

## PSF Engineering

**Overview:** With the recent award of the Nobel Prize to Betzig and Moerner there has been a significant increase in awareness of point spread function (PSF) engineering. In this case the SLM is placed in the emission arm of a microscope. Moerner demonstrated use of PSF engineering with a Meadowlark Optics SLM for super-resolution imaging and 3D localization of fluorescence emitters. PSF engineering has been demonstrated to enable a microscope to image a sample using multiple imaging modalities simultaneously change between modalities non-mechanically. This allows for imaging of structures with a weak refractive index, and for quantitative measurement of phase structures. Imaging modalities that have been demonstrated include: spiral phase imaging, dark field imaging, phase contrast imaging, differential interference contrast imaging, and extended depth of field imaging.

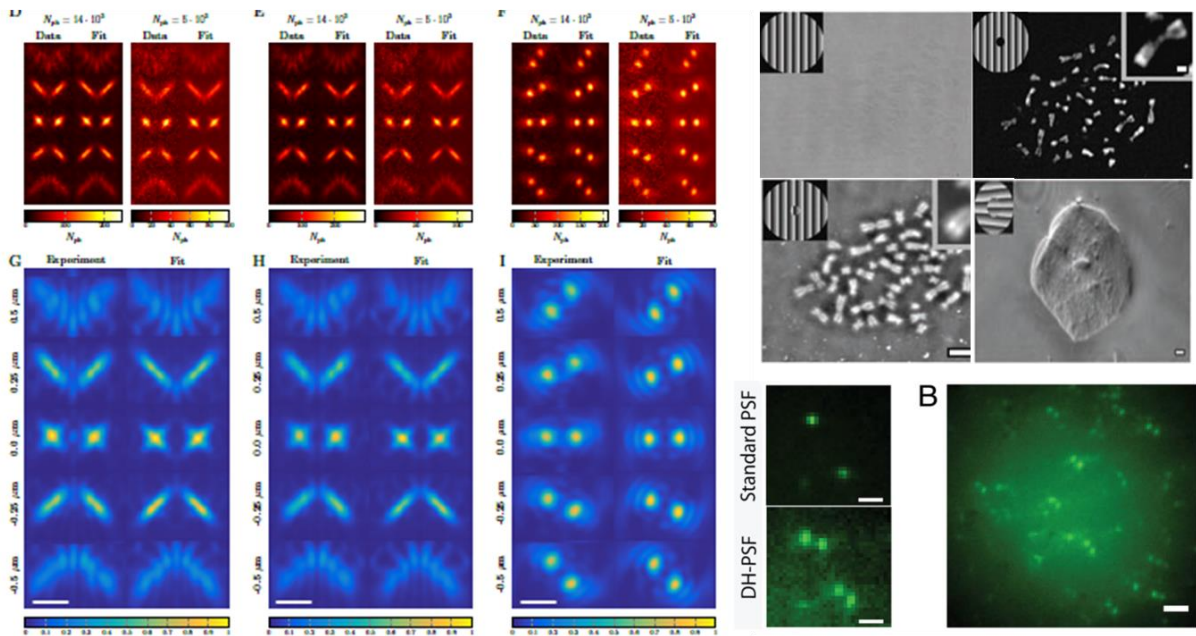


Figure 1 (left) Siemons, M., Hulleman, C. N., Thorsen, R. Ø., Smith, C. S., & Stallinga, S. (2018). High precision wavefront control in point spread function engineering for single emitter localization. *Optics express*, 26(7), 8397-8416. (right top) Maurer, Christian, et al. "What spatial light modulators can do for optical microscopy (right bottom) Pavani, Sri Rama Prasanna, et al. "Three-dimensional, single-molecule fluorescence imaging beyond the diffraction limit by using a double-helix point spread function."

**Critical requirements:** For this market it is important that the SLM minimize optical losses. PSF engineering uses a SLM to manipulate the wavefront in the emission path of the microscope. There is a lack of signal in fluorescence imaging without adding losses. Use of a SLM with a high fill factor minimizes losses to diffraction. High resolution SLMs are ideal for creating the complex phase functions required for 3D localization, and high speed allows for real time deep tissue super-resolution imaging.

### Recommended References:

1. Majeed, H., Kandel, M. E., Han, K., Luo, Z., Macias, V., Tangella, K. V., ... & Popescu, G. (2015). Breast cancer diagnosis using spatial light interference microscopy. *Journal of biomedical optics*, 20(11), 111210.

2. TANG, Z., O'DWYER, K., & HENNELLY, B. (2019, May). Calibration methods for Spatial Light Interference Microscopy. In *Digital Holography and Three-Dimensional Imaging* (pp. W1B-7). Optical Society of America.
3. Nguyen, T. H., Kandel, M., Shakir, H. M., Best-Popescu, C., Arikath, J., Do, M. N., & Popescu, G. (2017). Halo-free phase contrast microscopy. *Scientific reports*, 7, 44034.
4. Siemons, M., Hulleman, C. N., Thorsen, R. Ø., Smith, C. S., & Stallinga, S. (2018). High precision wavefront control in point spread function engineering for single emitter localization. *Optics express*, 26(7), 8397-8416.
5. Kandel, M., Teng, K. W., Selvin, P. R., & Popescu, G. (2016, March). Highly sensitive kinesin-microtubule motility assays using SLIM. In *Quantitative Phase Imaging II* (Vol. 9718, p. 97180T). International Society for Optics and Photonics.
6. Zhang, O., Lu, J., Ding, T., & Lew, M. D. (2018). Imaging the three-dimensional orientation and rotational mobility of fluorescent emitters using the Tri-spot point spread function. *Applied physics letters*, 113(3), 031103.
7. Kandel, M. E., Teng, K. W., Selvin, P. R., & Popescu, G. (2016). Label-free imaging of single microtubule dynamics using spatial light interference microscopy. *ACS nano*, 11(1), 647-655.
8. King, S. V., Yuan, S., & Preza, C. (2018). Performance evaluation of extended depth of field microscopy in the presence of spherical aberration and noise. *Journal of biomedical optics*, 23(3), 036016
9. Zhang, O., Ding, T., Lu, J., Mazidi, H., & Lew, M. D. (2018, February). Measuring 3D molecular orientation and rotational mobility using a Tri-spot point spread function. In *Single Molecule Spectroscopy and Superresolution Imaging XI* (Vol. 10500, p. 105000B). International Society for Optics and Photonics.
10. Lee, Y. J., Cintora, P., Arikath, J., Akinsola, O., Kandel, M., Popescu, G., & Best-Popescu, C. (2017). Quantitative assessment of neural outgrowth using spatial light interference microscopy. *Journal of biomedical optics*, 22(6), 066015.
11. Smith, C., Huisman, M., Siemons, M., Grünwald, D., & Stallinga, S. (2016). Simultaneous measurement of emission color and 3D position of single molecules. *Optics Express*, 24(5), 4996-5013.
12. King, S. V., Doblaz, A., Patwary, N., Saavedra, G., Martínez-Corral, M., & Preza, C. (2015). Spatial light modulator phase mask implementation of wavefront encoded 3D computational-optical microscopy. *Applied optics*, 54(29), 8587-8595.
13. Liu, L., Kandel, M. E., Rubessa, M., Schreiber, S., Wheeler, M. B., & Popescu, G. (2018). Topography and refractometry of sperm cells using spatial light interference microscopy. *Journal of biomedical optics*, 23(2), 025003.