Frontiers in Chronic Pain Assessment: What the Future Holds

National Academies Workshop | SSA Committee

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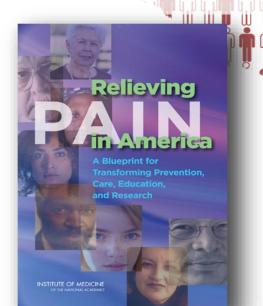
No relevant disclosures No industry COIs

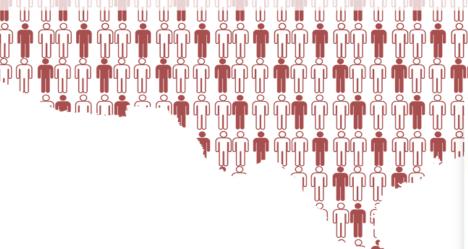
Public Health Crisis of Pain





50M-100M Americans has Chronic Pain 20M have High-Impact Chronic Pain





Centers for Disease Control and Prevention

Weekly / Vol. 67 / No. 24

Morbidity and Mortality Weekly Report
September 14, 2018

Pain Awareness Month — September 2018

September is Pain Awareness Month, when organizations work to raise awareness of how pain affects persons, families, communities, and the nation and to support national action to address pain. A 2011 Institute of Medicine report (https://www.ncbi.nlm.nih.gov/pubmed/22553896) has

Prevalence of Chronic Pain and High-Impact Chronic Pain Among Adults — United States, 2016

James Dahlhamer, PhD¹; Jacqueline Lucas, MPH¹; Carla Zelaya, PhD¹ Richard Nahin, PhD²; Sean Mackey, MD, PhD³; Lynn DeBar, PhD⁴; Robert Kerns, PhD⁵; Michael Von Korff, ScD⁴; Linda Porter, PhD⁶; Charles Helmick, MD⁷.

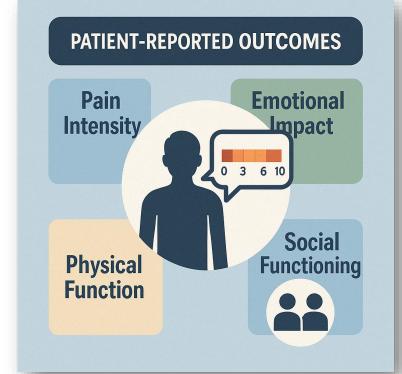
Emerging Tools & Technologies

- Patient Reported Outcomes (PROs)
- Biomarkers
 - Multiomics
 - -Wearables & Continuous Monitoring
 - Quantitative Sensory Testing (QST)
 - Neuroimaging (Brain, Spinal cord, Nerve)
- Learning Health Systems

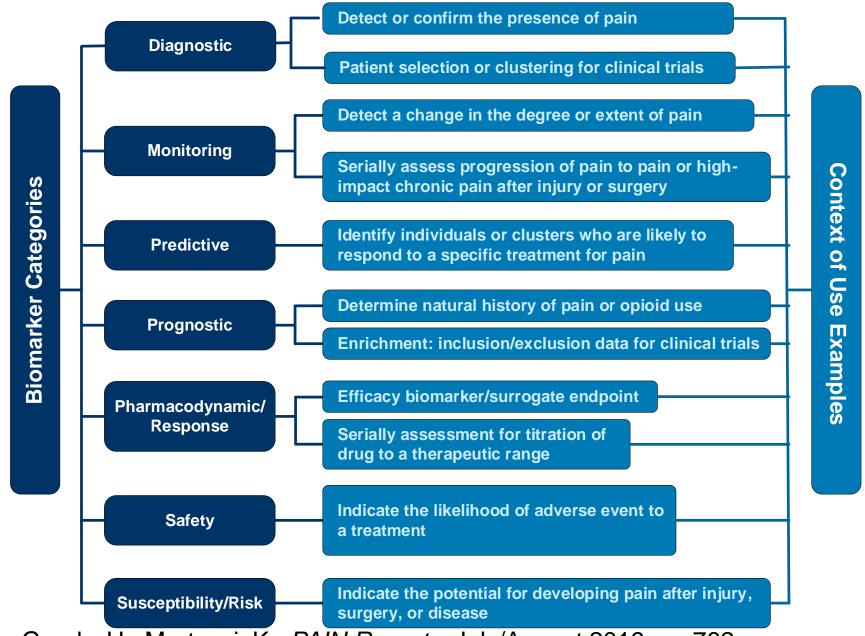
Challenges and Opportunities: Patient-Reported Outcomes

- Traditional 0-10 pain intensity scale overly simplistic with moderate consistency
- Pain as biopsychosocial phenomenon and importance of context
- Unmet Needs:
 - Better capturing daily fluctuations (e.g. experiential momentary assessment)
 - Tools reflecting true lived experiences of patients
 - Integrating patient perspective with objective measures

Foundation for exploring advanced measurement approaches



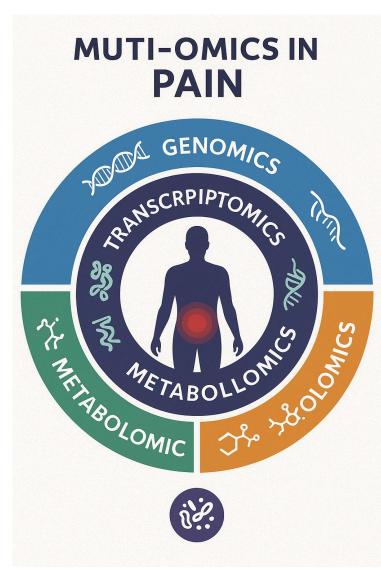
Pain Biomarker Categories and Examples of Use



Mackey, S., Greely, H., Martucci, K., PAIN Reports. July/August 2019, p e762

Biomarkers & Multi-omics – Objective Evidence

- Biomarkers: Objective biological indicators of health or disease
- Multi-omic Biomarkers: Genetics, cytokines, proteomics, metabolomics, lipidomics, microbiome, etc
- No 'omic biomarker yet established as a reliable biomarker of chronic pain
- SSA Implication:
 - Future inclusion in medical records (biomarker panels)
 - Currently cautious interpretations



Wearable Technology – Real-Life Impact

- Track functions: steps, activity, HR, HRV
- Pain-linked changes in activity/sleep patterns
- Objective digital biomarkers of functional impairment

- SSA Implication:
 - Real-world evidence of pain's functional limitations



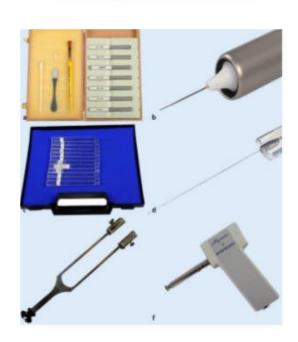
Quantitative Sensory Testing (QST)

- Standardized sensory testing (thresholds, sensitivity)
- Identifies pain mechanisms (neuropathic, inflammatory,

central)

Matches patients to targeted therapies

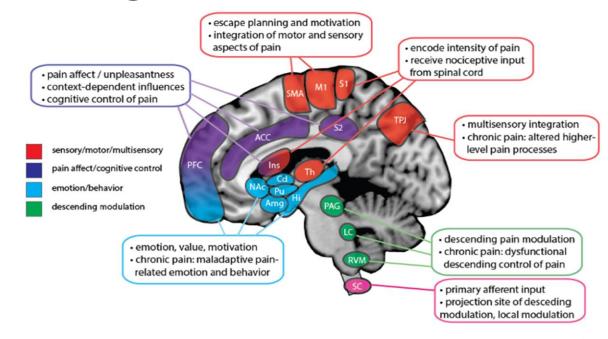
- SSA Implication:
 - Objective evidence of abnormal pain responses



Neuroimaging – Visualizing Pain

- Spinal Cord and Brain MRI/PET: "Pain signatures"
- Peripheral nerve imaging: Identifying pain sources
- EEG

 SSA Implication: Objective pain measures entering medical records



CONSENSUS

OPEN

Brain imaging tests for chronic pain: medical, legal and ethical issues and recommendations

Karen D. Davis^{1,2,3}, Herta Flor⁴, Henry T. Greely⁵, Gian Domenico lannetti⁶, Sean Mackey⁷, Markus Ploner⁸, Amanda Pustilnik^{9,10}, Irene Tracey¹¹, Rolf-Detlef Treede¹² and Tor D. Wager^{13,14}

NATURE REVIEWS | NEUROLOGY

ADVANCE ONLINE PUBLICATION | 1

Need for Composite Multi-Modal Biomarker Signatures

nature reviews neurology



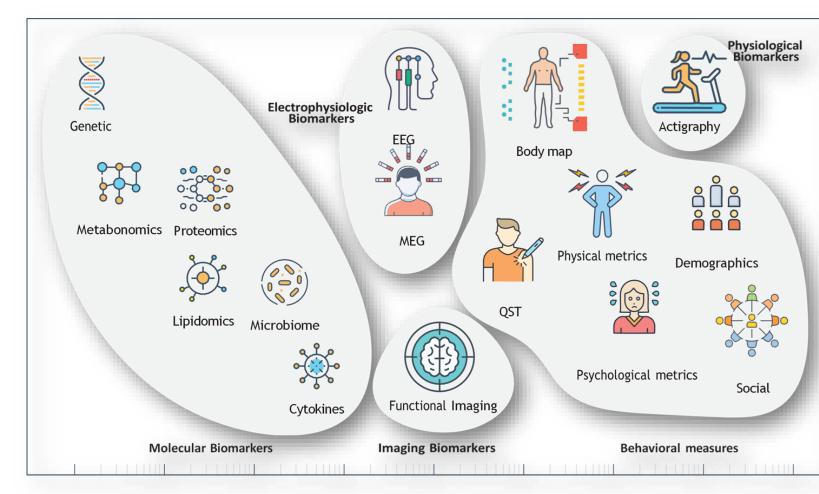
OPEN

Check for updates

Discovery and validation of biomarkers to aid the development of safe and effective pain therapeutics: challenges and opportunities

Karen D. Davis 6.28, Nima Aghaeepour³, Andrew H. Ahn 6.4, Martin S. Angst³,
David Borsook³, Ashley Brenton6, Michael E. Burczynski², Christopher Crean 6.8,
Robert Edwards9, Brice Gaudilliere 6.3, Georgene W. Hergenroeder 6.10,
Michael J. Iadarola11, Smriti Iyengar12, Yunyun Jiang 6.13, Jiang-Ti Kong³, Sean Mackey³,
Carl Y. Saab14, Christine N. Sang13, Joachim Scholz 6.16, Marta Segerdahl17,
Irene Tracey18, Christin Veasley19, Jing Wang20, Tor D. Wager21, Ajay D. Wasan22
and Mary Ann Pelleymounter 6.12

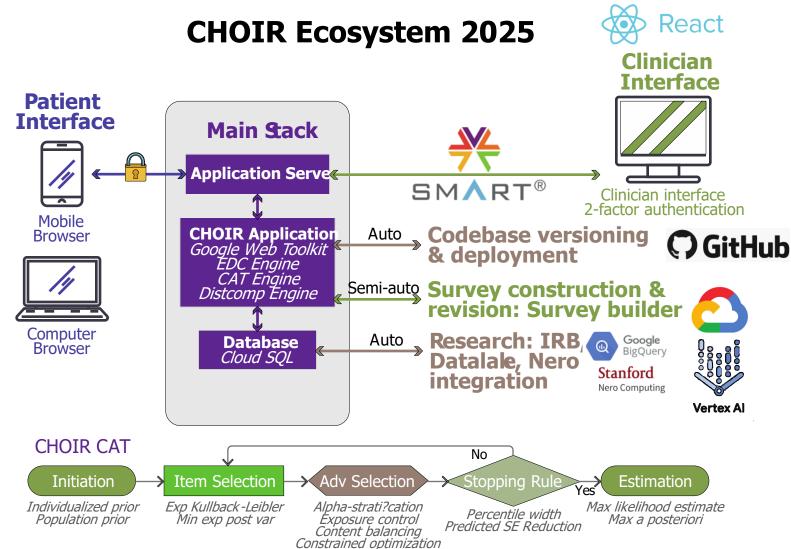
Nature Reviews Neurology 16, 381–400(2020)



Mackey, S., et al, Innovations in Acute and Chronic Pain Biomarkers: Enhancing Diagnosis and Personalized Therapy, *Regional Anesthesia and Pain Medicine* (2024)



Open source, open standard, flexible, and **free** platform for a Learning Health System (http://choir.stanford.edu)



Learning Health System | Example | Continuously Learning | Continuously Learn

Application of precision medicine



Biomarker Signatures Knowledge network

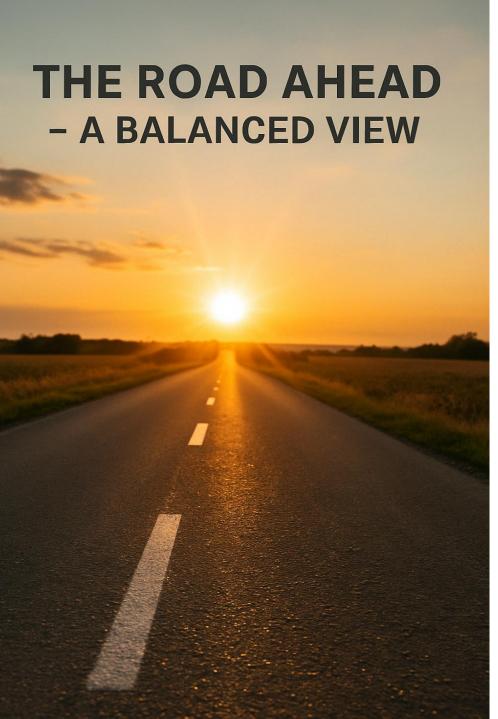


Point of Care: The learning engine

Research influences practice Practice influences research

> Research Studies





- New tools rapidly emerging, promising objectivity
- Many approaches still in research stages
- Future:
 - Multi-modal, objective chronic pain assessment and improved patient reported outcomes
 - Integration into learning health systems to make data actionable
- SSA to monitor validation and clinical acceptance

NIH HEAL grant: "Diagnostic and Prognostic Biomarkers of High-Impact Chronic Pain: Discovery and Validation" (NIH R61NS118651)

