

**Exploring the Ub/UBL landscape with activity-based probes** Witting, K.F.

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## Stellingen

- 1. Site-specific introduction of unnatural amino acids during protein translation to mimic posttranslational modifications is not trivial. It requires to go against Nature's original design. (this thesis)
- 2. A post-translational modification such as ubiquitination is much more than just a mark for degradation—it's a complex signaling language that determines virtually all aspects of cellular function and homeostasis. *(this thesis)*
- 3. Probing dynamic post-translational modifications is a challenging undertaking. Not only are these modifications temporally and spatially altered, but also react to changes in the proteome itself and the transcriptome. *(this thesis)*
- 4. Ribosomes are complex and adaptive to cellular protein synthesis demands. Decoration of the large ribosomal subunit in a critical position allows different biological outcomes. *(this thesis)*
- 5. Ribosomal proteins have more functions beyond protein synthesis—they orchestrate a complex choreography regulating ribosomal function and thus cellular homeostasis.
- 6. The plasticity of the proteome is an adaptation to changes in the cellular environment. (adapted from Kim *et al.*, 2011, Molecular Cell, p. 325).
- 7. Deeper understanding of cell biology often necessitates the use of chemistry by enabling the visualization and manipulation of cellular processes.
- Low abundance of a protein or a post-translational modification does not mean that it is insignificant—usually these proteins are highly specialized and expand the depth of biological insights immensely.
- 9. Dissecting the differences between similar studies should inspire other studies that illustrate the context-dependent subtleties of cellular processes.
- 10. Observation and accurate measurement of a scientific phenomenon is a conundrum manipulation of one factor in a biological system will lead to inevitable manipulation of another related factor. Thus, the "truth" depends entirely on the observer. (adapted from Erwin Schrödinger's and Eugene Wigner's thought experiments)
- 11. Groundbreaking scientific insights are only gained when brilliant minds share the same vision, collaborate, and remain engaged.