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Special Seminar Invitation

Probing protein structural changes on a proteome-wide scale

Speaker	Prof. Dr. Paola Picotti
	Institute of Molecular Systems Biology ETHZ

When May 9, 14.00 - 15.00

Where Forum Chriesbach, room D24, Eawag Dübendorf

Abstract Protein structural changes induced by external perturbations or internal cues can profoundly influence protein activity and thus modulate cellular physiology. Mass spectrometry (MS)-based proteomic techniques are routinely used to measure changes in protein abundance, post-translational modification and protein interactors, but much less is known about protein structural changes. In my talk, I will present a recently developed structural proteomics method that enables analysis of protein structural changes on a proteome-wide scale and directly in complex biological extracts. The approach relies on the coupling of limited proteolysis (LiP) tools and MS. LiP-MS can detect subtle alterations in secondary structure content, larger scale movements such as domain motions, and more pronounced transitions such as the switch between folded and unfolded states. I will describe different applications of the approach, including the unbiased identification of protein-small molecule interactions and the monitoring of amyloidogenic proteins involved in neurodegenerative diseases. I will describe how we recently used the approach to build a map of protein-metabolite interactions and binding sites in E. coli and how this enabled the unbiased identification of novel allosteric events. Last, I will discuss the power and limitations of the new approach.