

60th Friend E. Clark Lecture Series



sponsored by

Phi Lambda Upsilon Honorary Chemistry Society and
C. Eugene Bennett Department of Chemistry



Vicki Wysocki, Ph. D

*Ohio Eminent Scholar of Macromolecular
Structure and Function*

*Department of Chemistry and Biochemistry
Director, Campus Chemical Instrument Center
The Ohio State University*

**John Fenn made molecular
elephants fly and changed
biomedical research**

Where: Clark 312

When: Monday, March 4th at 3:30pm – 4:30pm

**Native mass spectrometry: A
structural biology tool**

Where: Clark 112

When: Tuesday, March 5th at 3:30pm – 4:30pm

Graduate students, please join us for
pizza with Dr. Wysocki

Where: CLC

When: Monday, March 4th at 12pm to 1:15pm



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Until the late 1980's, mass spectrometry was a useful measurement tool for measuring relatively small molecules. With the invention of electrospray ionization, mass spectrometry became a tool that can measure non-volatile, thermally unstable molecules and the use of mass spectrometry as a characterization tool changed dramatically. Today, mass spectrometry is a \$6 billion industry, is used for many applications, and new developments continue. This talk will illustrate several types of research that have been made possible because of improvements in ionization and in mass spectrometry instrument development. Examples will be selected from multiple fields of research including proteomics, metabolomics/gut microbiome, and native mass spectrometry of large protein complexes, including gene therapy platforms.

Native mass spectrometry: A structural biology tool

Where: Clark 112

When: Tuesday, March 5th at 3:30pm – 4:30pm

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Characterization of the overall topology and inter-subunit contacts of protein complexes, and their assembly/ disassembly and unfolding pathways, is critical because protein complexes regulate key biological processes, including processes important in understanding and controlling disease. Tools to address structural biology problems continue to improve. Native mass

spectrometry (nMS) and associated technologies such as ion mobility and variable temperature electrospray ionization are becoming increasingly important components of the structural biology toolbox. When the native mass spectrometry approach is used early or mid-course in a structural characterization project, it can provide answers quickly for small amounts of sample, even when heterogeneity is present. Integration of sample preparation/purification with effective dissociation methods (e.g., surface-induced dissociation, SID), ion mobility, and computational approaches provide an MS workflow that is enabling in biochemical, synthetic biology, and systems biology approaches. Native MS can determine whether a complex of interest exists in a single or multiple oligomeric states, and surface induced dissociation can provide characterization of topology/inter-subunit connectivity and other structural features. Examples will illustrate the coupling of SID to electron capture charge reduction and charge detection mass spectrometry for the characterization of protein and nucleoprotein complexes, including glycoproteins and adeno-associated virus capsids.

References

- [1] Snyder, Harvey, Wysocki, V. H., *Chemical Reviews*, 2022, 122, 8, 7442–7487.
- [2] Karch, Snyder, Harvey, Wysocki *Annu. Rev. Biophys.* 2022. 51, 157–79.