

COMPUTATIONAL SUPER-RESOLUTION MICROSCOPY: HISTORIC REVIEW, CURRENT STATUS, AND FUTURE APPLICATIONS

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KEY WORDS: Super-resolution microscopy, computational super-resolution theory

ABSTRACT

Many strategies for overcoming the optical diffraction limit (e.g. PALM, STORM, SOFI, STED, SIM) are now widely accepted tools in biological microscopy. However, a more elusive goal is to demonstrate a computational approach that can *significantly and verifiably increase the resolution (by three or more times) in a conventional microscope*, while imaging biological samples labeled with *standard, non-switching* fluorophores. Although seemingly impossible, such a concept was actually proposed, in principle, in as early as the 1980s. However, it is not until much more recently that we have demonstrated its capability in practice [1-2]. In this talk, we give a brief historic review of this super-resolution technique, present its current state of the art, and discuss some potentially promising uses in the field of biology.

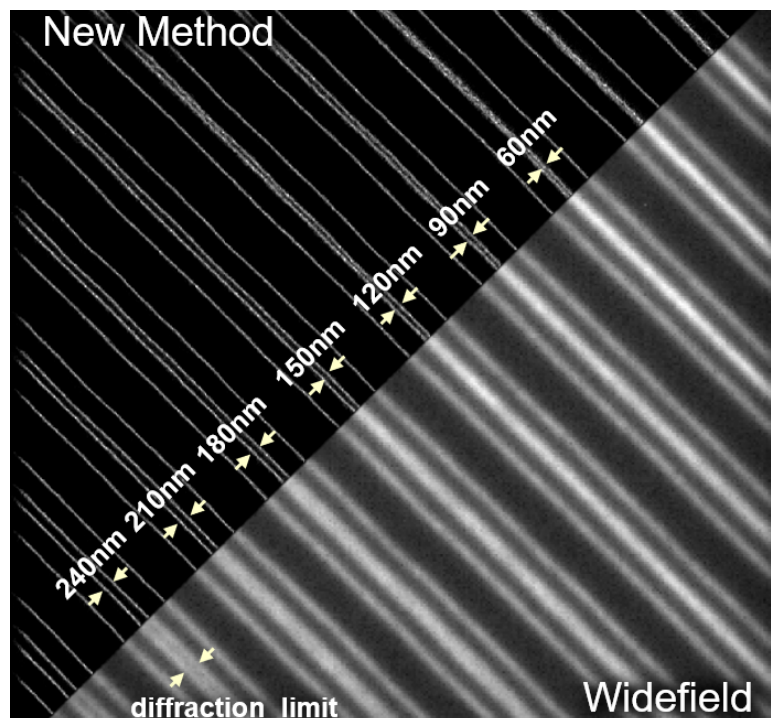


Fig. 1: Argolight fluorescent test slide shows our computational super-resolution approach can achieve 60nm lateral resolution which is nearly *four times improvement* over the 240nm diffraction limit. Objective: NA1.4, 100x.

[1] Yu, J.-Y., Becker, S. R., Folberth, J., Wallin, B. F., Chen, S., and Cogswell, C. J., "Achieving superresolution with illumination-enhanced sparsity," *Opt. Express* **26**, 9850-9865 (2018).

[2] Xing, J., Chen, S., Becker, S. R., Yu, J.-Y., and Cogswell, C. J., " ℓ_1 -regularized maximum likelihood estimation with focused-spot illumination quadruples the diffraction-limited resolution in fluorescence microscopy." *Opt. Express* **28**, 39413-39429 (2020).