LSU Integrated Musculoskeletal Biobank (LIMB)

Vinod Dasa, MD; Luis Marrero, PhD; Jessica Rivera, MD, PhD; and Jennifer Simkin, PhD



Objectives

- Integration of clinical measures, social parameters, and medical history with blood and tissue samples collected from affected anatomical sites in patients afflicted with musculoskeletal disease
- Answer clinical and basic science questions that account for biological and environmental variables in relation to:
 - Disparities in disease severity and progression
 - Disparities in surgical outcomes
 - ► Effectiveness, efficiency, and safety of novel interventions
 - ▶ Biologicals
 - Surgical techniques
 - ▶ Pre- and post-operative therapeutic strategies



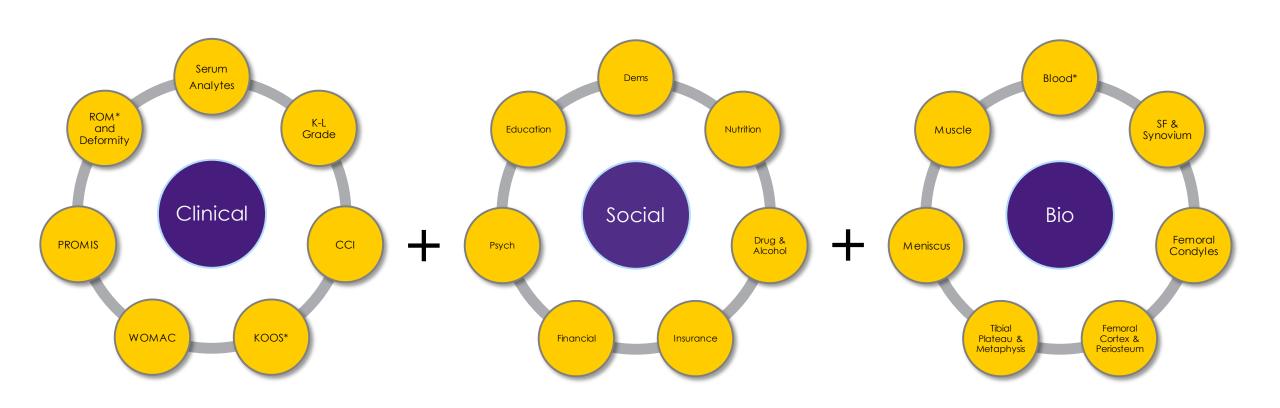
What is the LIMB?

- Central repository of de-identified samples and data currently approved by the IRB for sample collection from patients with osteoarthritis (OA) undergoing total knee arthroplasty (TKA).
- Standardized collection and preservation of most knee components during TKA by fellowship-trained arthroplasty surgeons
- Comprehensive collection of clinical data and validated questionnaires for storage in an encrypted database (RedCAP)
 - Complete demographics and medical records
 - Clinical lab results and metrics of knee function
 - Health indices and surveys
 - Results from assays executed in various research labs





Three comprehensive buckets



* Collected pre- and post-op





Patient Characteristics		Ν
Sex		
	Female	199
	Male	95
Race/Ethnicity		
	Black	107
	White	166
	Hispanic	8
	Asian	2
	Other	11
Age		
	<50	9
	50-65	193
	>65	92
BMI		
	<25	22
	25-35	151
	>35	121
K-L Score		
	1-3	56
	4	238
Deformity		
	Normal	30
	Varus	188
	Valgus	76

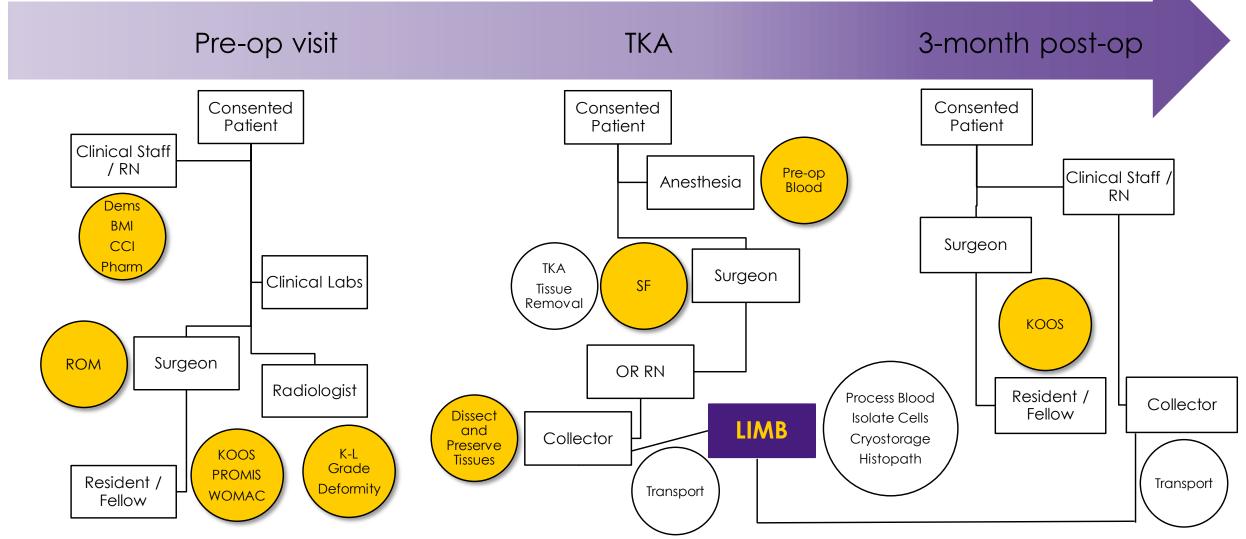
Collection target: 1000 patients







Seamless flow by a dedicated team







Compartment	Sample Type	Status
Pre-op Systemic		
	Blood Serum	Frozen*
	Blood Plasma	Frozen*
	Blood PBMCs	Frozen**
3m Post-op Systemic	:	
	Blood Serum	Frozen*
	Blood Plasma	Frozen*
	Blood PBMCs	Frozen**
Articular		
	Medial Femoral Condyle	Formalin-fixed, Paraffin Block, Frozen*, Frozen**
	Lateral Femoral Condyle	Formalin-fixed, Paraffin Block, Frozen*
	Tibial plateau	Formalin-fixed, Paraffin Block, Frozen*
	Anterior Meniscus	Paraffin Block, Frozen*
	Posterior Meniscus	Paraffin Block, Frozen*
Near Bone Shaft		
	Femoral Cortex	Frozen*
	Femoral Periosteum	Frozen**
	Tibial Metaphysis / Marrow	Frozen**
Joint Capsule		
	Synovial Fluid	Frozen*
	Synovium	Formalin-fixed, Paraffin Block, Frozen*, Frozen**
Intra-articular		
	Articularis Genu	Formalin-fixed, Paraffin Block, Frozen*

Standardized sample processing and storage





^{*}cryopreserved at -135°C in LN_2 (vapor phase) ** cryopreserved in freezing media at -135°C in LN_2 (vapor phase) for tissue culture





Analysis Target	Assay	Output
Cartilage / Synovium / Periosteum	Tissue culture	Cells for profiling and testing biologicals
Tissue / Cells - Pathology		
	H&E stained tissue sections	Semi-quantitative OA severity scores
	Safranin O stained tissue sections	Semi-quantitative OA severity scores
	Picrosirius Red stained tissue sections	Semi-quantitative synovitis scores Fibrosis measurements
	Immunoperoxidase (tissue)	Qualitative protein detection in situ
	Immunofluorescence (tissue/cells)	Qualitative and quantitative protein detection
Tissue / Cells - Microscopy		
	Brightfield	Semi-quantitative
	Deconvolution	Quantitative detection of proteins
	Confocal	Quantitative detection proteins
	Laser Capture Microdissection	Contactless tissue isolation for RNA extraction
Bone		
	μCΤ	Mineral density measures
	Reference point indentation	Strength measures
Synovial Fluid / Serum		
	Protein multiplex against ~17 analytes	Quantitative protein analysis
Tissue / Cells - Molecular		
	RT-PCR	Quantitative gene expression
	Next generation sequencing	Quantitative gene expression
	In-gel near infrared western blot	Quantitative protein expression

Multiple assays and analyses

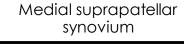


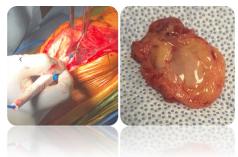


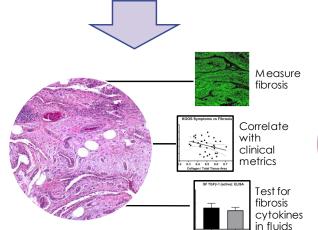




Sample studies



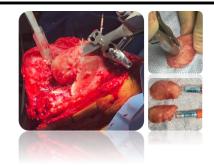


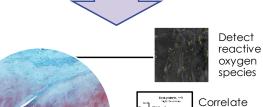


Hodgeson et al. Racial differences in severity of synovial fibrosis in patients with knee osteoarthritis. Submitted to JOR 10/2020.

Synovitis scoring

Femoral condyles





Oxygen species

Oxygen species

Correlate with signaling molecules

Profile and test

cellsin

vitro

Mix et al. Oxidative stress induces nuclear

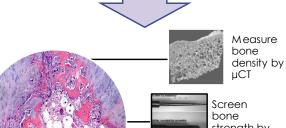
receptor 4A2 in human articular

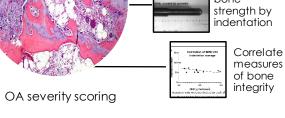
chondrocytes. OA and Cartilage. 2019

OA severity scoring

Tibial plateau

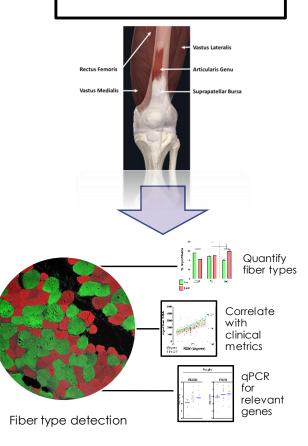






Valentino et al. Is Poor Bone Quality Associated with Pain after Total Knee Arthroplasty (TKA)? ORS 2020

Intra-articular muscle



Crawford et al. Using the Articularis Genu to test peri-articular muscle health during knee osteoarthritis. ORS 2021





533 Bolivar St.
Clinical Sciences Research Bldg., 5th floor
New Orleans, LA 70112
Phone: 504-568-2597
E-mail: lmarre@lsuhsc.edu

https://www.medschool.lsuhsc.edu/ortho/mus culoskeletal_sample_repository.aspx





