Breakout Session 2: Track A

Enabling AI/ML Readiness and Modernization of Longitudinal Pregnancy and Cardiovascular Health Data: Lessons Learned

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Enabling AI/ML Readiness and Modernization of Longitudinal Pregnancy and Cardiovascular Health Data

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Project summary



Overview of study timeline



nuMoM2b

AIMS – Nulliparous Pregnancy Outcomes Study: Monitoring Mothers-to-be

- 1. Determine maternal characteristics, including genetics, epigenetics, and physiological response to pregnancy as well as environmental factors that influence and/or predict adverse pregnancy outcomes (APOs)
- 2. Identify specific aspects of placental development and function that lead to APOs
- 3. Characterize genetic, growth, and developmental parameters of the fetus that are associated with APOs

- Sponsor: NICHD
- Multi-site prospective cohort study
- 10,038 nulliparas with singleton pregnancies were enrolled at $6 13^{6/7}$ weeks gestation
 - Enrollment 2010-2013; deliveries 2010-2014
- Age at enrollment: 13-45 years
- Participants were followed through 14 days postpartum and newborn infants were followed through hospital discharge
- Study visits at up to 4 timepoints:
 - ▶ 6⁰ 13⁶
 - ► 16⁰ 21⁶
 - ► 22⁰ 29⁶
 - Delivery



Pregnancy: a window to future cardiovascular health?

Prior studies: association between APOs (PTB, SGA, GDM) and maternal cardiovascular disease later in life

- Some APOs have underlying vascular mechanisms
- Is there a common cause?
- Is pregnancy a "stress test" that might reveal cardiovascular risk?





nuMoM2b Heart Health Study (HHS1)

- Goal: Evaluate the association between APOs and maternal health later in life in an effort to develop future screening and preventive strategies.
- Funded jointly by NHLBI and NICHD
- Continued follow-up (2014-2020) of nuMoM2b participants who agreed to recontact, with:
 - Known pregnancy outcomes
 - Age 18 years or more at time of first recontact
 - (In-person visit) Not currently pregnant and at least 6 months postpartum
- Interval contacts (phone or online survey) every 6 months until in-person visit, then at 12-month intervals
- In-person visit at 2 or more years after nuMoM2b pregnancy
- In-home sleep breathing substudy after the in-person visit (for nuMoM2b sleep breathing study participants)
- Chart abstraction for subsequent pregnancies if APO or multiple gestation pregnancy was reported
- · Chart review for reports of CV-related event or death
- Central laboratory analysis of biospecimens: lipids, glucose, HbA1c, insulin, hsCRP, NT-proBNP, urine albumin & creatinine



Domain	Instrument or Clinical Measure	Dataset by Study Visit					
		1 st	2 nd	3 rd	D	IC	IP
Medical and sociodemographic history	Medical and Sociodemographic History	V1A*	V2A*	V3A*	V4A	Txx*	V5 <u>A,V</u> 5S
	Lactation History					Txx*	
	Family Medical History		V2A				V5A
	Residential History	V1A*	V2A*	V3A*			V5A
	Medical Record Abstraction: Pregnancy/Neonatal				Cxx*	P5A-F*	P5A-F*
	Medical Record Abstractions: CVD events				CMA,CMB	CRX,CRM	CRX,CR
Physical activity	Physical Activity Log (BRFSS)	V1A*	V2A*	V3A*			V5A
	Modifiable Activity Questionnaire (MAQ)						V5Q*
Nutrition	Modified Block FFQ	V1D*					V5D*
	Three Factor Eating Questionnaire - Disinhibition						V5P
Health literacy	Rapid Estimate of Adult Literacy (REALM-SF)		V2A				
Psychosocial	Experiences of Discrimination (EOD)		V2A				
	Reaction to Race	V1A					
	Multidimensional Social Support (MSPSS)	V1G					
	Pregnancy Experience Scale, Brief (PES-Brief)			V3J			
	Connor-Davidson Resilience Scale (CD-RISC)		V2I				
Stress & mental health	Perceived Stress Scale (PSS-10)	V1A		V3A			V5R
	Edinburgh Postnatal Depression Scale (EPDS)	V1C		V3C			V5C
	State-trait Anxiety Index (Trait) (STAI-T)	V1H					
Sleep	Sleep Breathing Substudy	V1K		V3K			V5N
	Sleep Patterns and Quality Substudy		V2M				
	WHI Insomnia Rating Scale	V1F/V1L		V3F/V3L			V5N
	Berlin Questionnaire for Sleep Apnea	V1F/V1L		V3F/V3L			V50
	Epworth Sleepiness Scale, and Restless Legs	V1F/V1L		V3F/V3L			V5N
	Restless Legs Syndrome Diagnostic Criteria	V1F/V1L		V3F/V3L			V5N
	PROMIS Sleep Disturbance & Impairment						V5N
Clinical measures	Blood Pressure	V1B*	V2B*	V3B*			V5B*
	Pulse						V5B*
	Weight	V1B*	V2B*	V3B*			V5B*
	Height	V1B*					V5B*
	Waist, Hip, and Neck Circumference	V1B*					V5B*

Overview of data collection across nuMoM2b & HHS1

* Ancillary datasets contain derived content from these forms and from biospecimen assays and electronically captured data

Enabling AI/ML Readiness ... Aims

- 1. To convert existing epidemiologic datasets to machine-readable structures with complete metadata, with modernized documentation, and to provide access to the resulting data and materials on BioData Catalyst.
- 2. To complete the preparation and submission of currently disparate multi-omics datasets to dbGaP and BioData Catalyst.
- 3. To expand awareness and improve usability of the AI/ML ready data by creating a template workflow for the application of ML tools and completing analyses for a clinical use case that will be made available as a public markdown project.



Challenges



Enabling AI/ML Readiness ... Aim 1

Aim 1: To convert existing epidemiologic datasets to machine-readable structures with complete metadata, with modernized documentation, and to provide access to the resulting data and materials on BioData Catalyst.

- Machine-readability: Convert existing files (130+) from SAS datasets to JSON format
 - In existing data, missing data codings were applied inconsistently and many variables were not labeled. Extensive work to update prior to JSON conversion, still underway
- Harmonization: assign clinical ontologies to clinical characteristics and endpoints
 (obstetric and cardiovascular)
 - Deferred until addition progress made on machine-readability
- Documentation modernization: Convert current documentation (300+ page word document + supporting metadata) to file-specific datasheets
 - Proceeding well, but time-consuming



Enabling AI/ML Readiness ... Aim 2

Aim 2: To complete the preparation and submission of currently disparate multi-omics datasets to dbGaP and BioData Catalyst.

- Data sharing: submission of GWAS, exosome and plasma proteomics, WGS, and RNASeq data and/or datasheets for public access on BioData Catalyst
 - TOPMed data (WGS, RNAseq) delayed
 - Technical issues with dbGaP accessions delayed submission of GWAS data
- Data interoperability: support multi-omic analyses by performing common group comparisons and storing results under common metadata structures
 - Deferred start on this
- Data harmonization: create phenotype datasets using new metadata standards used by other major cardiovascular health cohort studies
 - Deferred start on this



Enabling AI/ML Readiness ... Aim 3

Aim 3. To expand awareness and improve usability of the AI/ML ready data by creating a template workflow for the application of ML tools and completing analyses for a clinical use case that will be made available as a public markdown project.

All Aim 3 activities pending additional progress on JSON dataset creation and omics data submissions.

- Template workflow: create template for basic ML workflow and analysis using commonly used cohort datasets
- Data paper: submit a data paper describing the shared datasets and project information
- Use case: complete analysis for a clinically relevant application of the AI/ML ready datasets and make it available as a public markdown project



Future work



Future work

- Complete existing planned work
- Apply processes to new datasets to prepare them for public use
- Update / maintain new resources as appropriate



