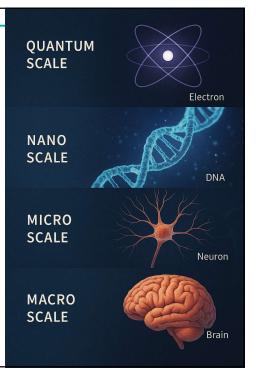




- ☐ Quantum-scale complex-time reps of repeated measurement processes
- Nano-scale
- □ Micro-scale
- Macro-scale
- □ Neuro Data, Brain Networks, MLP & AI
- □ Case-Studies integrated experimental, theoretical, computational & data sciences



### **Multiscale Neuroscience**



Feature	Quantum Scale	Nano Scale	Micro Scale	Macro Scale
Size Range	10 <sup>-15</sup> - 10 <sup>-9</sup> m	10 <sup>-9</sup> - 10 <sup>-7</sup> m	10 <sup>-6</sup> - 10 <sup>-3</sup> m	10 <sup>-3</sup> m and larger
Components	Atoms, ions, subatomic particles	Biomolecules, nanoparticles, nanoscale devices	Neurons, glia, synapses, microcircuits	Brain regions, large- scale networks
Focus	Fundamental quantum phenomena	Molecular interactions, nanoscale tools & structures	Cellular physiology, local circuits	Systems-level function, brain regions & networks
Key Techniques	Quantum modeling, Spectroscopy, Kime-phase tomography (KPT)	Nanofabrication, high-resolution microscopy, nanoscale sensors Cryotomography	Electrophysiology (patch-clamp), optical microscopy, micro-connectomics	fMRI, EEG, MEG, PET, macro- connectomics, lesion studies
Complexity	Lowest (individual particles)	Intermediate (molecules, small devices)	Higher (cells, small circuits)	Highest (brain regions, whole brain)

3

### Quantum Scale: Complex-time (Kime) Reps | VVC | Dare





Using QM principles to model quantum variability

Kime-Phase Simulation - Repeated Spacetime Measurements

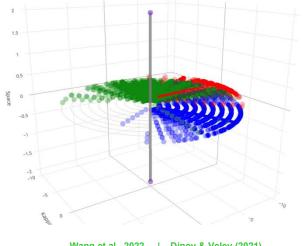
3 Processes - Green, Red and Blue colors (scatter)

At a given spatial location, x,  $\kappa = te^{i\varphi} \in \mathbb{C}$ , where the magnitude (t > 0) is time and the event phase  $\varphi{\sim}\Phi(t)_{[-\pi,\pi)}$  is an angular displacement, event direction, reflecting a random sampling index

1 Fixed spatial location (vertical axis is 1D space)

Repeated IID Measurements colocalized in 4D ST

3 Different Kime-Phase distributions (color-coded) Radial displacement  $t = \underline{\text{time}}$ Angular (**phase**) location  $\varphi \sim \Phi_{[-\pi,\pi)}(t)$ 



Wang et al., 2022 | Dinov & Velev (2021) https://kime.statisticalcomputing.org

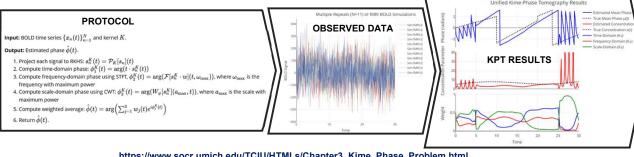
### Quantum Scale: Quantum → Kime-Phase Tomography



**Measurable Observables**: Repeated longitudinal measurements (time-series),  $f_i(t)$ :  $\mathbb{R}^+ \to \mathbb{C}$ ,  $\forall i \in \mathbb{N}$ 

**Representation**: Complex-time (kime) representation,  $\kappa = te^{i\theta} \in \mathbb{C}$ , parameterizing the time-series using event ordering (time,  $t \in \mathbb{R}^+$ ) and random draws from a time t -dependent kime-phase distribution  $\theta \sim \Phi(t)$ 

Problem: The kime-phase is unobservable; its recovery requires indirect kime phase tomography (KPT). Similarly to the quantum mechanical approach for recovering the wavefunction phase, KPT takes repeated measurements in different non-commutative bases and distribution action on kime-test functions



https://www.socr.umich.edu/TCIU/HTMLs/Chapter3\_Kime\_Phase\_Problem.html

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### **Quantum Scale: Complex-time (Kime) Reps**



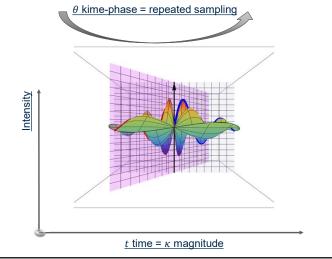


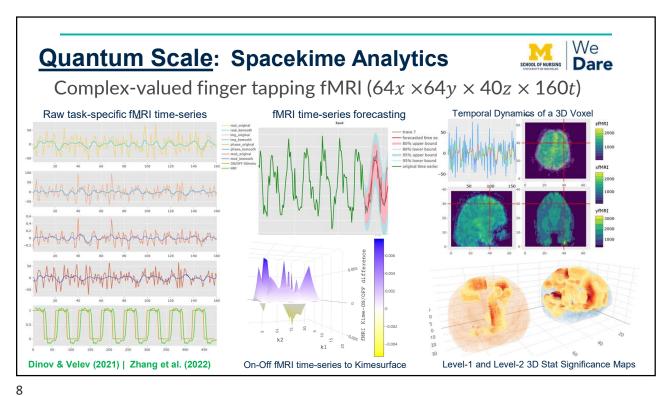
Observable Longitudinal Data Time-Series → Kime-Surfaces (not curves)

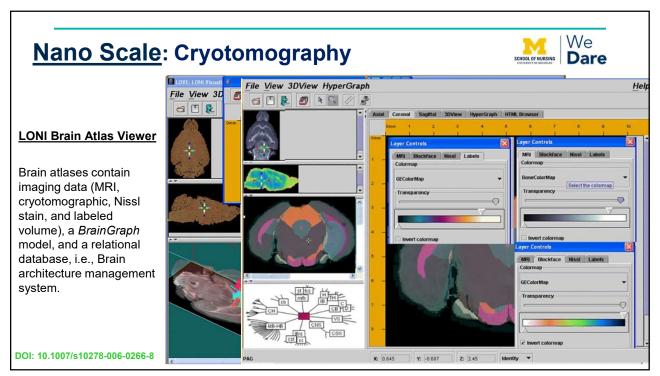
In the 5D spacekime manifold, timeseries curves extend to kime-series, i.e., surfaces parameterized by kimemagnitude (t) and the kime-phase ( $\theta$ ).

Kime-phase aggregating operators that can be used to transform standard timeseries curves to spacekime kimesurfaces, which can be modeled, interpreted, and predicted using advanced spacekime analytics.

Zhang et al., 2022 | Dinov & Velev (2021)







### Micro Scale: Confocal microscopy



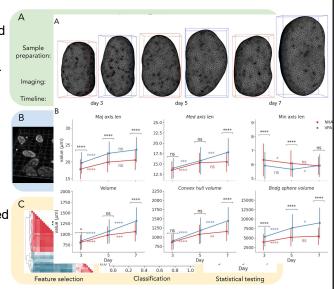
<u>Fig 1</u>: A schematic overview of a confocal microscopy experiment, cancer data collection, and analysis.

- (A) Sample preparation, treatment, and imaging.
- (B) 3D nuclear segmentation, shape modeling, and feature extraction.
- (C) Feature selection, and univariate statistical and machine learning analysis.

#### Fig 2:

- (A) Reconstructed surfaces of representative NHA (normal) and VPA (valproic acid ) treated nuclei
- (B) Time-dependent changes in morphometric measures of nuclear sizes (mean  $\pm SE$ )

DOI: 10.1091/mbc.E20-08-0502



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### **Macro Scale**

Common neuroimaging protocols & computational statistical mapping

Invasive neuroimaging

Functional (in vivo)

Anatomical (ex vivo)

Structural (ex vivo)

Noninvasive neuroimaging

Structural

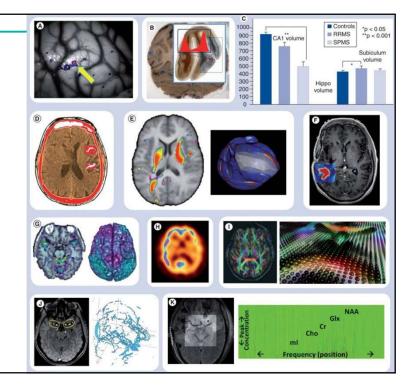
**Functional** 

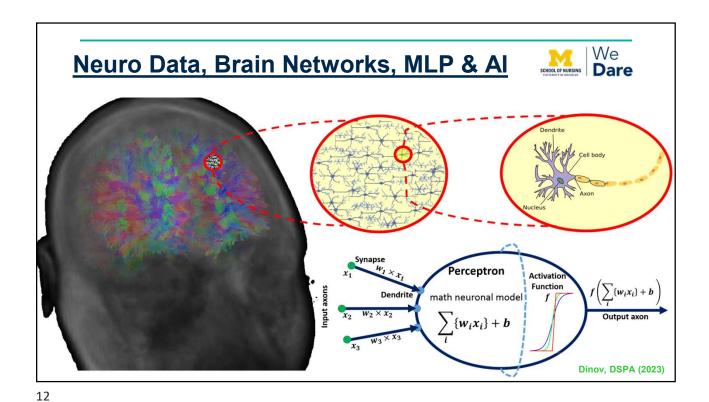
Diffusion

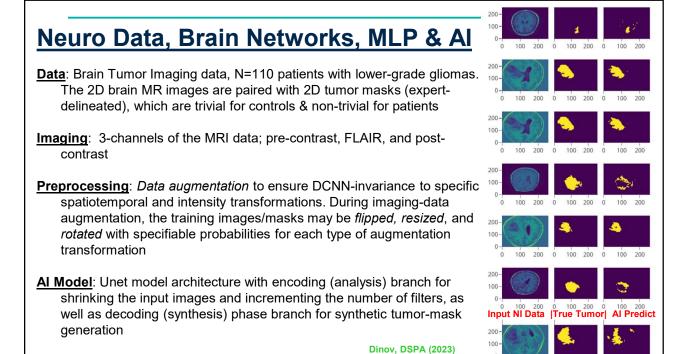
Angiography

Spectroscopy

DOI: 10.2217/iim.11.37







# Cohort Study: Normal & Pathological Aging | We Dare



- □ Problem Model age-related cognition in 3 participant cohorts (1) Asymptomatic Controls, (2) Mild Cognitive Impairment, (3) Dementia
- ☐ Evidence (data types) clinical evaluation (tables), genetic information (sequences), and 3D/4D neuroimaging (spatiotemporal)
- ☐ Status-quo of clinical care independent analysis of the 3 different data types followed by inference pooling
- ☐ Challenge introduce new holistic Health-Analytics Protocol for AI modeling, Dx, classification, and Tx plan using the joint distribution of the entire observed data.

Refs: https://www.socr.umich.edu/people/dinov/publications.html Apps: https://socr.umich.edu/HTML5/ **Input Data** Results Alzheimer's disease dementia rs11257238 ECHDC3 rs6014724 CASS4 rs8093731 SUZ12P1 0.0007

End-to-end Pipeline Workflow

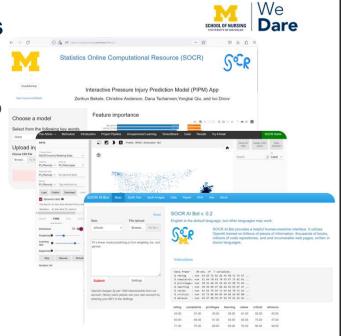
15

3D Neuroimaging

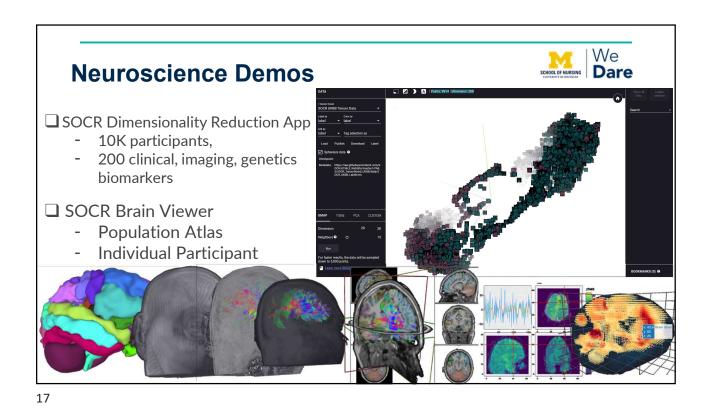
## **Live SOCR AI Webapps**

Genetics

- ☐ Interactive Pressure Injury Prediction Model (PIPM) App (RShiny)
- ☐ Visual Exploratory Data Analytics (SOCR Tensorboard Webapp)
- Quantitative Al-driven Analytics (SOCR AI Bot)



**Neuroimaging-Genetics Associations** 







## **Available Al Resources**

SOCR Motto – "It's Online & Freely Accessible, Therefore it Exists!"		
Pubs:	https://socr.umich.edu/people/dinov/publications.html	
GitHub:	https://github.com/SOCR/PressureInjuryPrediction	
PIPM App:	https://rcompute.nursing.umich.edu/PIPM_v2/	
Al Apps:	https://socr.umich.edu/HTML5/	
SOCR AI Bot	: https://rcompute.nursing.umich.edu/SOCR_AI_Bot/	
Demos:	https://DSPA2.predictive.space (Appendix 9 - OpenAl Synth Text Img & Code)	
<b>Tutorials:</b>	https://TCIU.predictive.space & https://SpaceKime.org	
Websites:	https://nursing.umich.edu & https://socr.umich.edu	