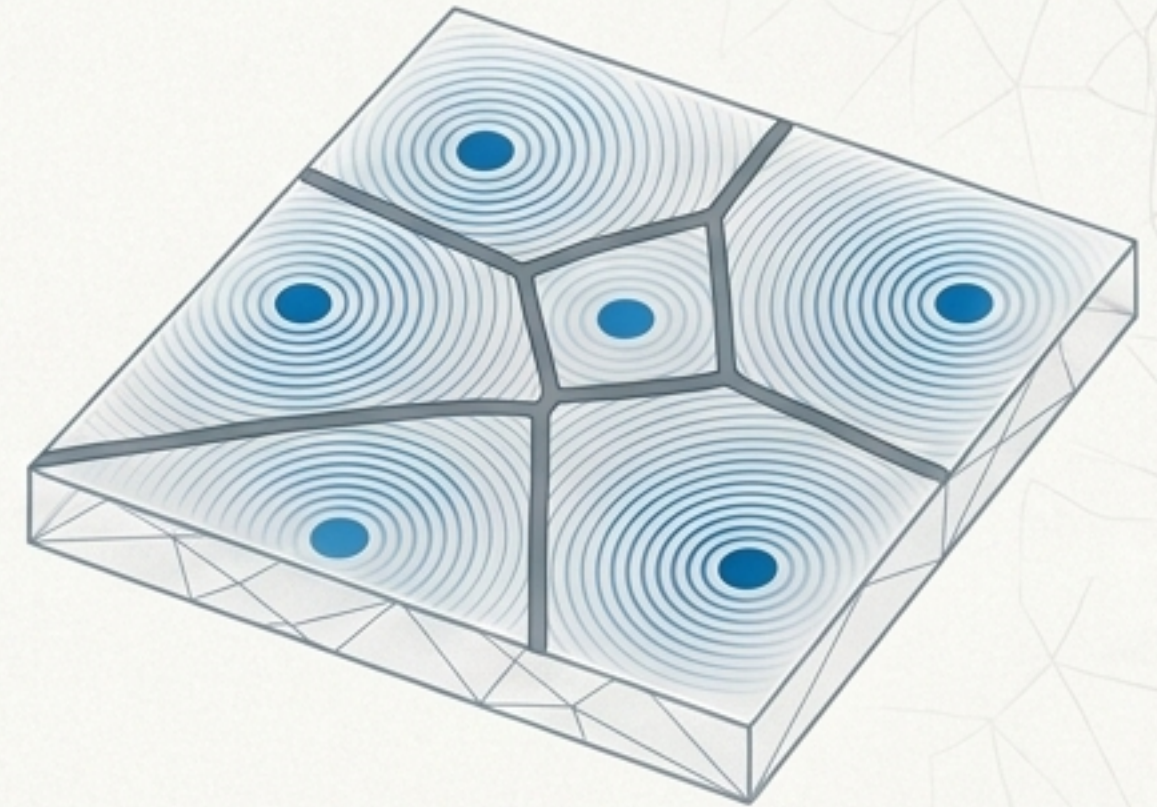
# Scale I: The Spatial Geometry of Partition in Leaves



1. Secretory hydathodes emit propagating auxin waves.



2. Waves collide, halting propagation and forming a structural ridge.



3. Localized collisions resolve into global Voronoi vein partitions.

## The Blueprint Rejection

The *Pilea* leaf does not encode vein locations; it positions hydathodes. The geometry emerges from wave competition. The Voronoi model out-performs hierarchical biological k-d tree alternatives by a massive margin (0.72 vs 0.40 Jaccard overlap index).

# Scale II: Signaling Through a Changing Material Medium

## Active Regulatory Media

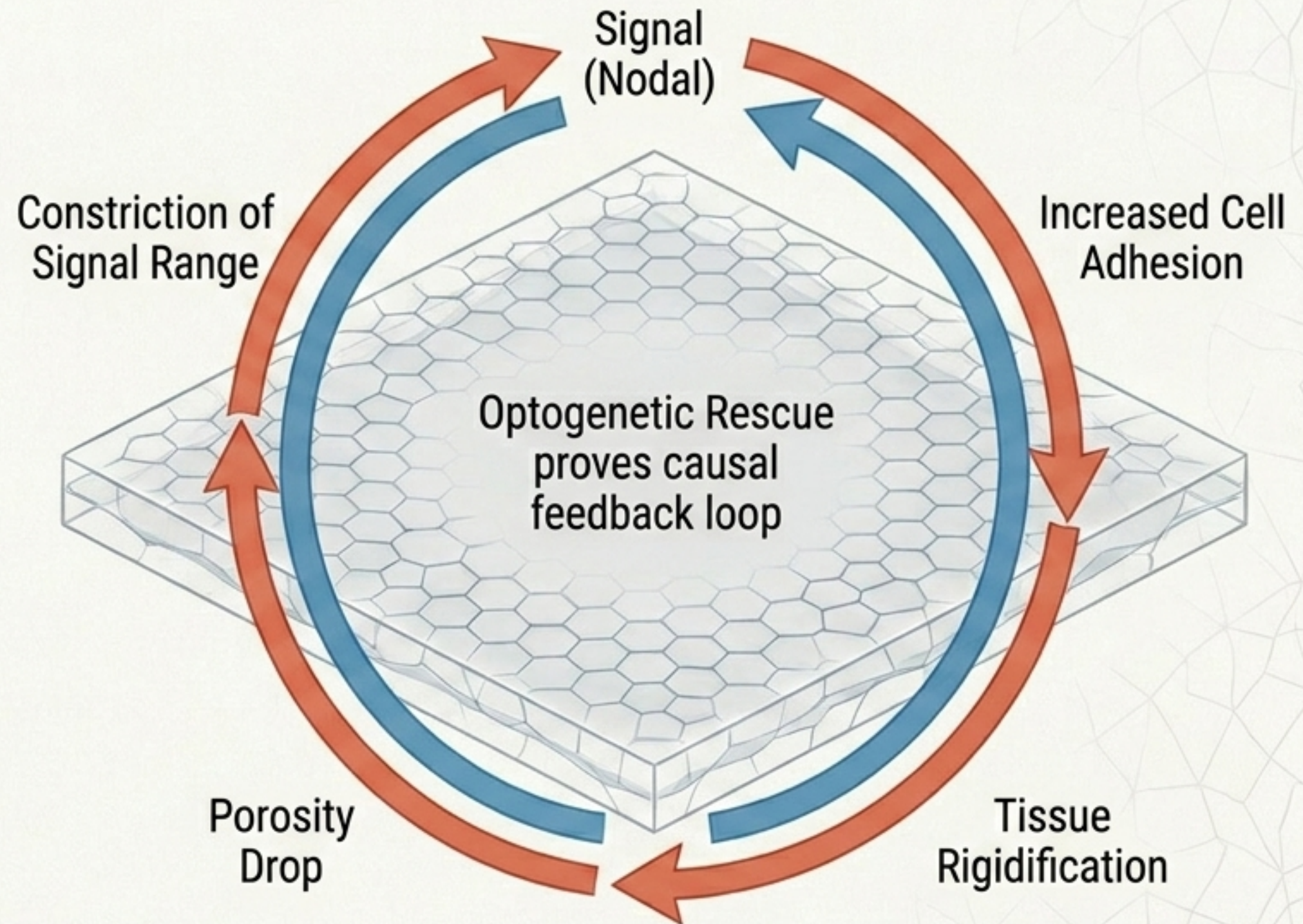
Tissue mechanics are not passive backdrops. In the zebrafish blastoderm, the physical medium actively participates in signal regulation.

## Fluid Tissue (Mutant):

Without rigidity, Nodal diffuses infinitely through porous channels.

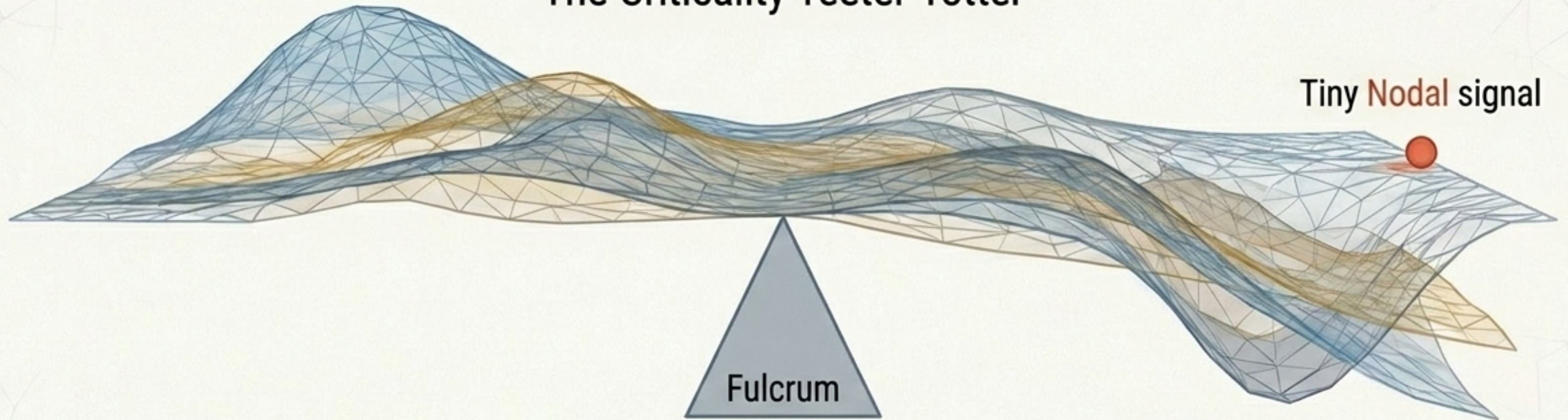
## Rigid Tissue (Wild-Type):

Tissue rigidification collapses interstitial porosity, physically trapping the signal and shaping the morphogen gradient. Restoring rigidity optogenetically fully restores geometric signaling.



# Maximum Constraint Susceptibility

## The Criticality Teeter-Totter



### The Edge of Chaos

Embryonic tissues deliberately poise themselves near phase transition critical points (e.g., cell connectivity  $k_c \approx 4$ ).

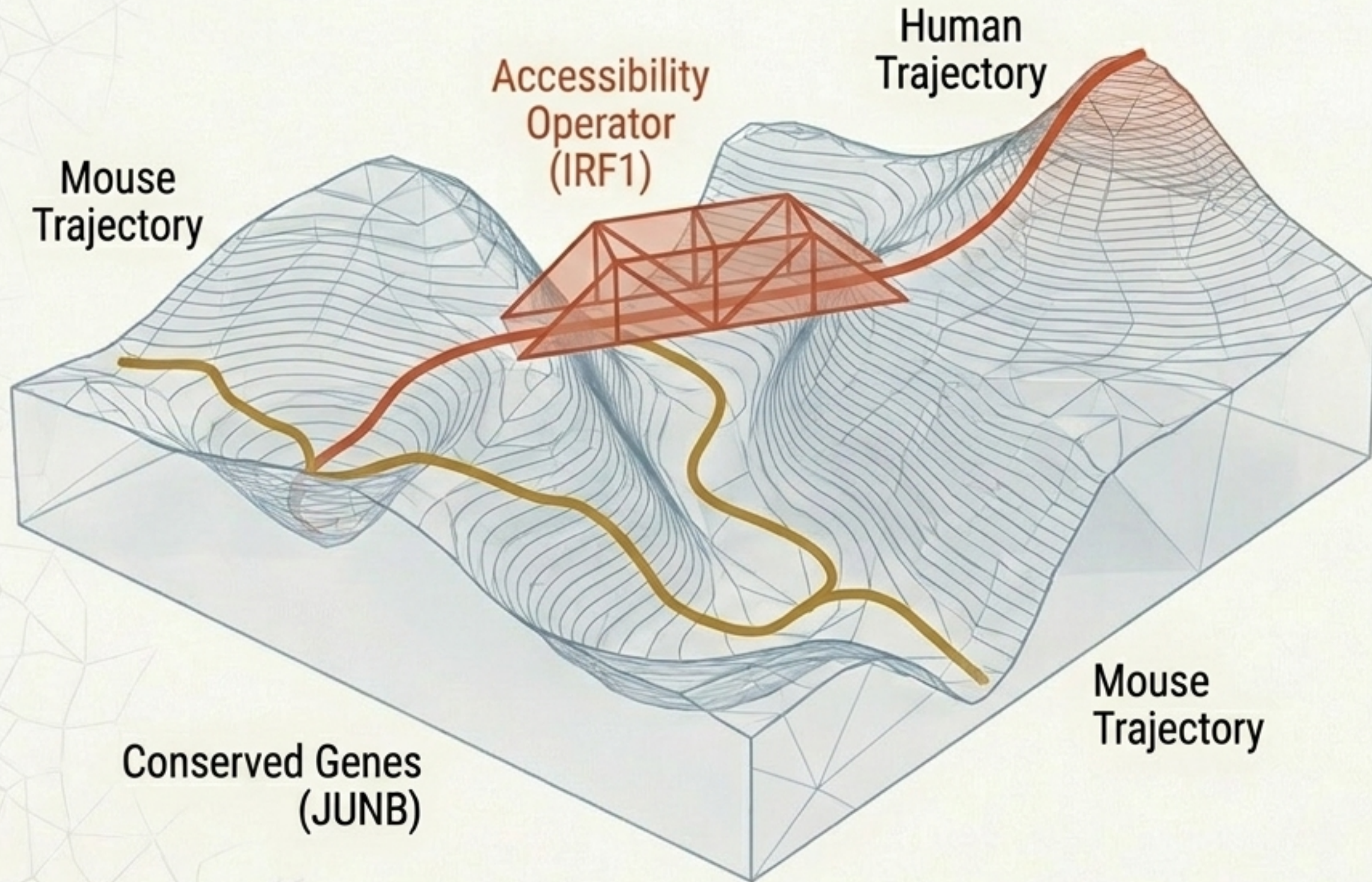
### Infinite Susceptibility

Near criticality, constraint susceptibility diverges. The tissue is in a state of maximum geometric plasticity.

### Informational Leverage

A tiny molecular signal causes massive reorganization of the accessible state space. Biology leverages instability for precise structural control.

# Scale III: Navigating the Developmental Manifold

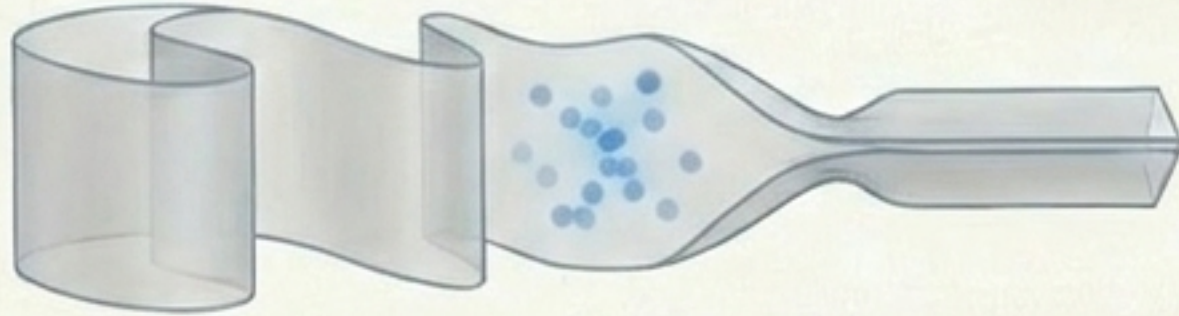


## Evolution Through Accessibility

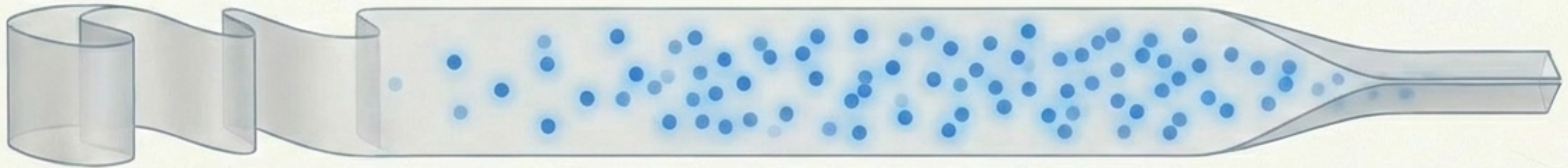
- Evolution does not invent new genes to build a human neocortex. It uses upstream regulators like IRF1 to open access to latent, poised programs (JUNB) in early progenitor populations.
- The manifold is deeply conserved; the accessibility changes.

# Time is a Structural Component of Accessibility

Mouse: Rapid Constriction



Human: Extended Progenitor Persistence



## Heterochrony as Geometry

Delaying the transition from symmetric to asymmetric progenitor division physically expands the geometric capacity of the developmental compartment.

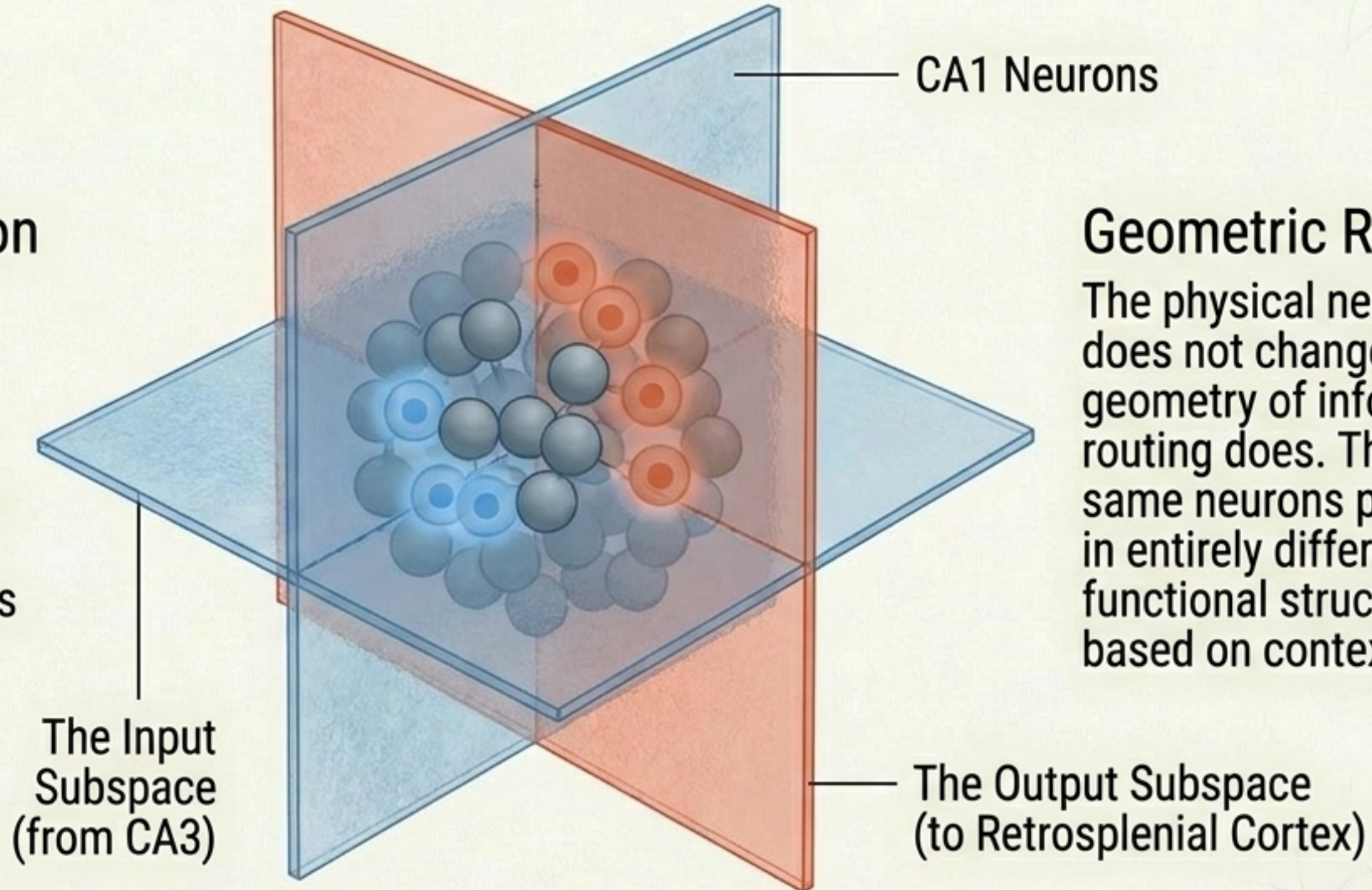
## Structural Output

This temporal dilation generates the massively expanded upper layers of the primate neocortex **without** introducing novel molecular machinery.

# Scale IV: Rotating Communication Subspaces in the Brain

## Dynamic Organization

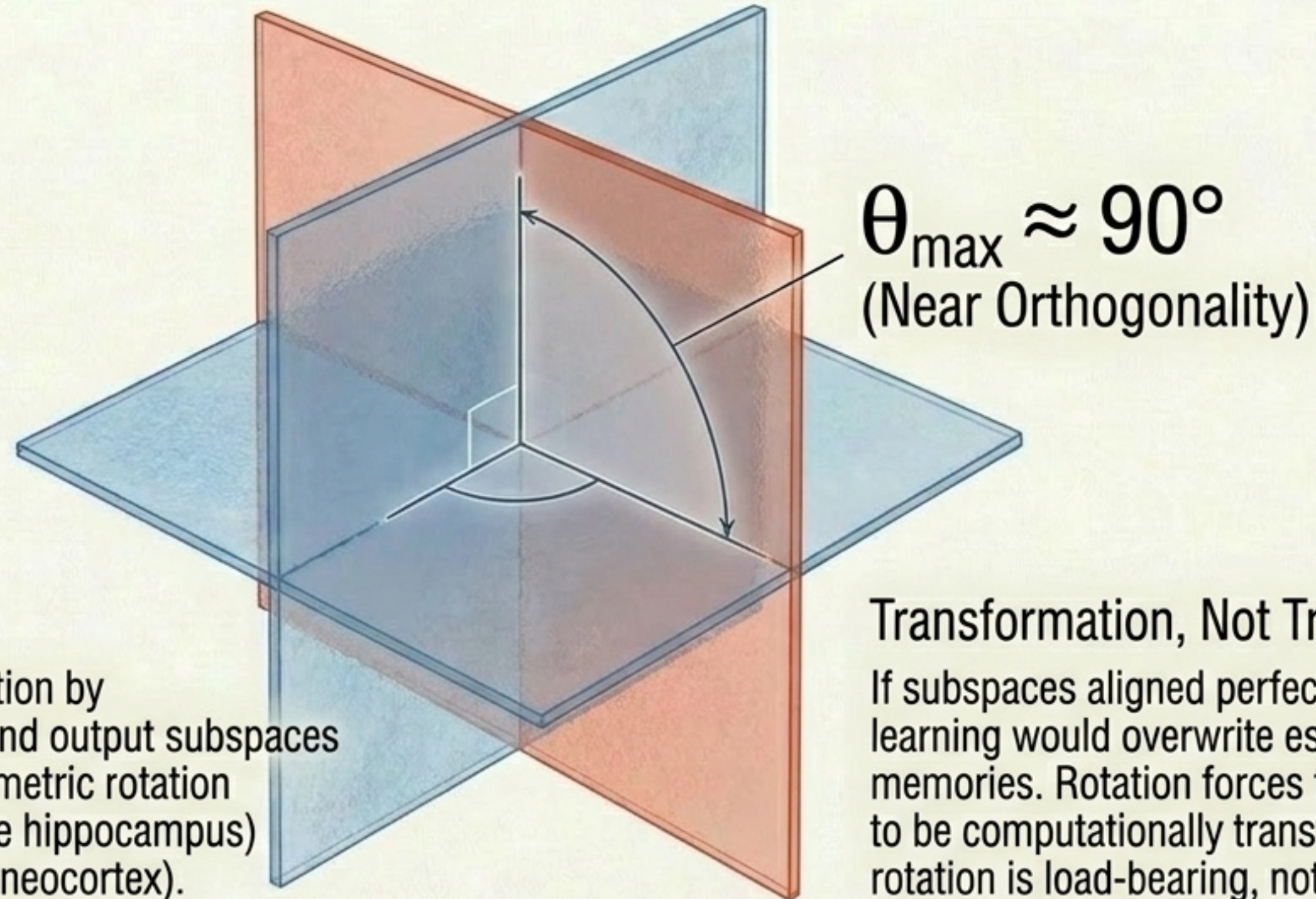
Inter-regional communication is not determined by static anatomical wiring. Information flows through distinct, low-dimensional communication subspaces (identified via pCCA).



## Geometric Routing

The physical neural tissue does not change; the geometry of information routing does. The exact same neurons participate in entirely different functional structures based on context.

# The Computational Necessity of Rotation




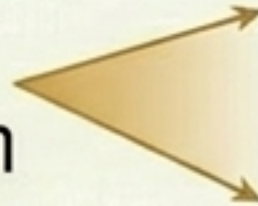
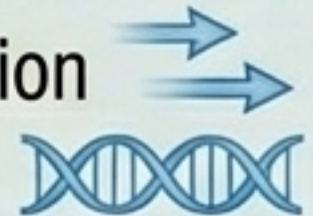
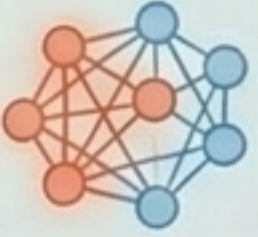
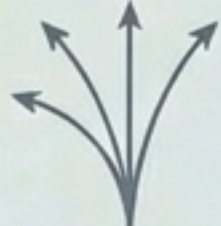
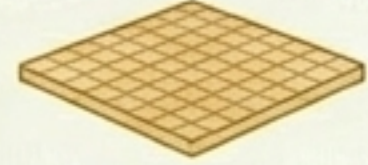


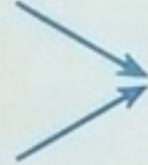
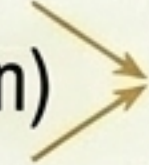
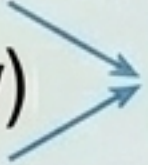
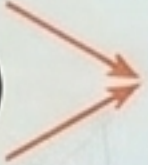

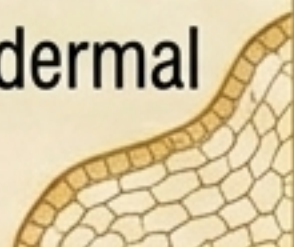

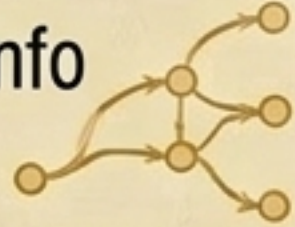
## Separating Functions

The CA1 region routes information by intentionally rotating its input and output subspaces to near-orthogonality. This geometric rotation separates rapid plasticity (in the hippocampus) from long-term stability (in the neocortex).

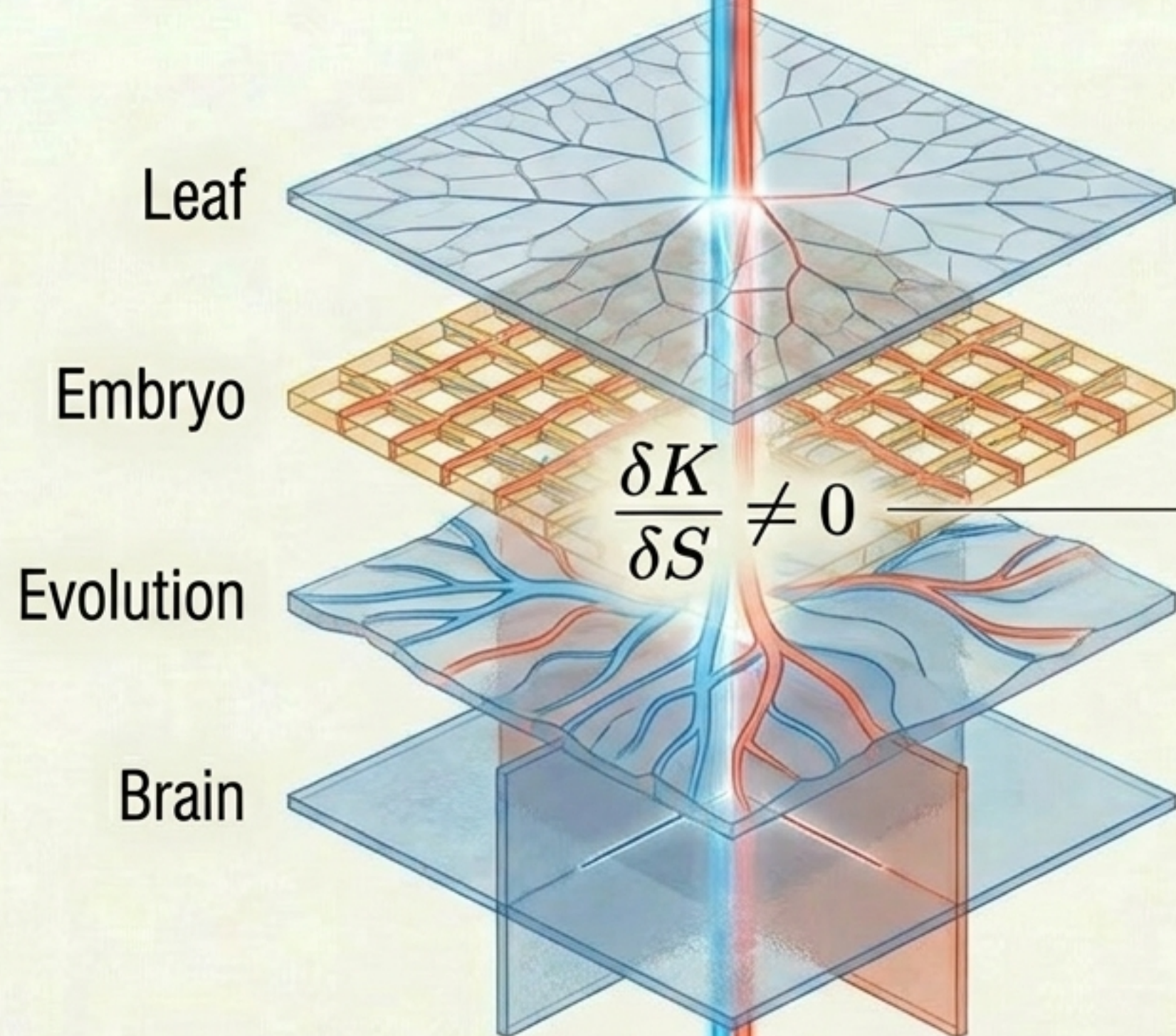
## Transformation, Not Transmission

If subspaces aligned perfectly, rapid learning would overwrite established memories. Rotation forces the information to be computationally transformed, proving rotation is load-bearing, not epiphenomenal.

# The Unified Constraint Geometry Matrix

	Scale I: Spatial (Leaf)	Scale II: Material (Embryo)	Scale III: Regulatory (Evolution)	Scale IV: Dynamical (Brain)
The Signal	Auxin Waves 	Nodal Morphogen 	Transcription Factors 	Neural Population Activity 
The Geometry Modifier	Hydathode Position 	Tissue Rigidity 	IRF1 Accessibility Gate 	Context / Experience 
The Operator	$C_V$ (Partition) 	$C_R$ (Percolation) 	$C_A$ (Regulatory) 	$C_S$ (Projection) 
The Emergent Outcome	Venation Polygons 	Meso-endodermal Boundary 	Species-Specific Cortex 	Selective Info Routing 

# Biological Systems are Self-Modifying Propagation Media



Biological systems **do not compute** by sending signals through fixed wires.

They compute by sending signals that change the nature of the wires themselves.

The propagation **kernel (K)** structurally evolves as a function of the signals (**S**) that pass through it.

# Development as an Evolving Kernel

The Future State

Morphogen concentration,  
gene expression, or  
neural firing rate.


$$S(x, t + \delta t) = \int K_t(x, y) S(y, t) dy$$

The Constraint Geometry Operator  
This is the primary object of biological  
regulation. Development inherently modifies  
the kernel itself, not just the inputs.

The History

The localized signal input  
traversing the medium.

# How to Build a Neocortex



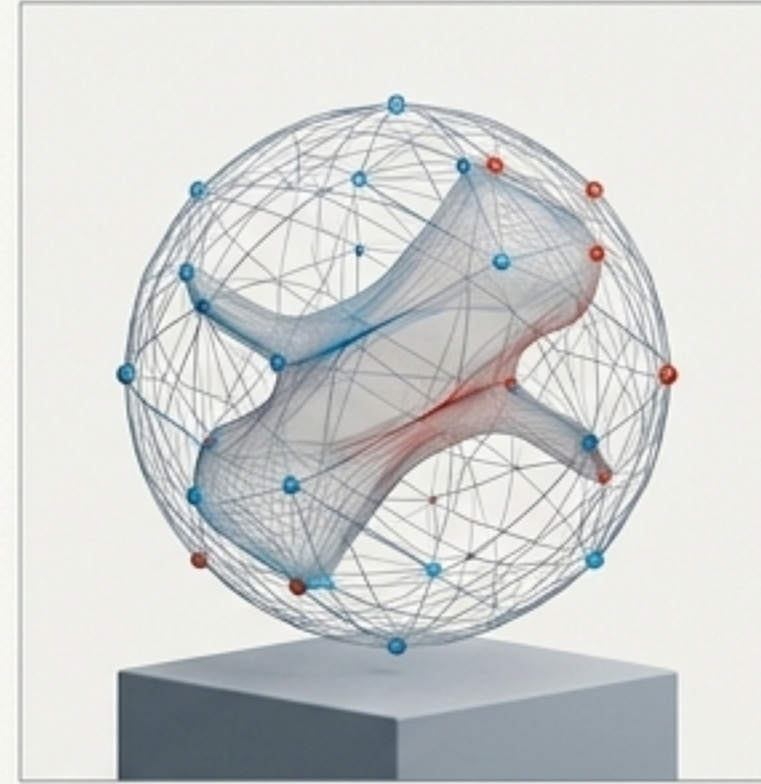
## 1. Regulate the Medium, Not the Outcome

Upstream manipulation of propagation geometry is always more powerful than direct manipulation of downstream signals.



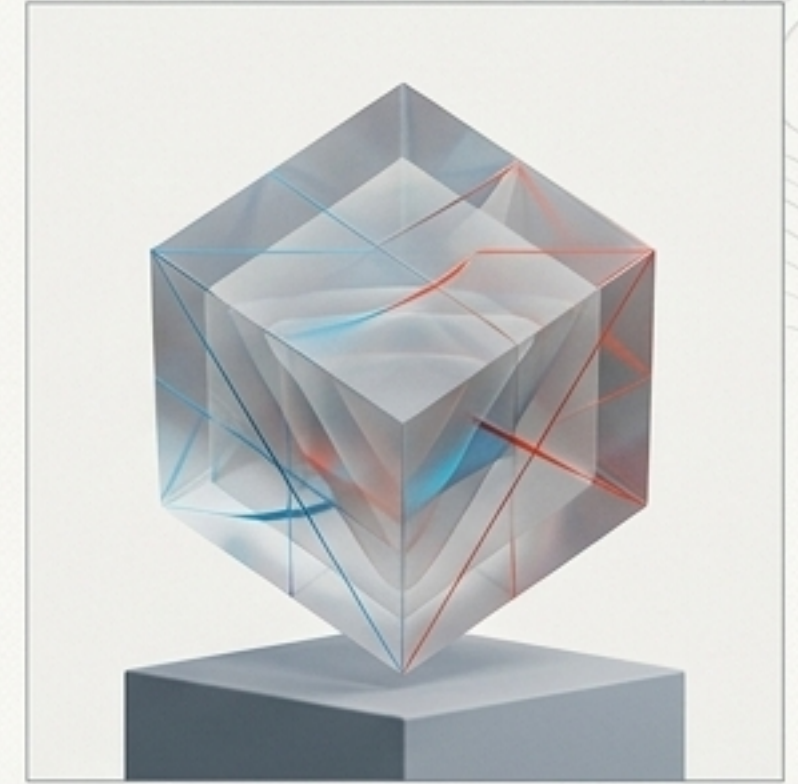
## 2. Unlock Latent Potential

Morphological evolution primarily relies on latent programs waiting for accessibility boundaries to shift, not wholesale molecular novelty.



## 3. Exploit the Edge of Chaos

Maximize informational leverage by maintaining spatial and material tissues near phase-transition critical points.



## 4. Compute via Rotation

Achieve cognitive flexibility not by rewiring anatomical graphs, but by rapidly reorienting communication subspaces.

The neocortex is not a blueprint executed.  
It is a sequence of constraint geometries realized.