

Monocyte Anisocytosis: The Future of Screening for Sepsis in Children

Lael Yonker, MD
Pediatric Pulmonary
Massachusetts General Hospital
Boston, MA

Disclosures

- Funded by Human Health Services(HHS) / Office of Assistant Secretary for Preparedness and Response (ASPR) / Biomedical Advanced Research and Development Authority (BARDA), cost share with Beckman Coulter

Background



- 30 million children evaluated in the Emergency Department annually
- Top cause: fever/signs of infection
- Sepsis is rare but potentially devastating outcome

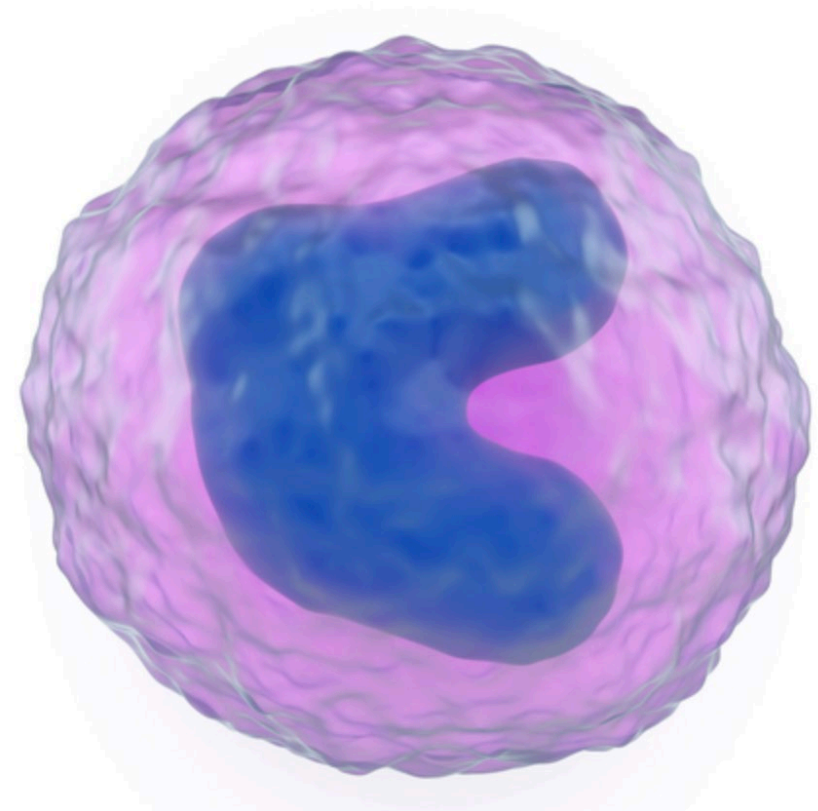
Background



- Clinical evaluations can screen for children at risk of sepsis
- Variability in clinical assessments:
 - Fearful child
 - Provider variability/interpretation
 - Comfort of provider with pediatric patients
- Work up for sepsis/infection is variable
- Reliable biomarkers are needed

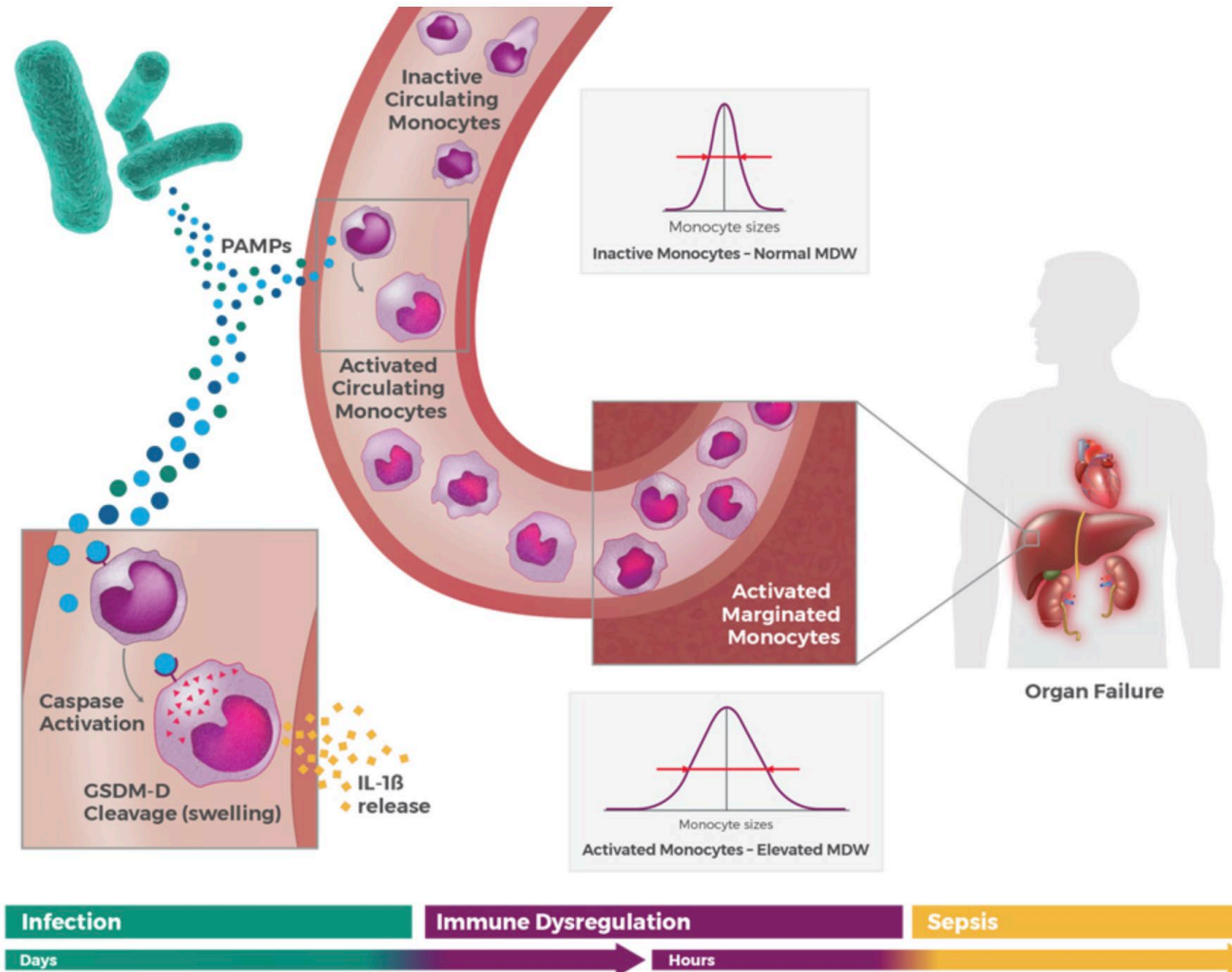
The Monocyte

- Key role in antigen presentation
- Research shows severe sepsis has a shift in monocyte profiles by flow cytometry / gene expression
- Monocytes release multiple cytokines associated with severe inflammation

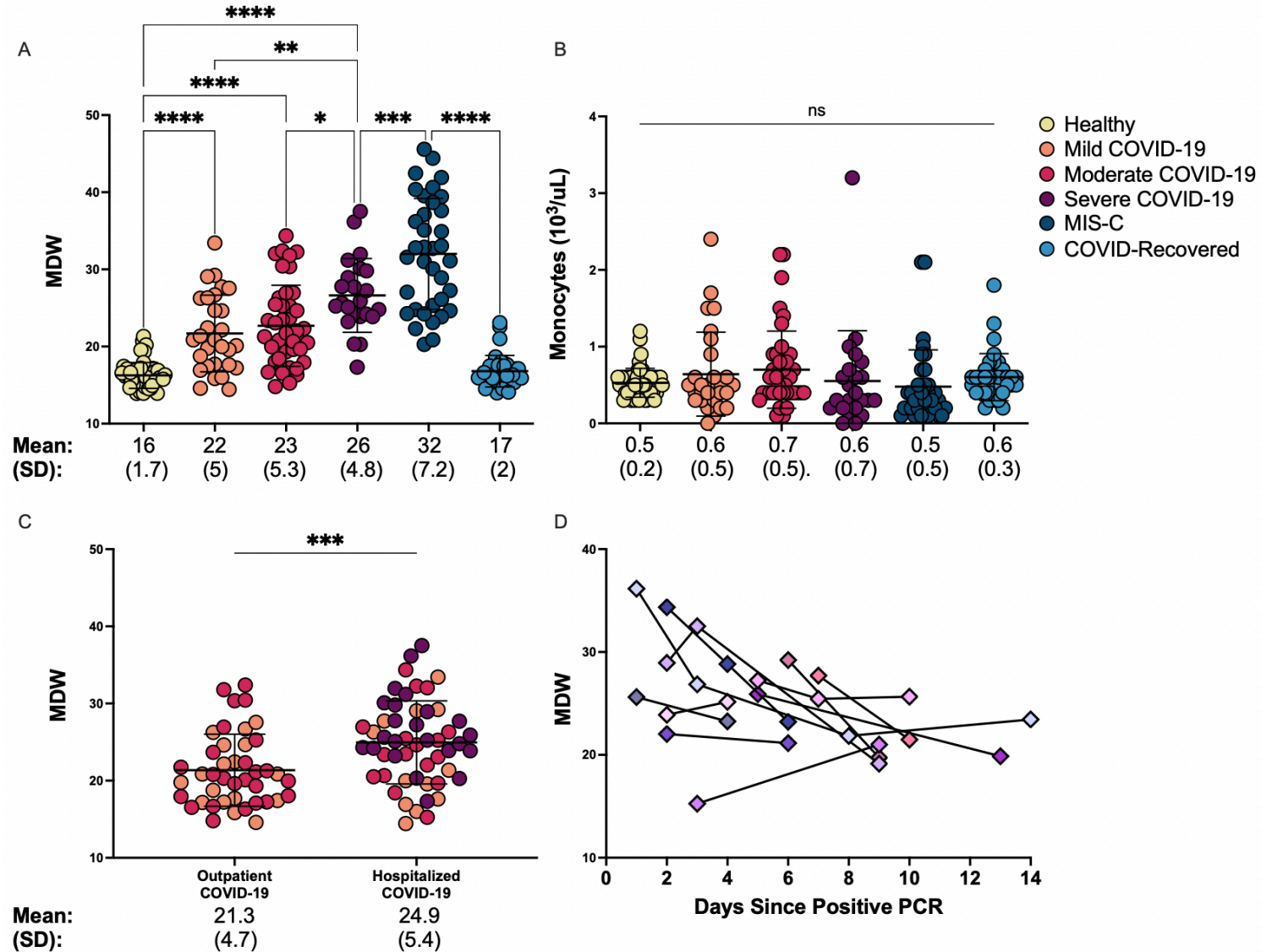


What is monocyte anisocytosis?

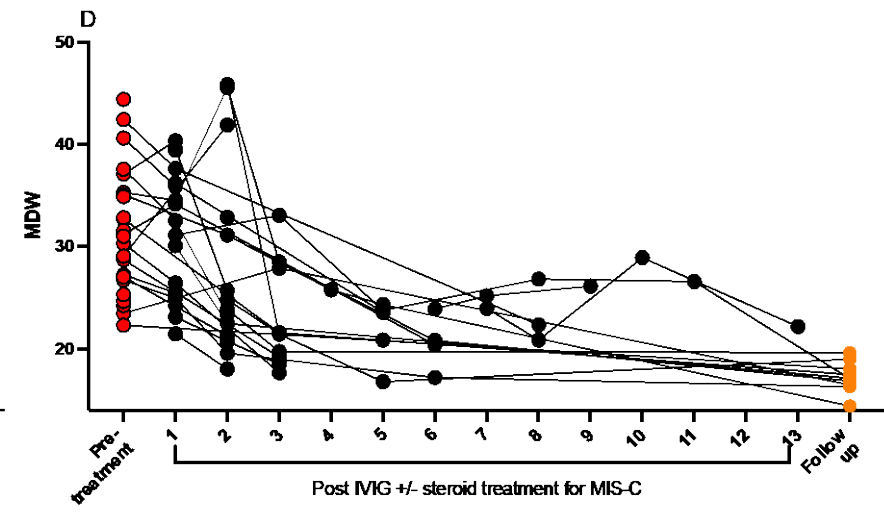
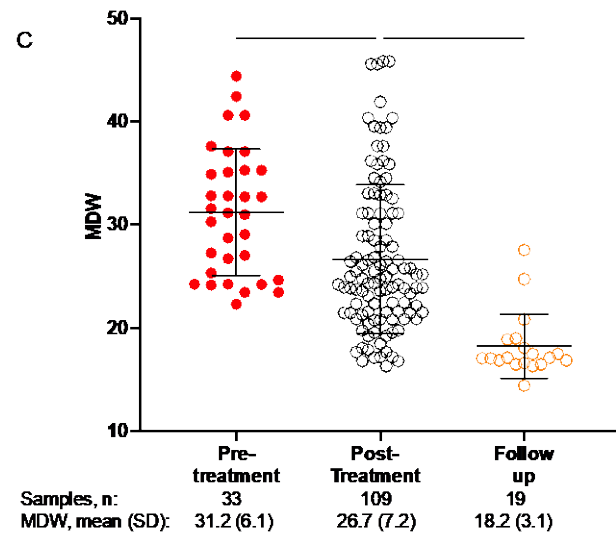
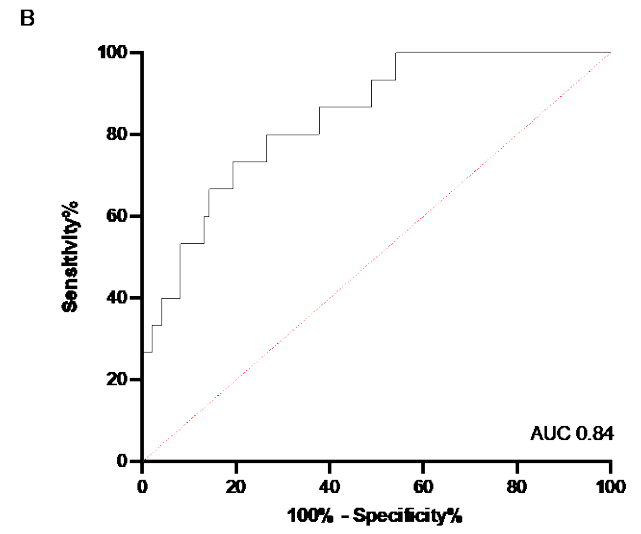
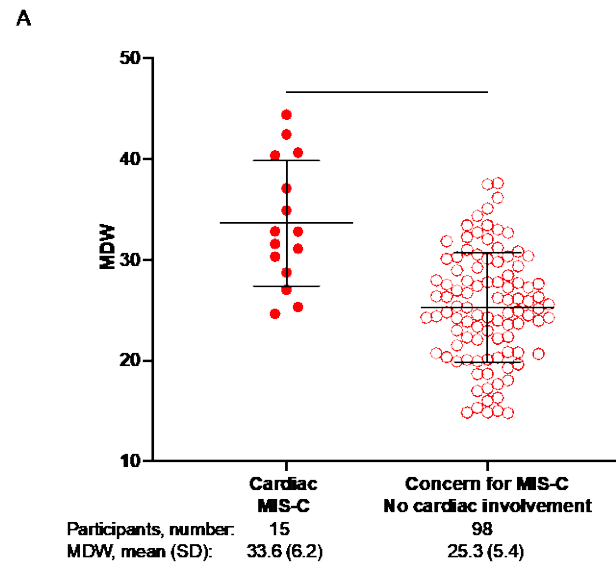
Monocyte distribution width (MDW)



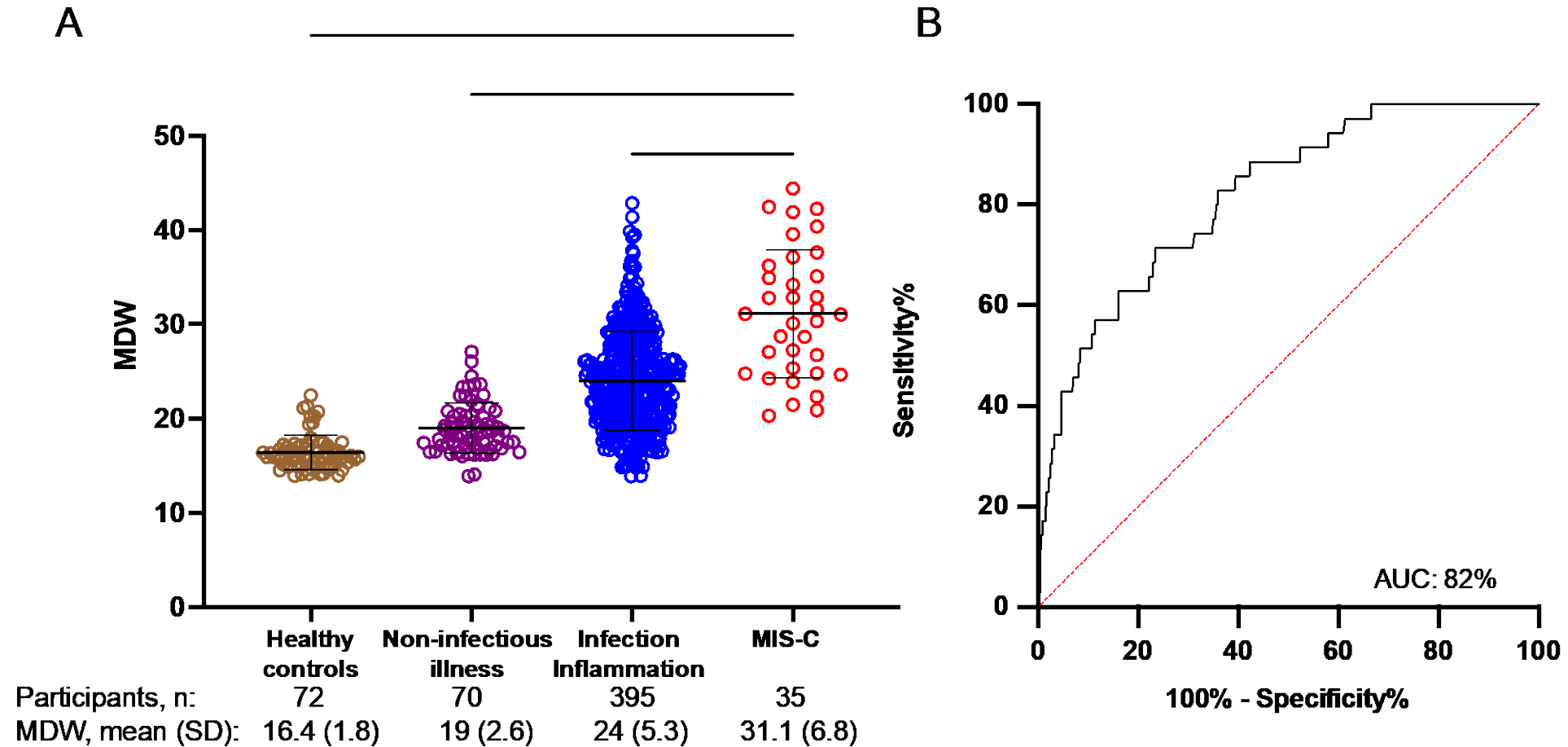
Monocyte anisocytosis is increased with severity of COVID-19 in children



Monocyte anisocytosis is increased in MIS-C, especially with cardiac involvement



Monocyte anisocytosis is increased in MIS-C *but* MDW is broadly elevated in infection and inflammation



Does MDW correspond more broadly with sepsis?

- Retrospective data
- 2 clinical sites (Johns Hopkins and MGH)
- Assigned sepsis 2 scores (Goldstein 2005)
- Assigned pSOFA scores

Total enrolled (N=394)	Pediatric Sepsis (n=110)	Non-septic pediatric patients (n=284)
Age, years: mean (SD)	7.5 (5.5)	8.7 (5.6)
Sex at birth		
Female, n (%)	54 (49)	131 (46)
Male, n (%)	56 (51)	153 (54)
Race		
White	48 (43)	142 (50)
Black	23 (21)	55 (19)
Asian	7 (6)	11 (4)
Other	34 (31)	80 (28)
Ethnicity		
Hispanic	30 (27)	85 (30)
Non-Hispanic/Unknown	80 (72)	199 (70)

Conclusion

- MDW is a biomarker that is clinically accessible on hematology analyzers
- Monocyte anisocytosis is strongly associated with sepsis in children
- MDW may serve as a useful, objective tool in screening for children at risk of sepsis

Thank you!



Team:

Alessio Fasano
Lauren Guthrie
Rosie Lima
Bryan Alvarez
Yanki Okuduco
Madeleine Godfrey

Collaborators:

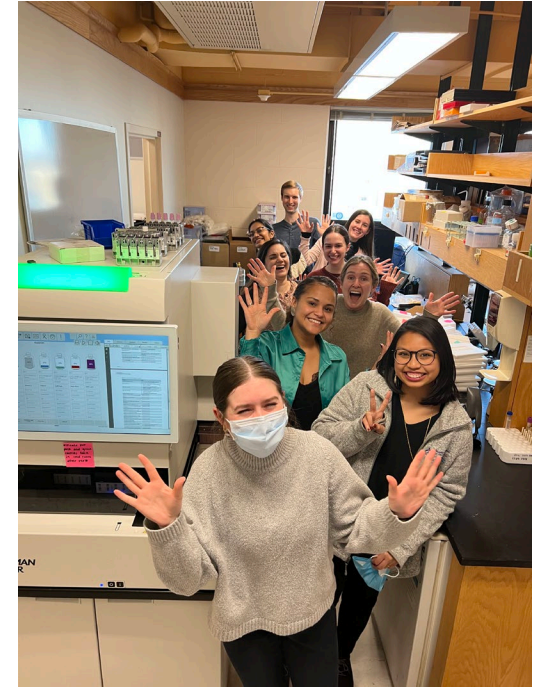
Daniel Irimia
Kemi Badaki
Kent Lewandrowski
John Higgins

Lab Alumni

Denis De la Flor
Eva Farkas
Mimi Burns
Abby Kane
Jameson Davis
Maggie Loiselle
Allison Fialokowski
Brittany Boribong
Matthew Dunn

Funding:

NIH/NHLBI
ASPR/BARDA
Polybio Research



X/Twitter: @LaelYonker

Email: lyonker@mgh.harvard.edu