

# Computational Super-Resolution Microscopy: Combining Noise Models, Regularization, and Object Sparsity to Achieve Highest Resolution

Jian Xing<sup>1</sup>, Simeng Chen<sup>1</sup>, Stephen R. Becker<sup>2</sup>, Jiun-Yann Yu<sup>1</sup>, and Carol J. Cogswell<sup>1</sup>

<sup>1</sup>Department of Electrical, Computer and Energy Engineering, and

<sup>2</sup>Department of Applied Mathematics

University of Colorado,

Boulder, CO 80309-0425, USA

E-mail: [jian.xing@colorado.edu](mailto:jian.xing@colorado.edu) or [cogswell@colorado.edu](mailto:cogswell@colorado.edu)

**KEY WORDS:** Super-resolution microscopy, computational super-resolution, regularization

## ABSTRACT

Super-resolution fluorescence microscopy has obvious benefits in advancing biological study of small structures beyond the diffraction limit. Currently, super-resolution microscopy techniques that achieve more than two times resolution improvement over the diffraction limit have generally required stimulated emission or switchable fluorophores. Using standard fluorophores and a minimally modified widefield microscope, we demonstrated [1] it is possible to achieve more than two times resolution improvement over the diffraction limit using numerical post-processing.

Following a careful examination of the imaging system, we present an algorithm with improved performance over our previous work, and present verified ground-truth results.

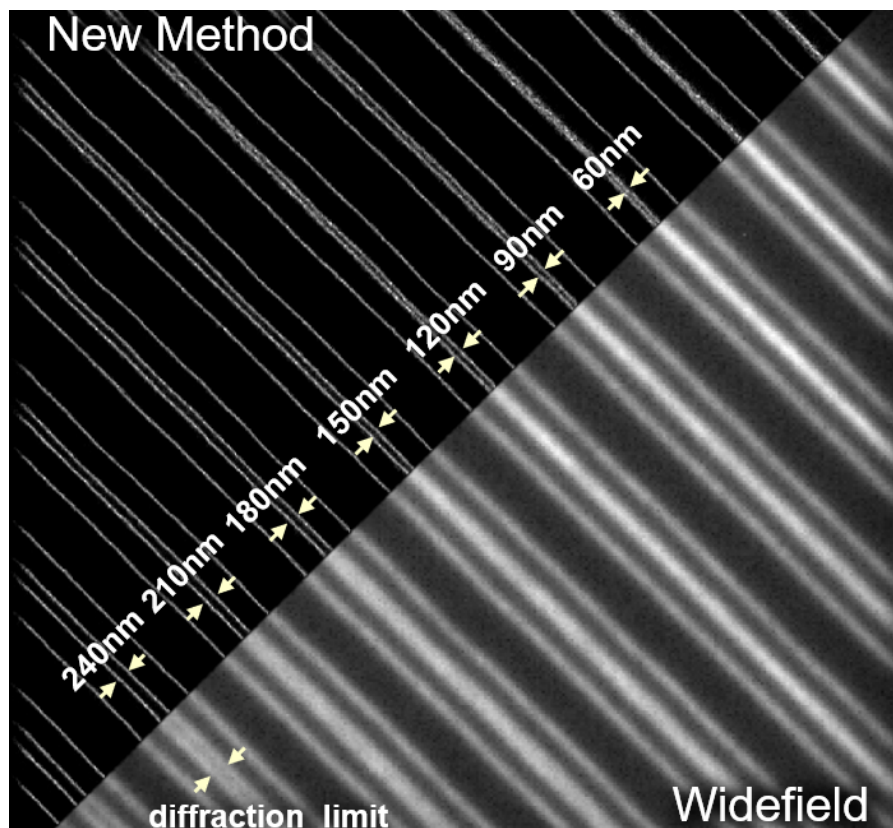


Fig. 1: Argolight fluorescent test slide shows our computational super-resolution approach can achieve 60nm lateral resolution which is nearly four times improvement over the 240nm diffraction limit. Objective: NA1.4, 100x.

[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 (Apr 2018).