

Andrea Bonini

Department of Chemistry “Ugo Schiff”, University of Florence, Florence (FI), Italy
a.bonini@rug.nl

Following the successful application of biological nanopores in DNA sequencing, nanopore-based technologies are now emerging as powerful analytical tools for observing and sensing proteins at the single-molecule level, with the potential to become the first method capable of sequencing full-length proteins one molecule at a time [1,2].

After years of dedicated research, biological nanopores are now beginning to overcome many of the longstanding challenges associated with protein analysis. This progress is opening the door to a wide range of new applications in chemistry, biology, and medicine [3–6].

This tutorial introduces the fundamental principles of biological nanopores and explains how they function as nanoconfined spaces to study biomolecules at the single-molecule level [7]. It examines the key differences and current challenges in analyzing proteins compared to DNA and provides an overview of the expanding applications of nanopore-based methods, from DNA sequencing to the emerging frontier of protein detection and sequencing [8].

References

- [1] Bayley, H., Cremer, P., *Nature*, 413 (2001) 226–230
- [2] Dorey, A., Howorka, S, *Nat. Chem*, 16 (2024) 314-334
- [3] Bonini, A., Sauciuc, A. & Maglia, G. *Nat Methods*, 21 (2024) 16–17.
- [4] Sauciuc, A., Morozzo della Rocca, B., Tadema, M.J. et al. *Nat Biotechnol*. 42 (2024) 16-17.
- [5] Vreeker, E., Grünewald, F., van der Heide, N. J., Bonini, A., Marrink, S. J., Tych, K., & Maglia, G. *Advanced Materials*, 37 (2025).
- [6] Sauciuc, A., Whittaker, J., Tadema, M., Tych, K., Guskov, A., & Maglia, G. *Proceedings of the National Academy of Sciences*. 121 (2024).
- [7] Maglia, G. *Methods in enzymology*, 475 (2010) 591-623
- [8] Lu, C., Bonini, A., Viel, J. H., & Maglia, G. *Nature biotechnology*. 43(3) (2025) 312-322