

# Protein discrimination using aptamer-functionalized nanopores

**C, Raillon<sup>1\*</sup>**

Lucile Reynaud<sup>1</sup>, Aurélie Bouchet-Spinelli<sup>1</sup>, Jean-Marc Janot<sup>2</sup>, Arnaud Buhot<sup>1</sup>, Sébastien Balme<sup>2\*</sup>

*1 University Grenoble Alpes, CEA, CNRS, INAC-SyMMES, 17 Rue des Martyrs, 38000 Grenoble, France*

*2 IEM, Institut Européen des Membranes, UMR 5635 Université Montpellier, CNRS, ENSCM, Place Eugene Bataillon, F-34095 Montpellier cedex 5, France*

Correspondence: [camille.raillon@cea.fr](mailto:camille.raillon@cea.fr)

## Abstract

Protein detection and identification at the single-molecule level is a major challenge in many biotechnological fields. Solid-state nanopores have raised attention as label-free biosensors with high sensitivity. Here, we use solid-state nanopore sensing to discriminate two closely related proteins,  $\alpha$ - and  $\gamma$ -thrombin. We show that aptamer functionalization improves protein discrimination thanks to a significant difference in the relative current blockade amplitude. To enhance discrimination, we post-processed the signals using machine learning and training algorithms and we were able to reach an accuracy of 98.8 % of the signals generated by  $\gamma$ -thrombin and  $\alpha$ -thrombin respectively.