



# Molecular programming with DNA: DNA-based nanoassembly and synthetic biochemical circuits

Friedrich C. Simmel

Physics Department - TU München  
[simmel@tum.de](mailto:simmel@tum.de)

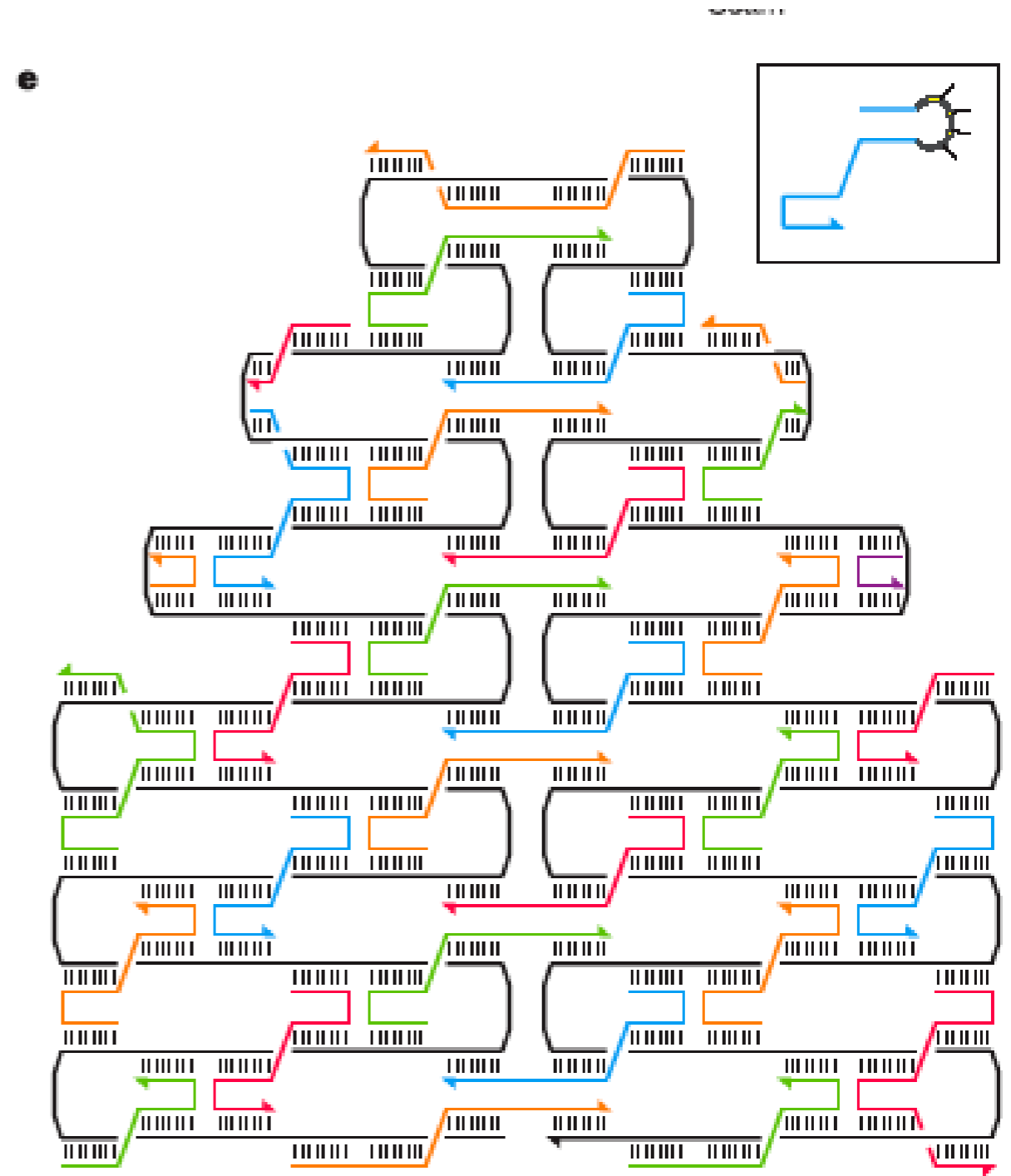
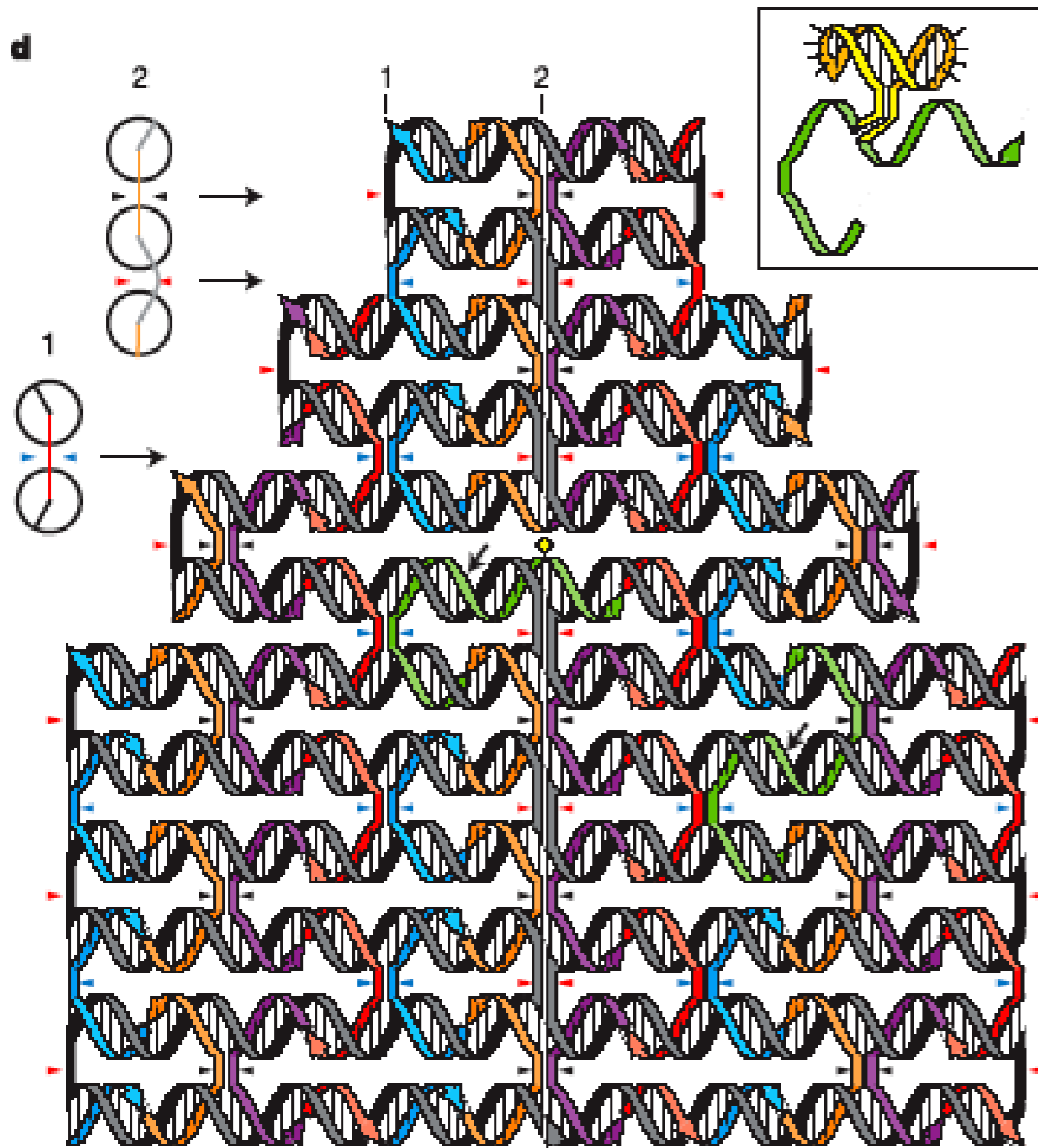
---

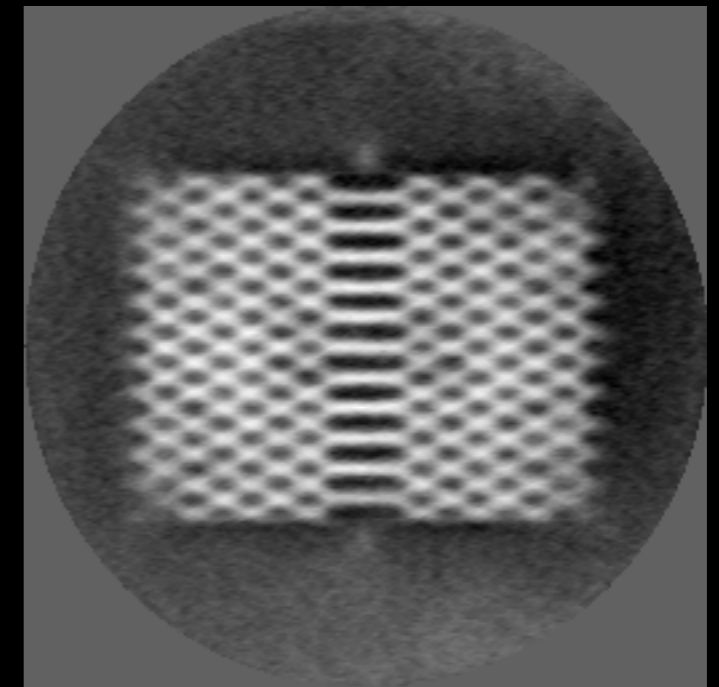
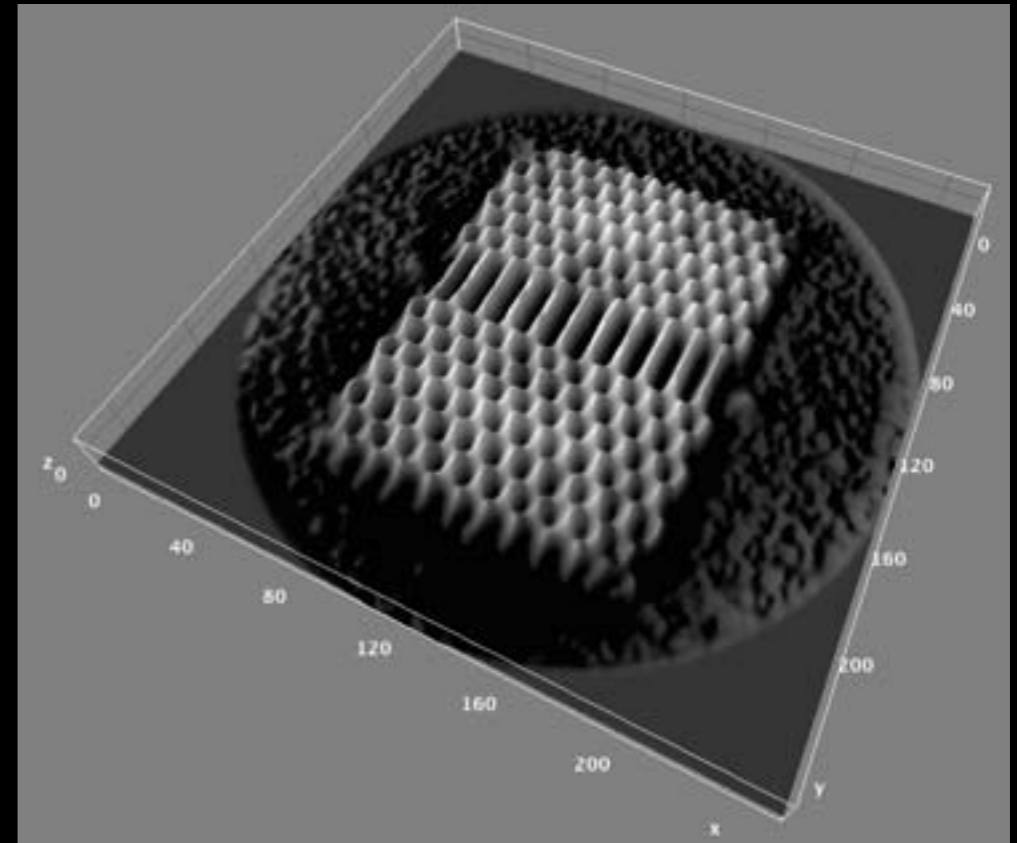
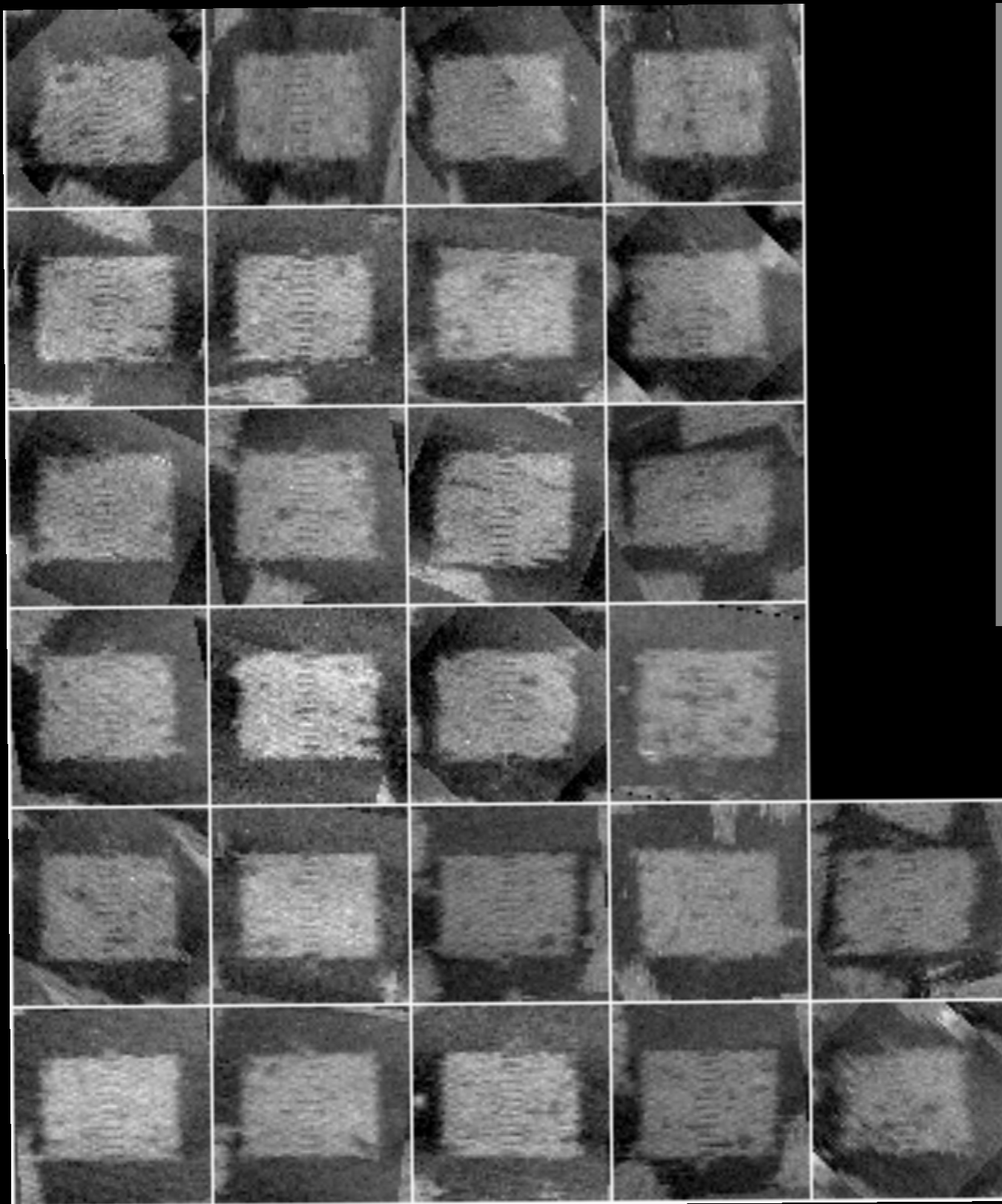


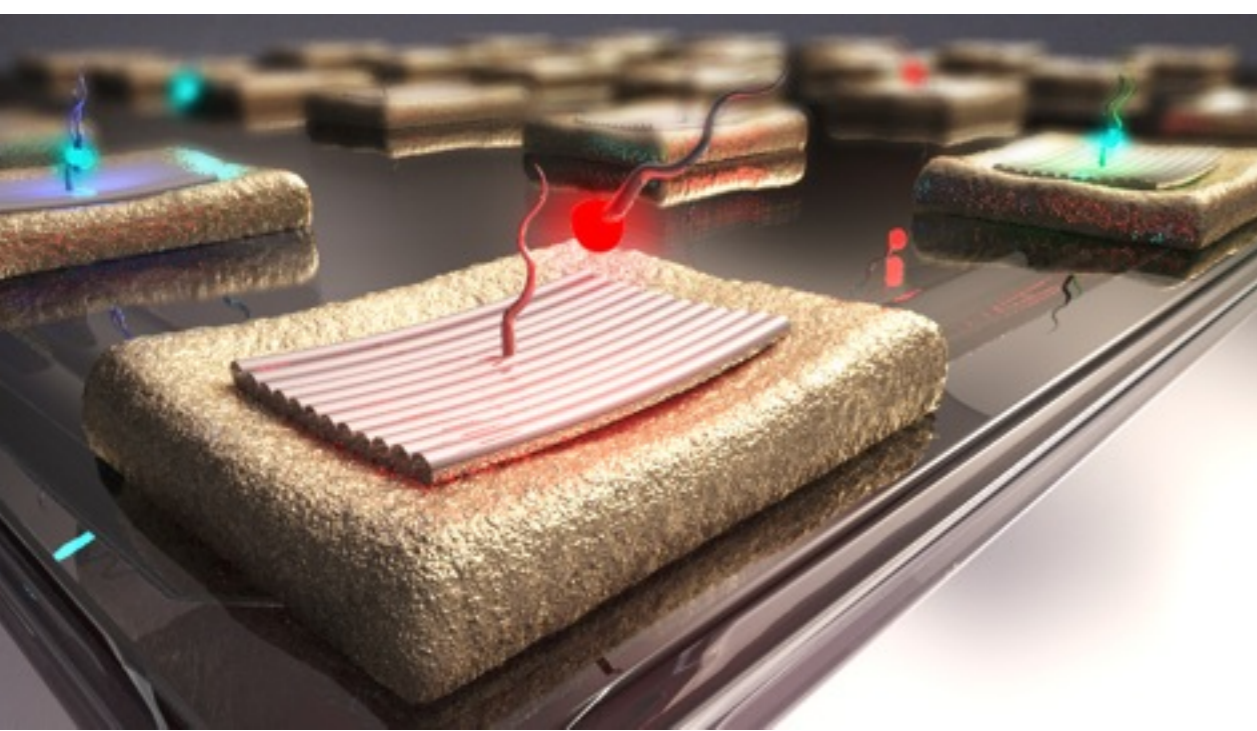
# 1. Nanostructures & nanodevices

---

# DNA origami







# **DNA strand displacement on lithographic DNA origami arrays**

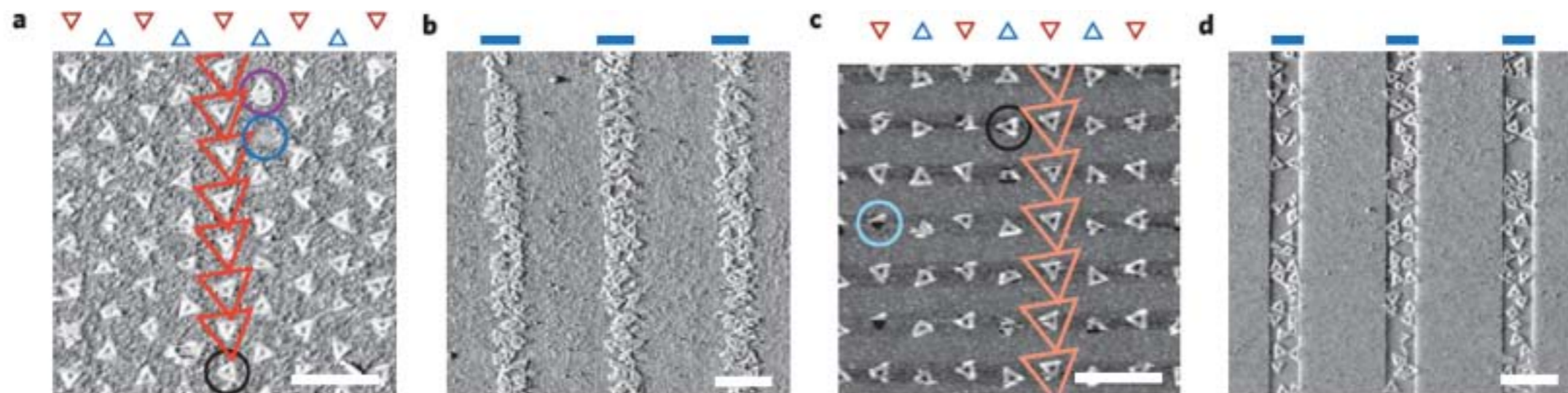
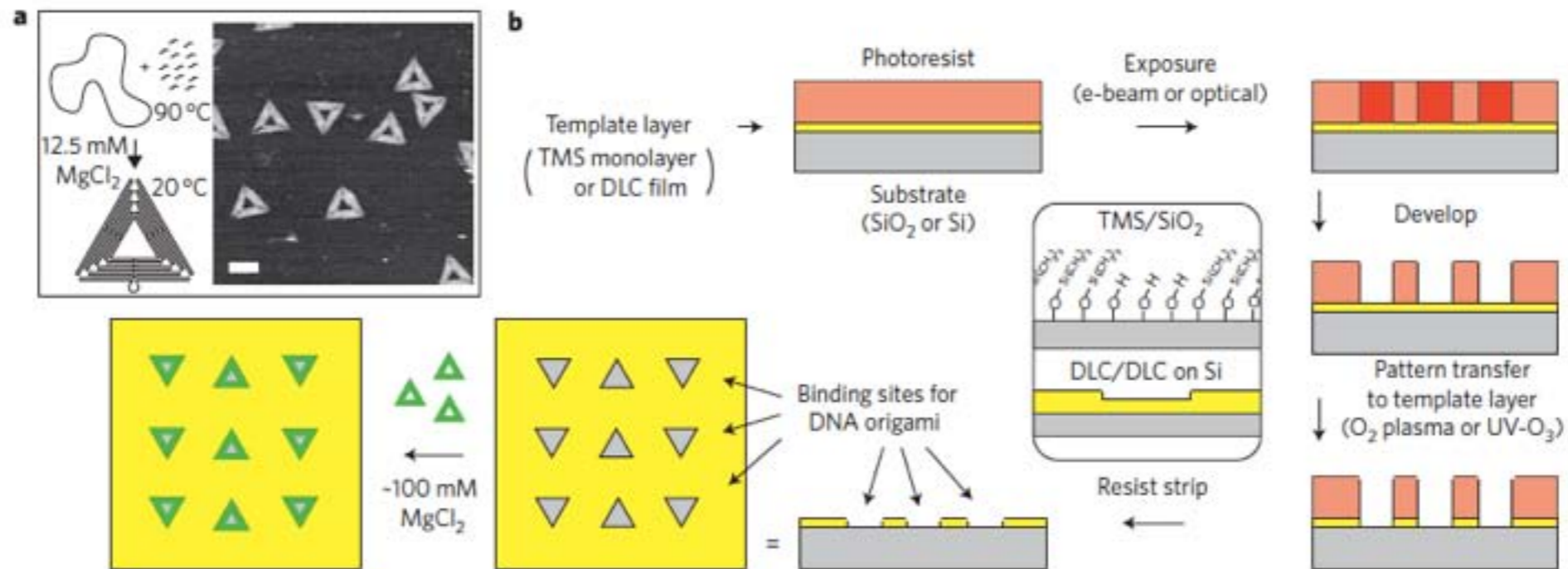
Max Scheible, Günther Pardatscher, Anton Kuzyk

M. B. Scheible, G. Pardatscher, A. Kuzyk, F. C. Simmel, *Nano Lett.* **2014**, *14*, 1627-1633.

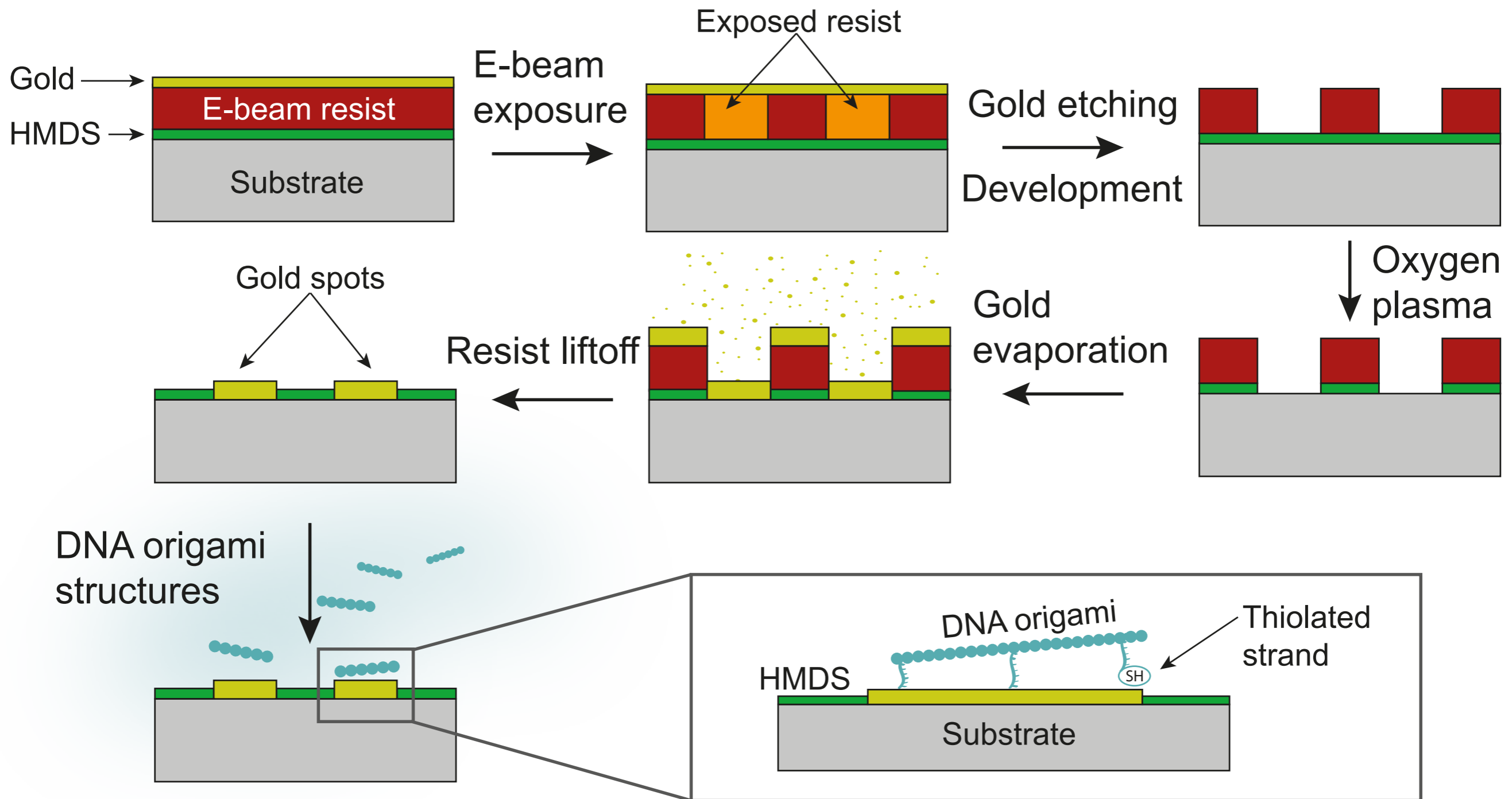
# Bridging the scales with lithography

## Placement and orientation of individual DNA shapes on lithographically patterned surfaces

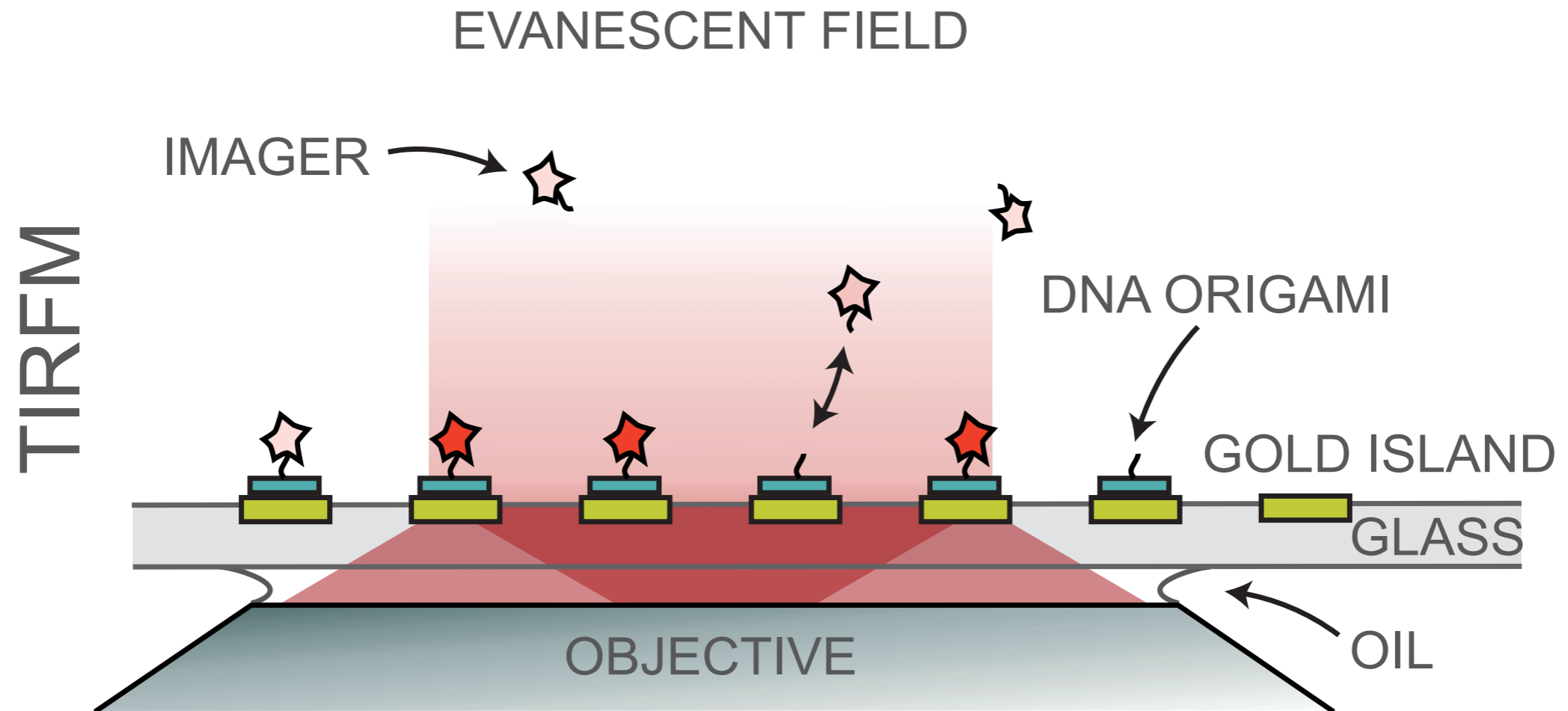
Ryan J. Kershner<sup>1</sup>, Luisa D. Bozano<sup>1</sup>, Christine M. Micheel<sup>1</sup>, Albert M. Hung<sup>1</sup>, Ann R. Fornof<sup>1</sup>, Jennifer N. Cha<sup>2\*</sup>, Charles T. Rettner<sup>1</sup>, Marco Bersani<sup>1</sup>, Jane Frommer<sup>1</sup>, Paul W. K. Rothemund<sup>2\*</sup> and Gregory M. Wallraff<sup>1\*</sup> Nature Nanotech 2009



# E-beam lithography on glass substrates

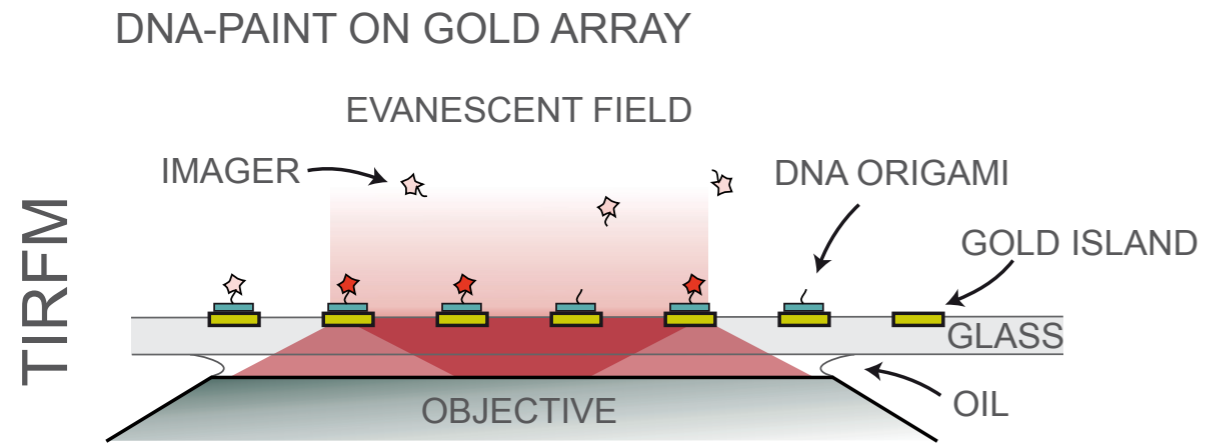


# TIRF and DNA PAINT on lithographic origami arrays

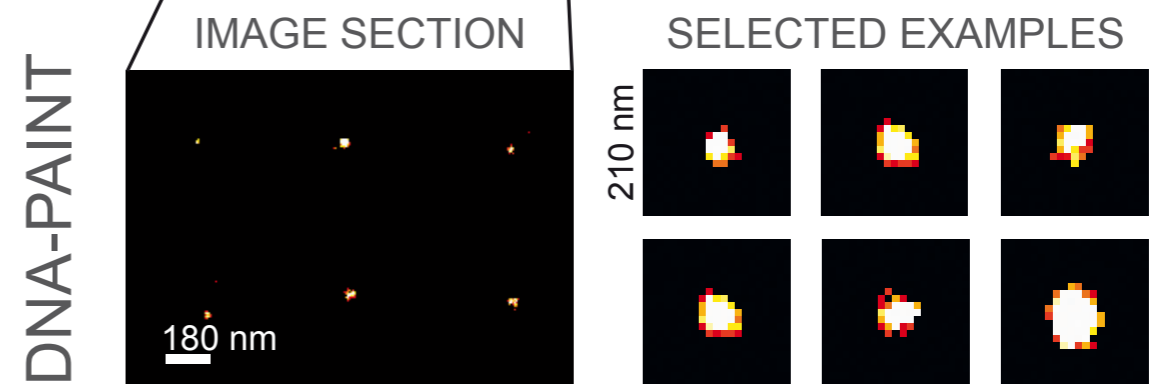
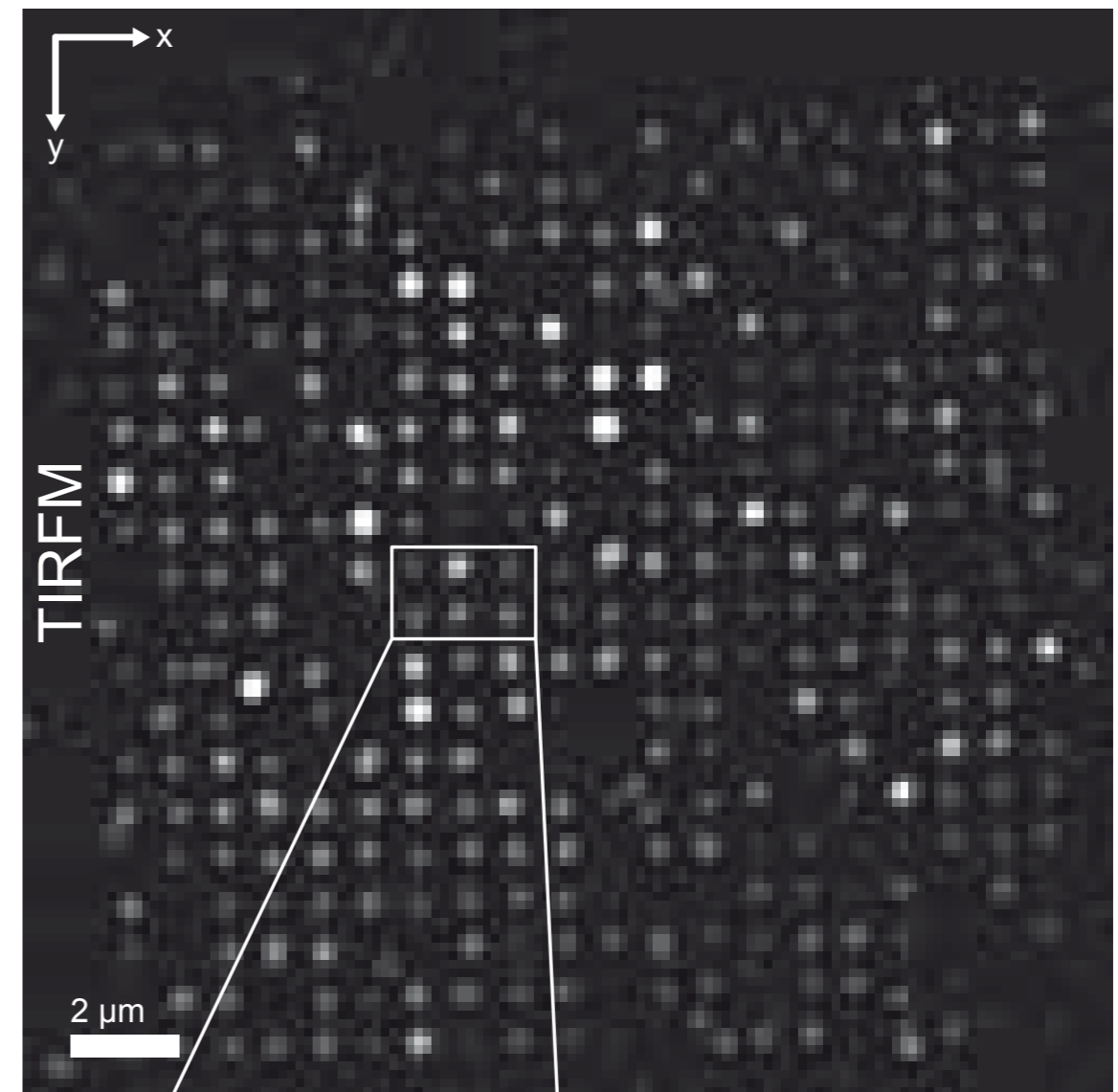
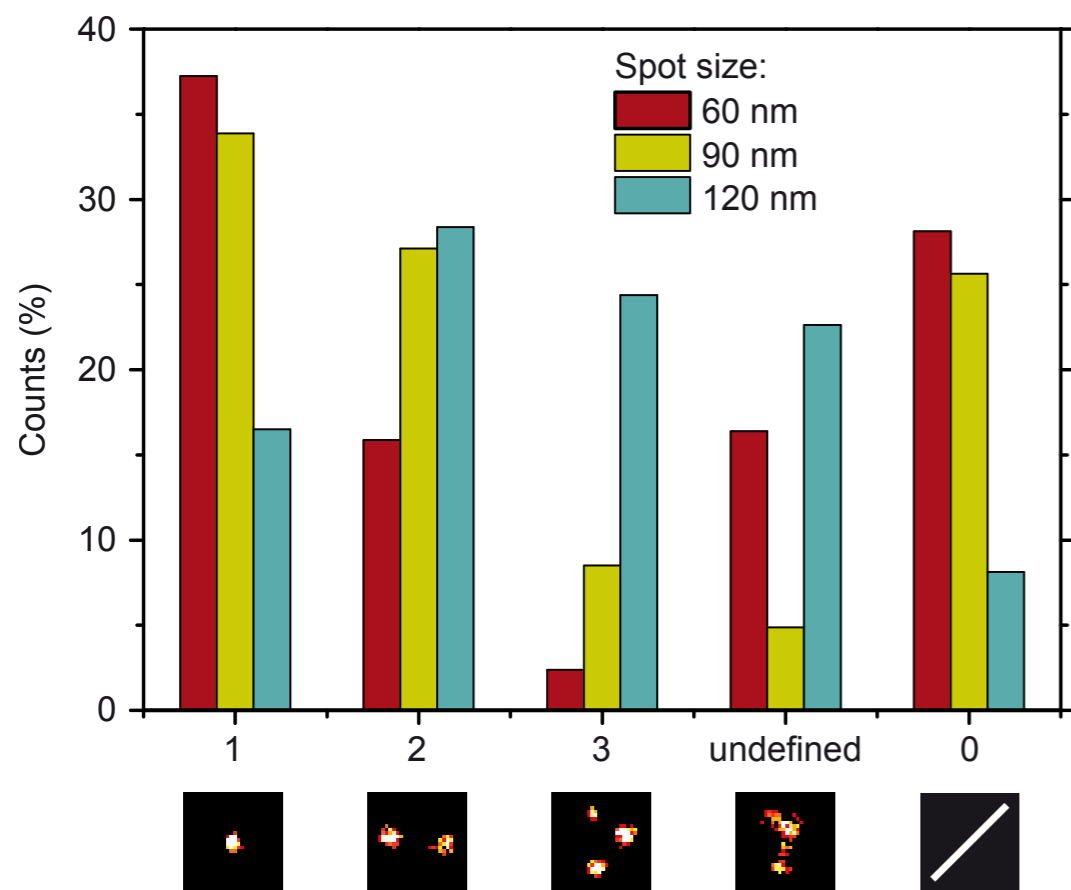
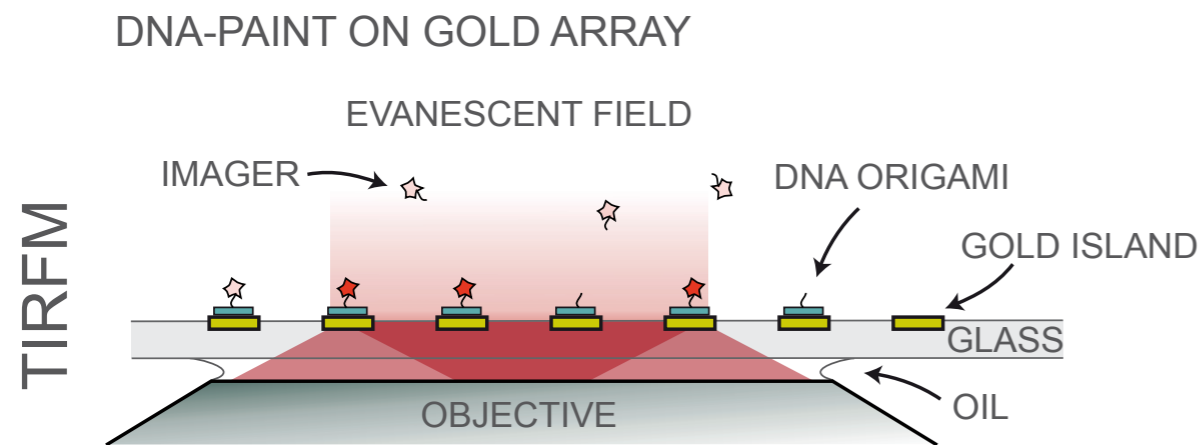




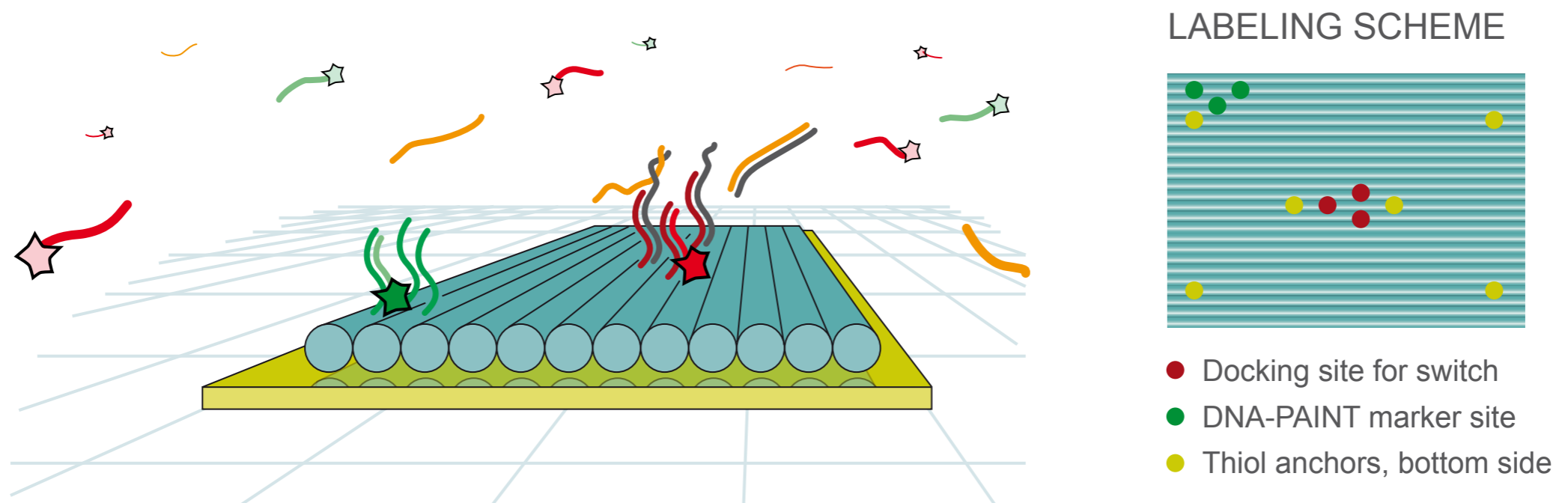
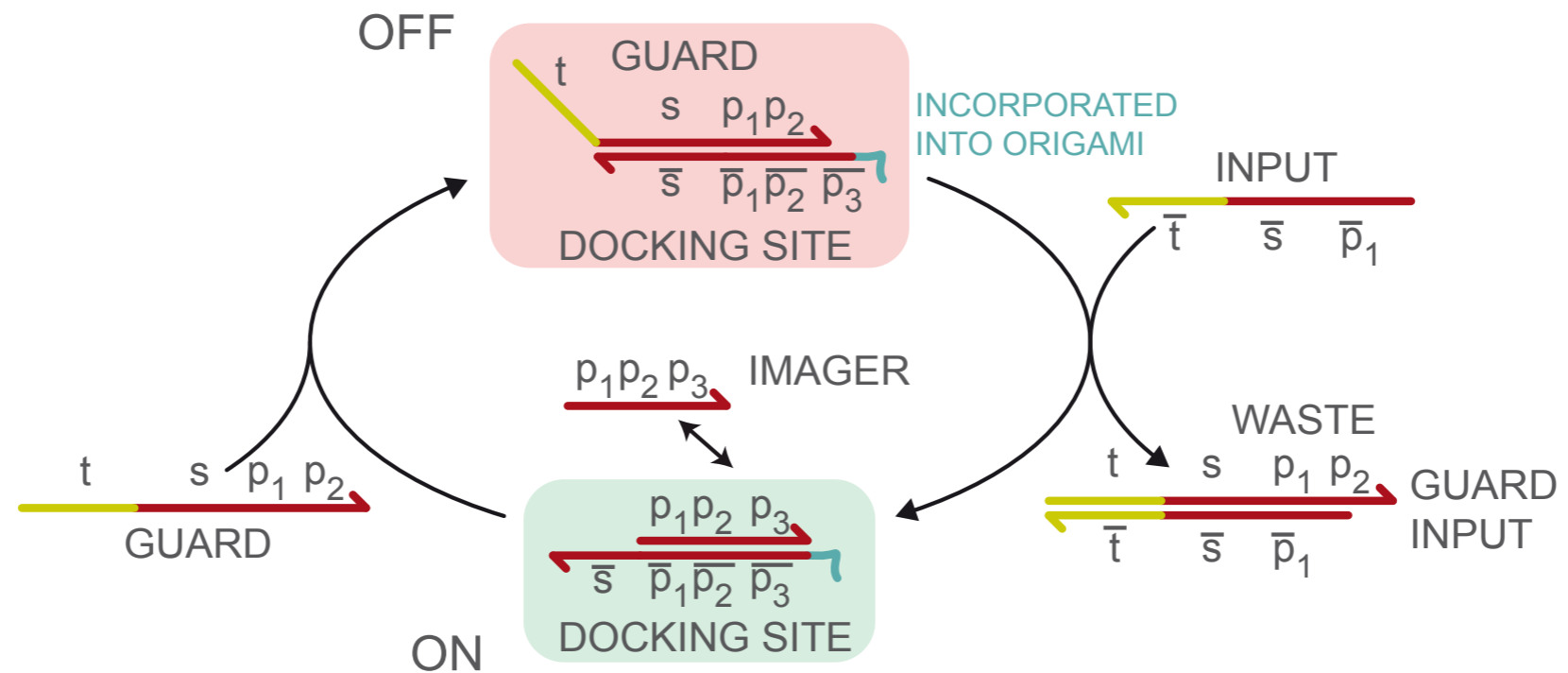
# TIRF and DNA PAINT on lithographic origami arrays



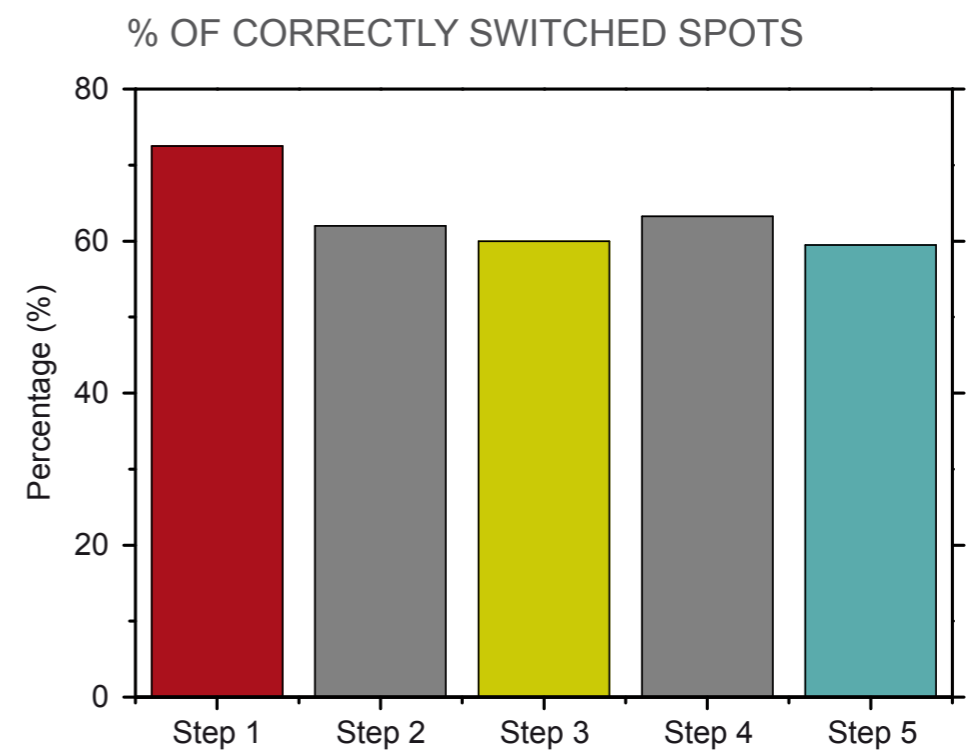
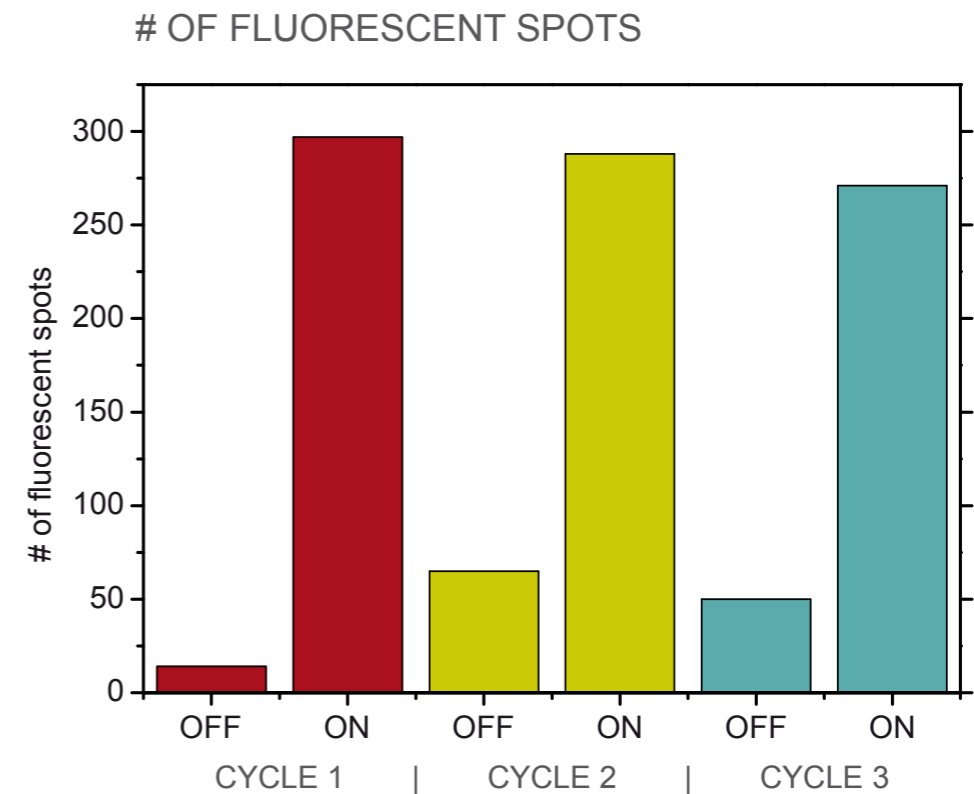
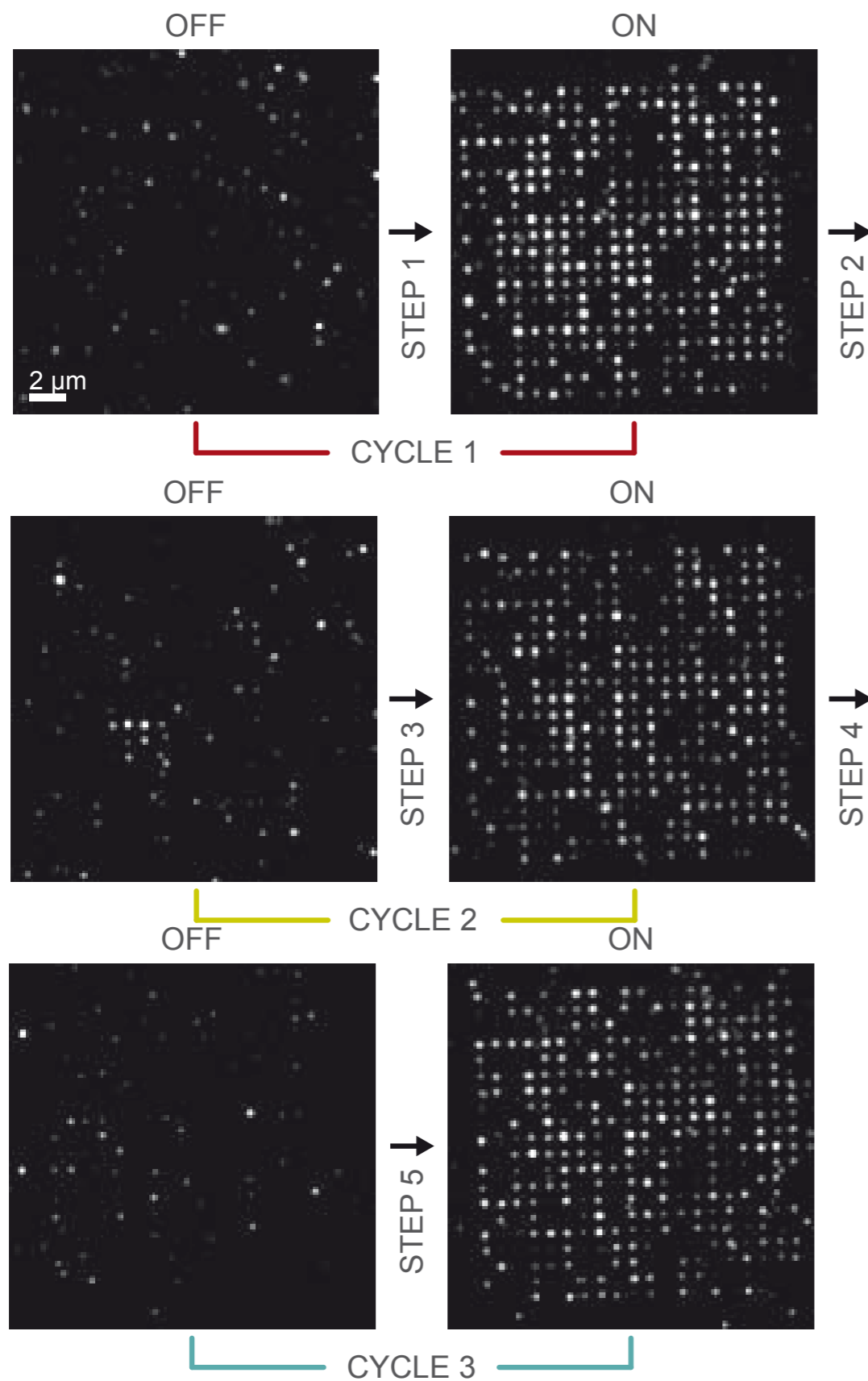
# TIRF and DNA PAINT on lithographic origami arrays



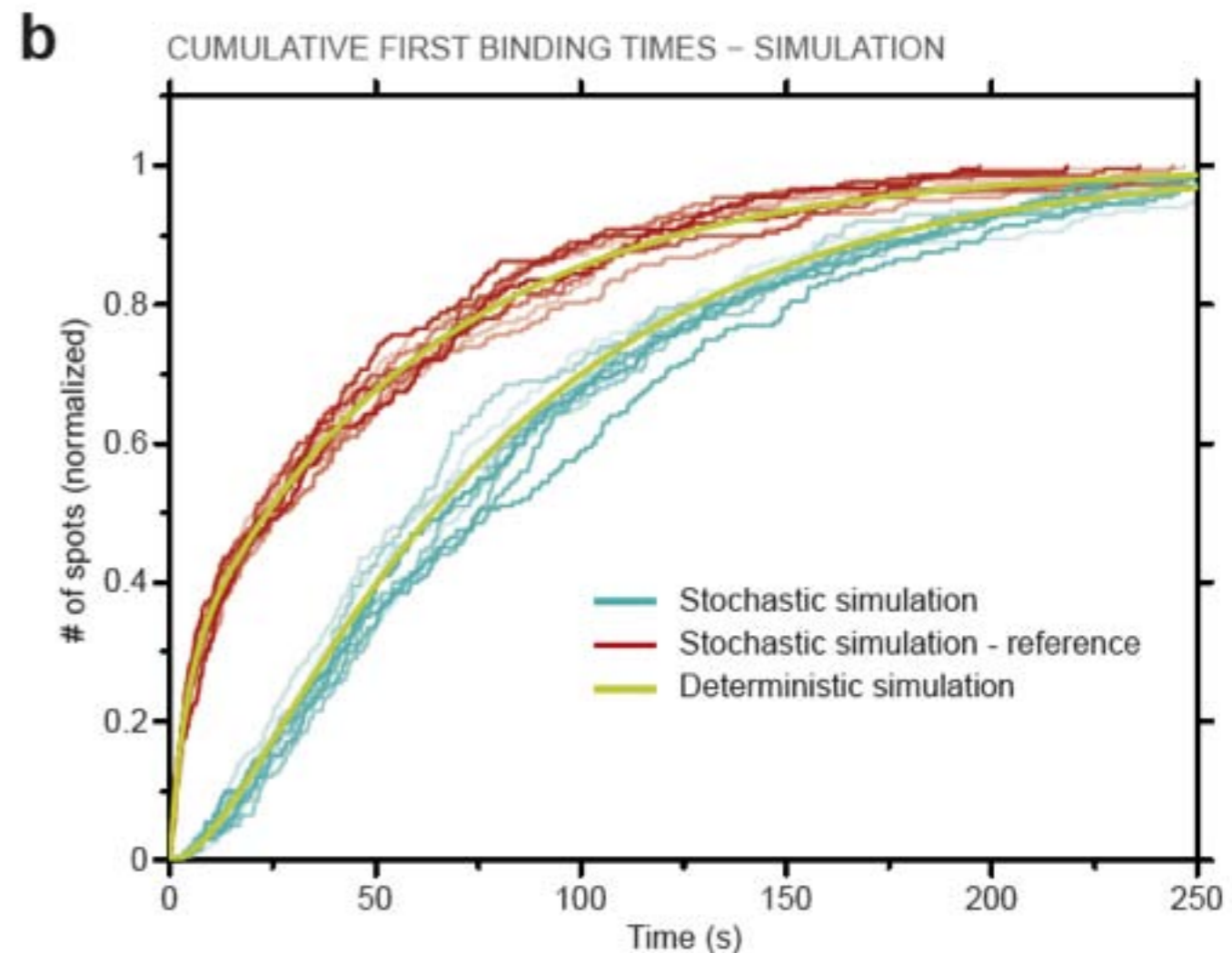
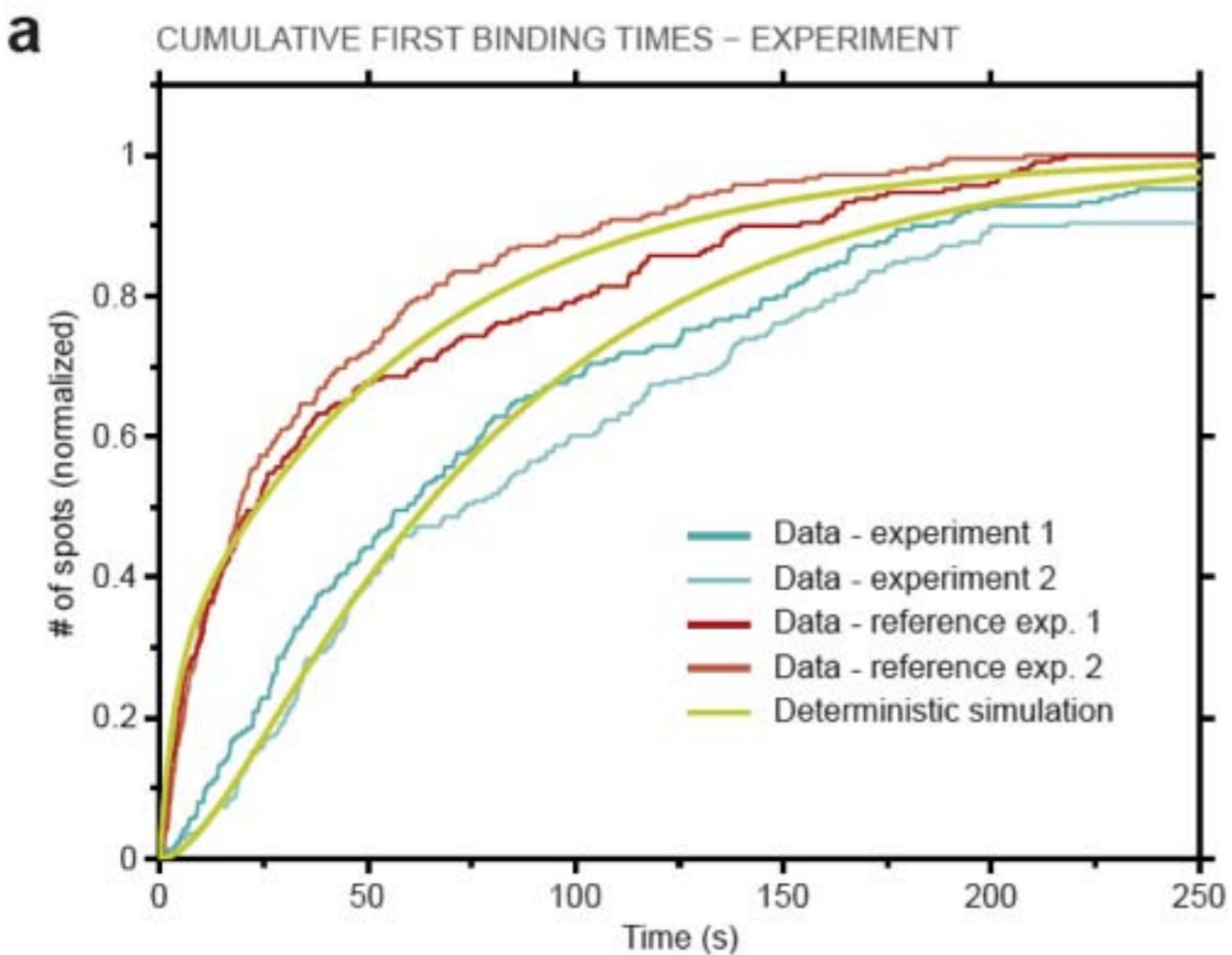
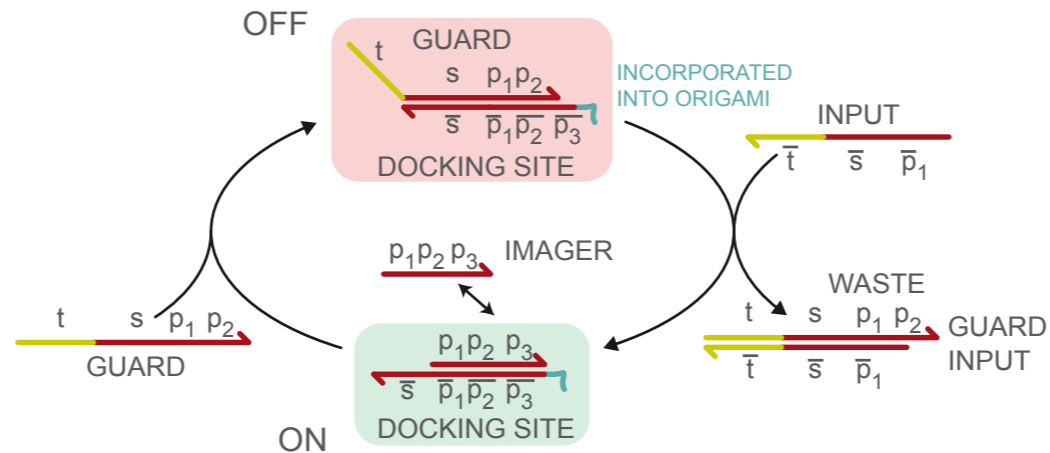
# Strand displacement on origami arrays



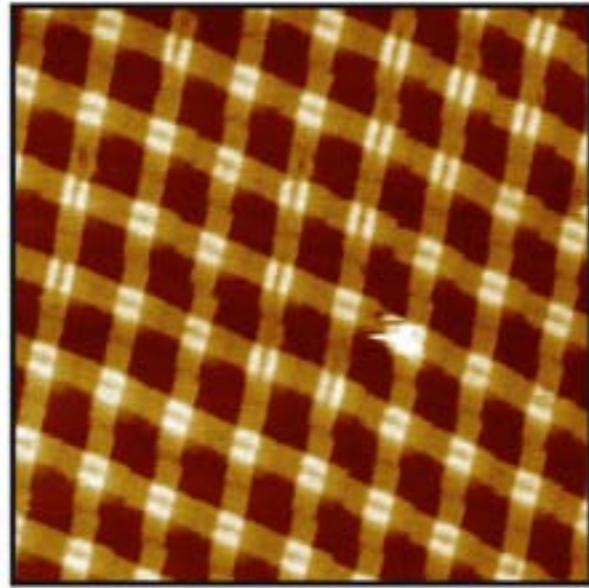
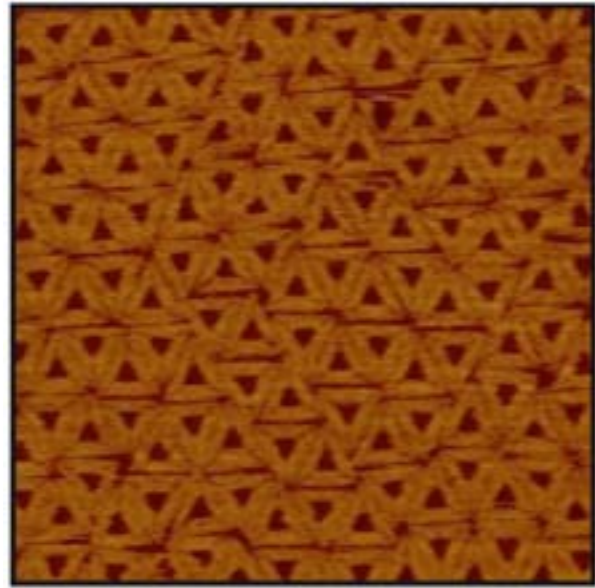
# Strand displacement on origami arrays



# Kinetics of strand displacement on the array



(simulations with Visual DSD)

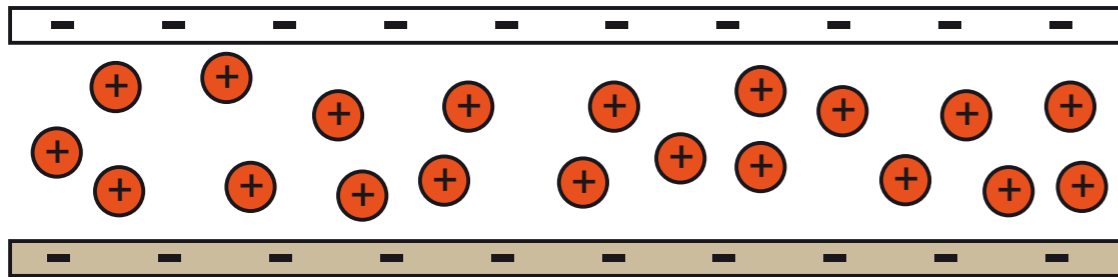
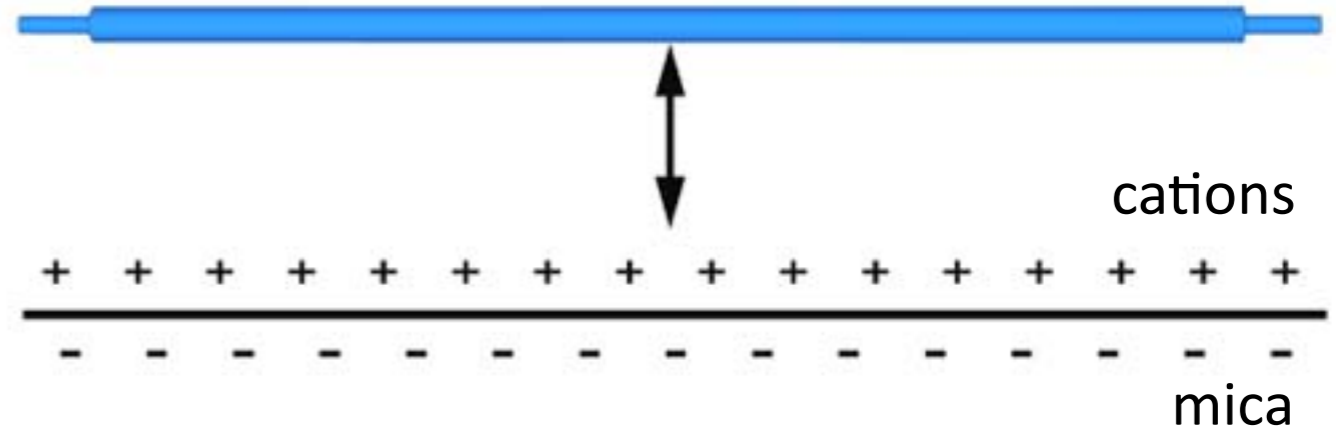


# Large scale assembly of DNA origami tiles on surfaces

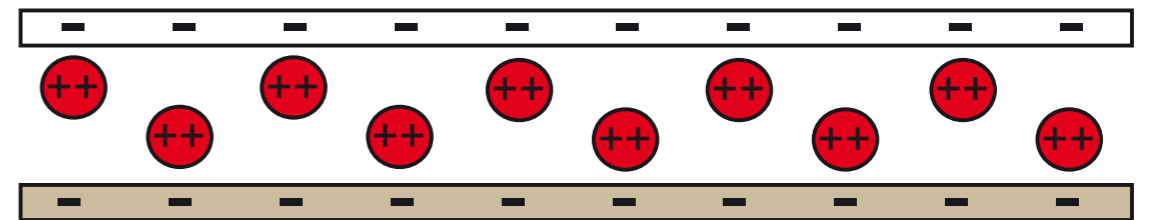
Ali Aghebat Rafat, Tobias Pirzer, Max Scheible, Anna Kostina

A. Aghebat Rafat, T. Pirzer, M.B. Scheible, A. Kostina, F.C. Simmel, *Angewandte Chemie Int. Ed.*, **53** (29), 7665-7668 (2014)

# DNA on mica surfaces

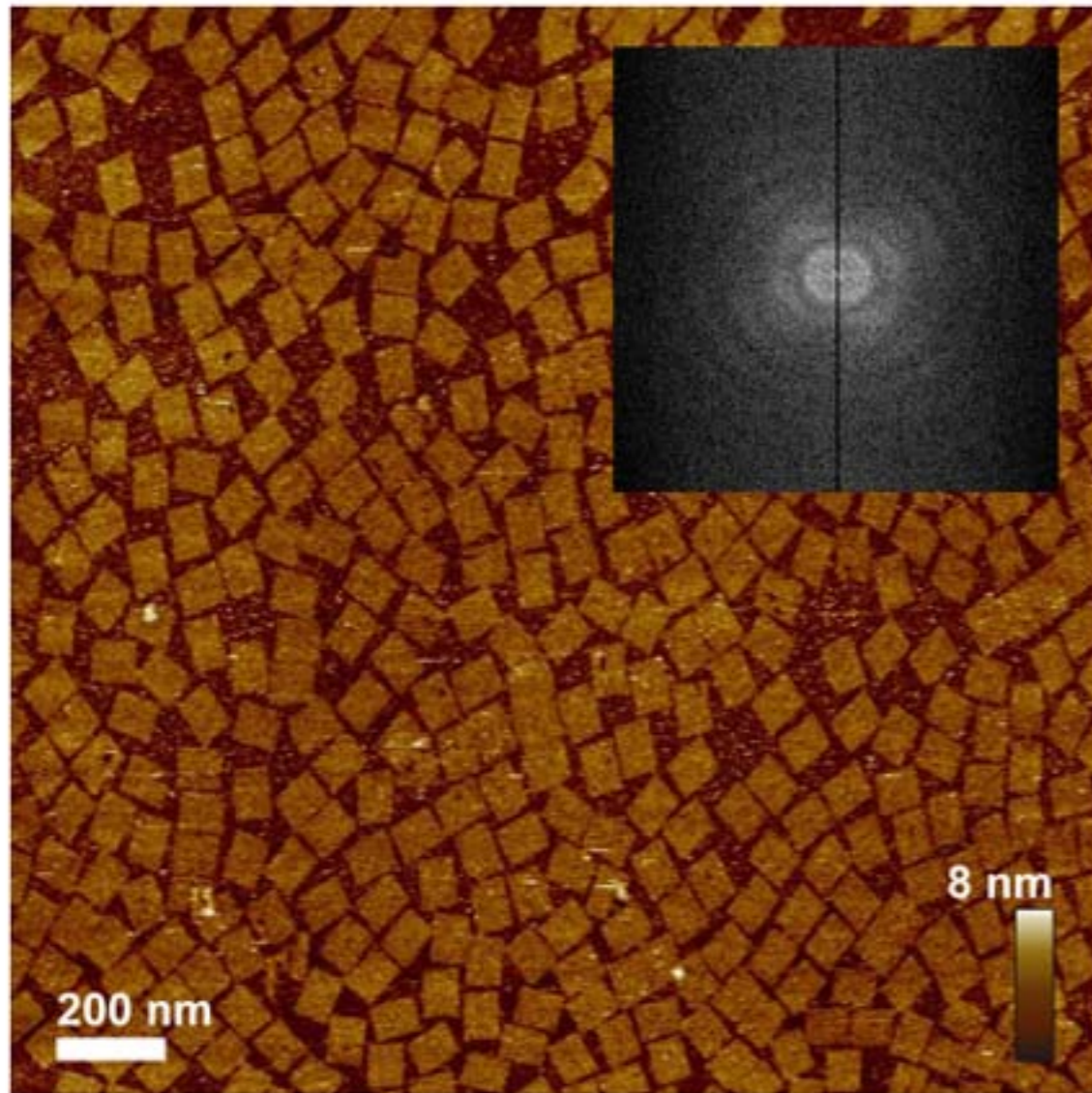


monovalent cations

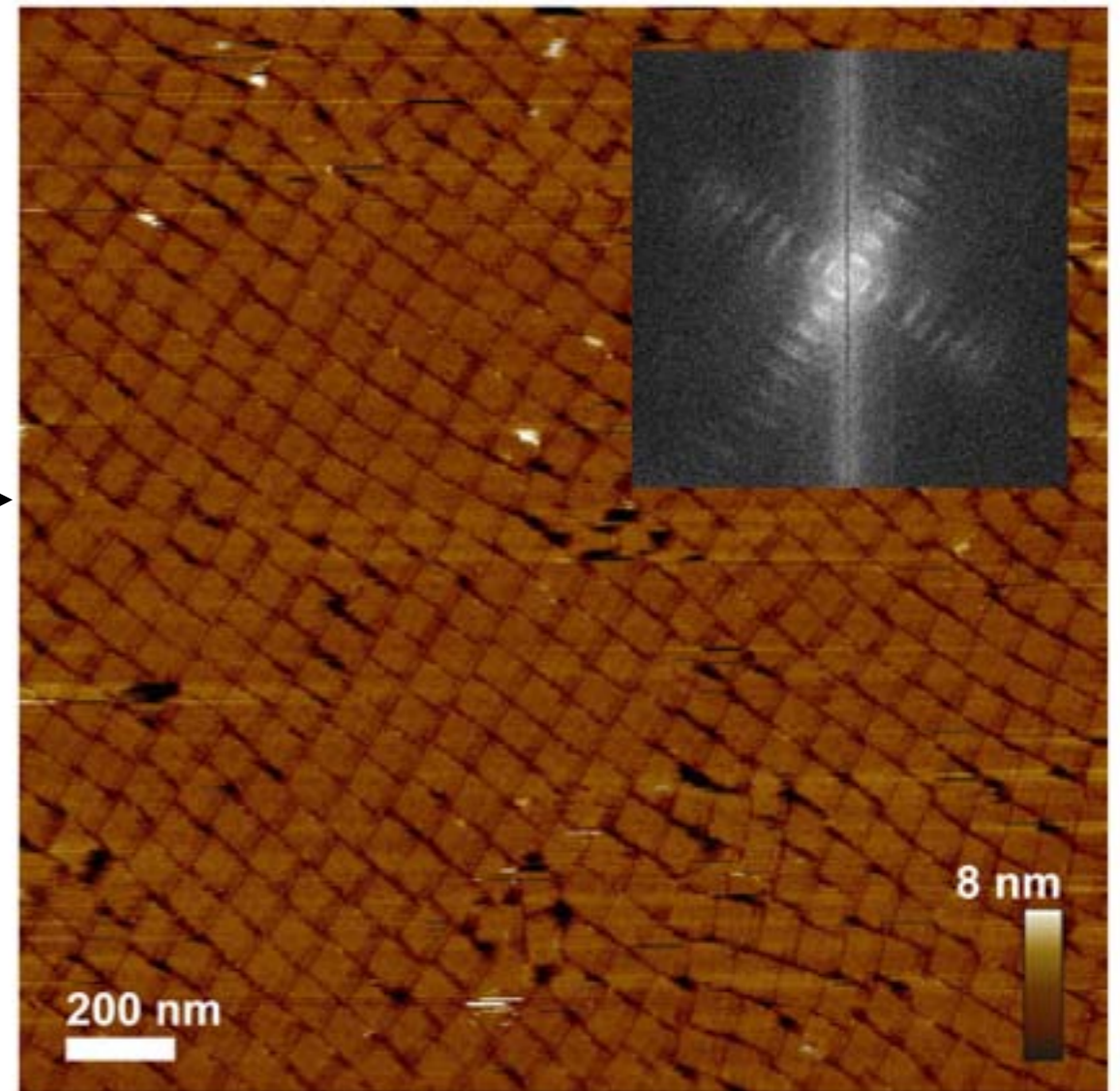


divalent cations

# Surface-assisted arrangement of DNA origami tiles



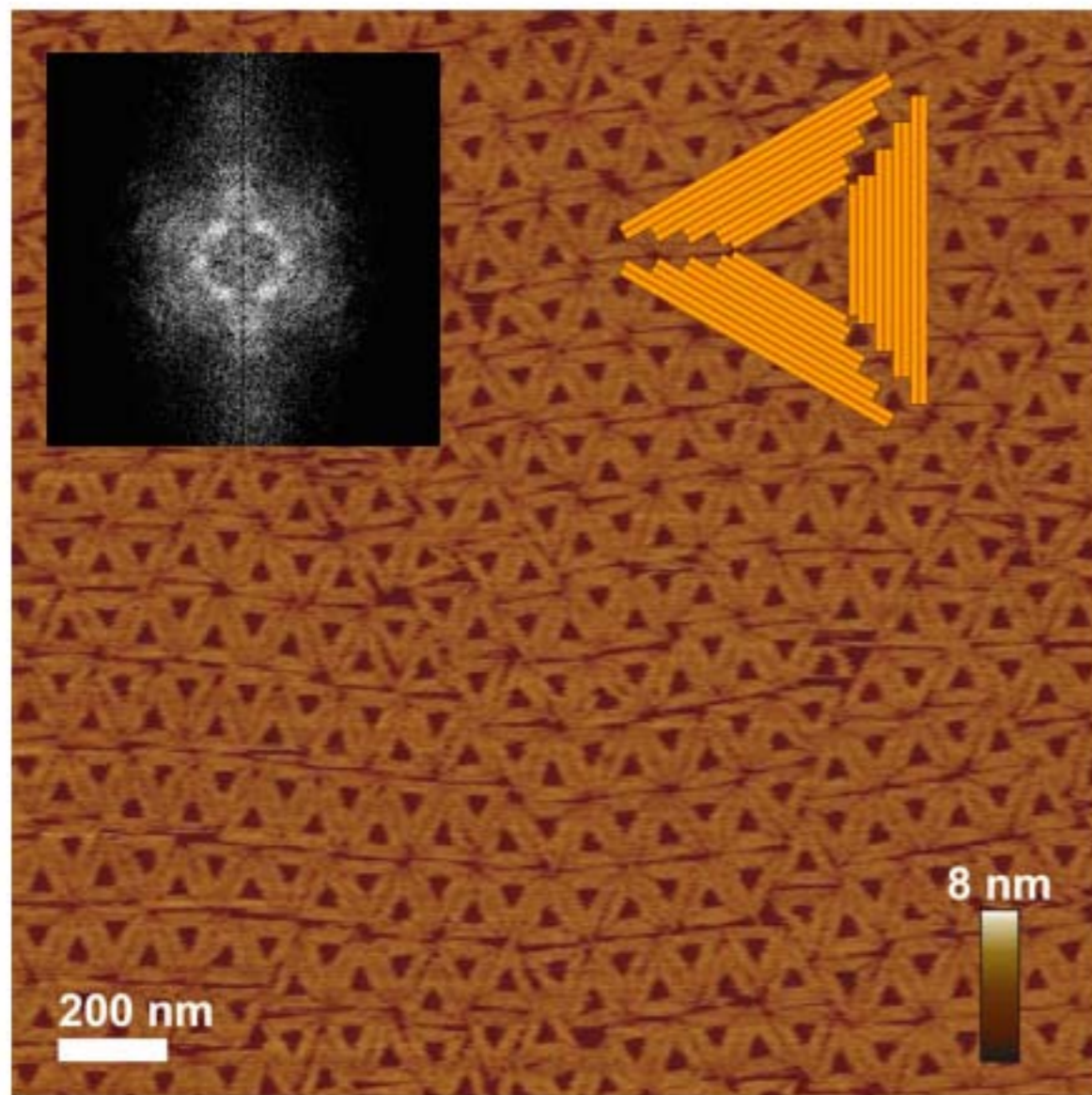
Na<sup>+</sup>



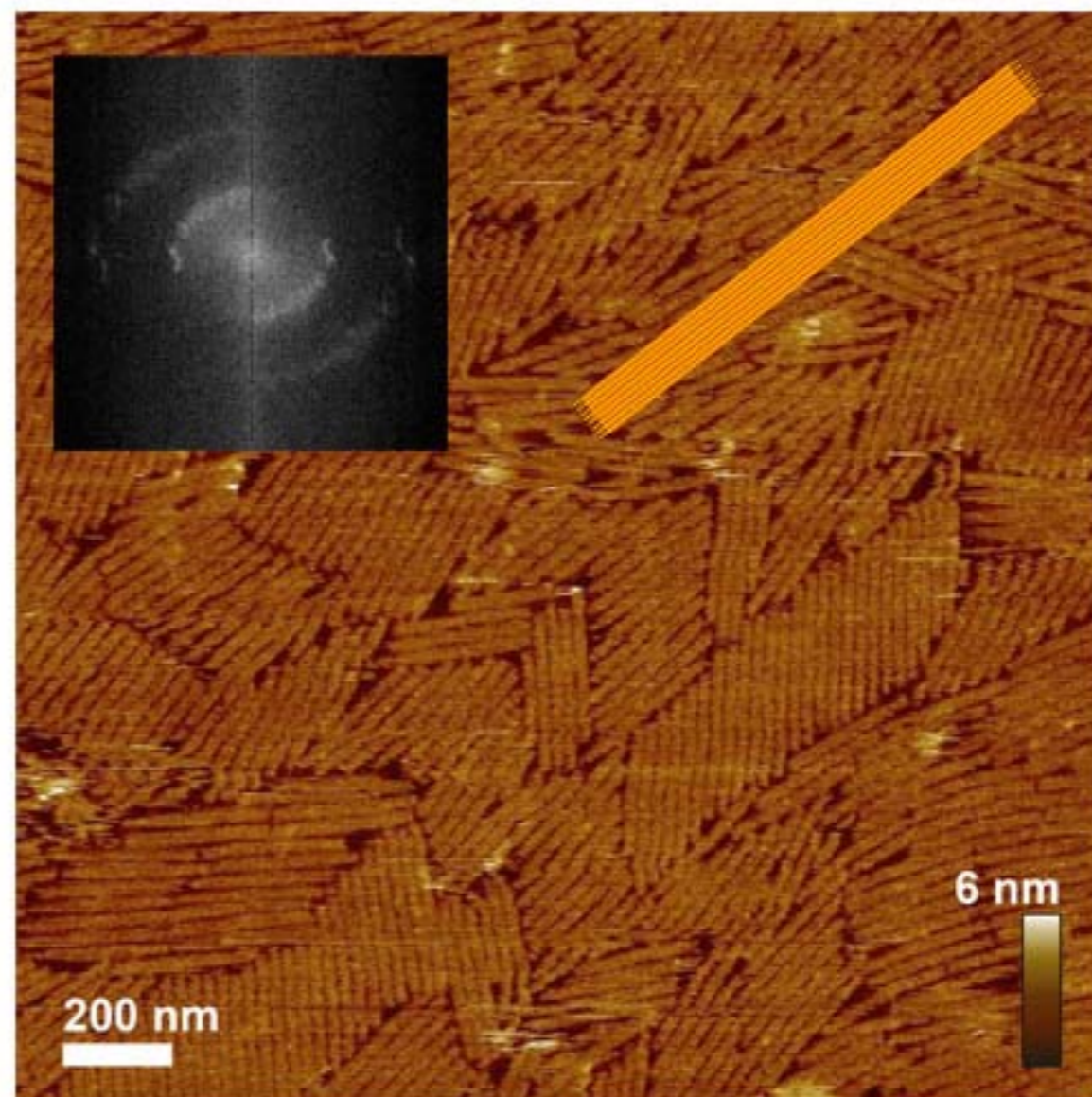


# Surface-assisted arrangement of DNA origami tiles

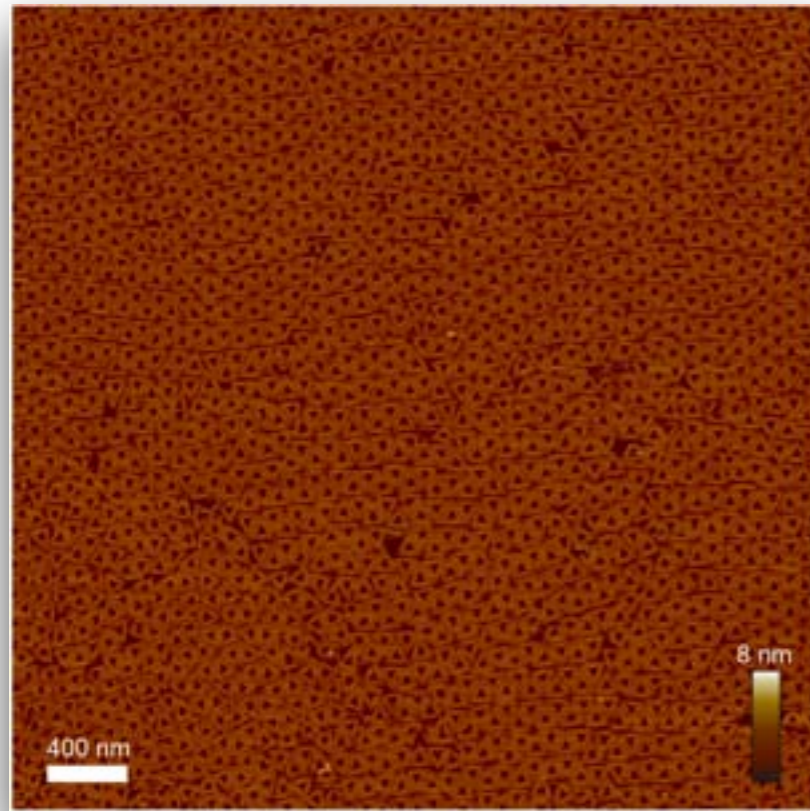
A



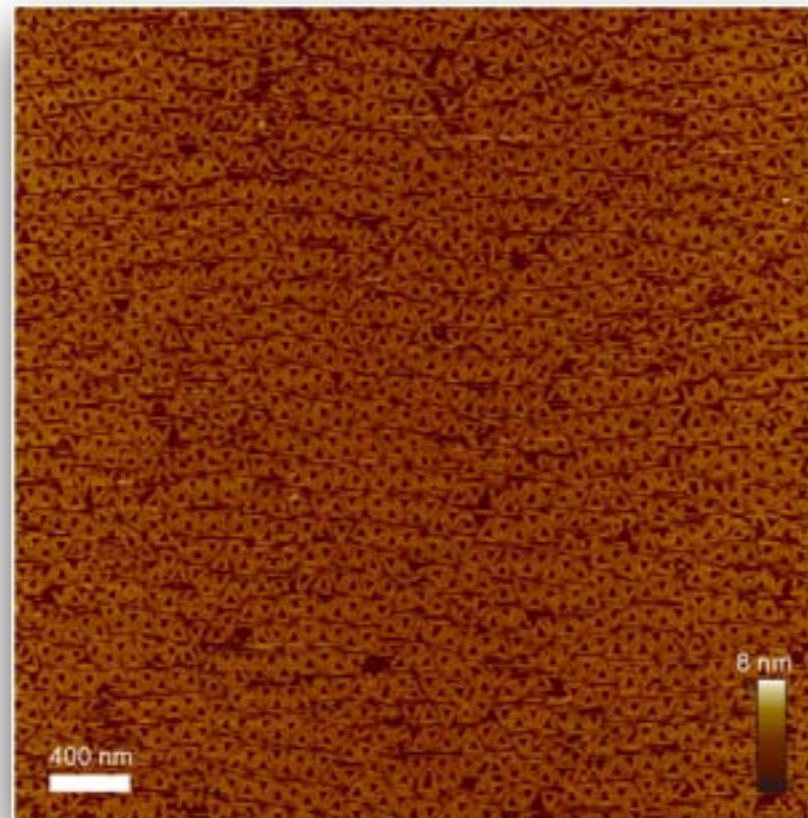
B



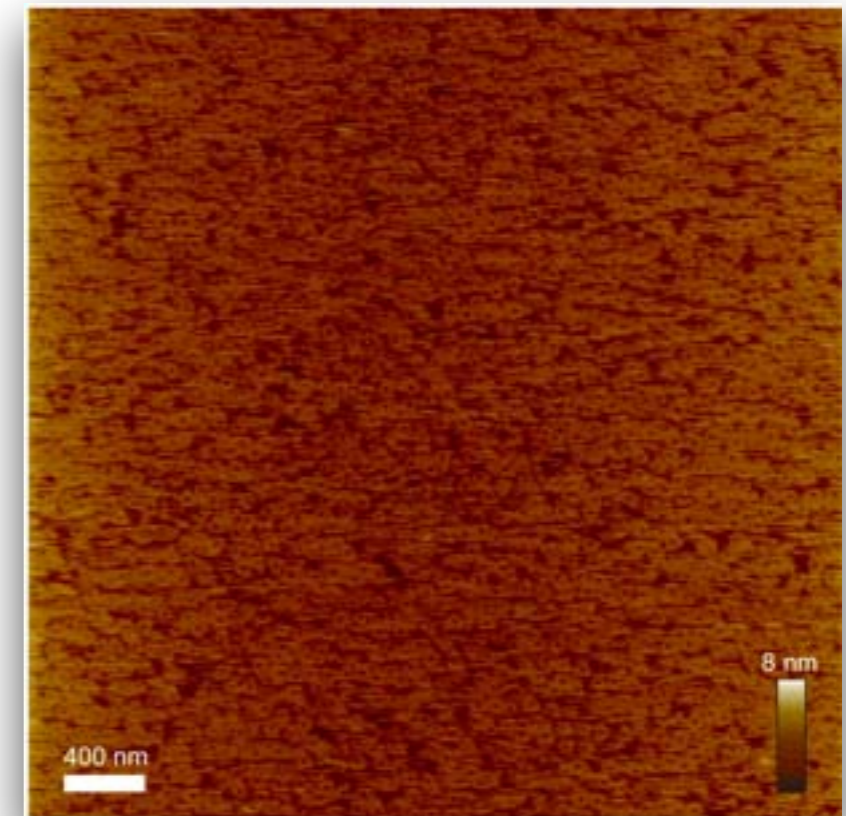
# Surface-assisted arrangement of DNA origami tiles



$[\text{Na}^+] = 100 \text{ mM}$



$[\text{Na}^+] = 120 \text{ mM}$



$[\text{Na}^+] = 200 \text{ mM}$

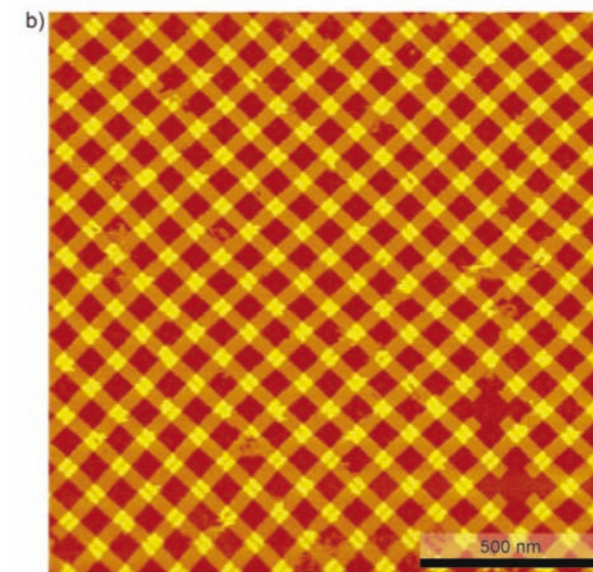
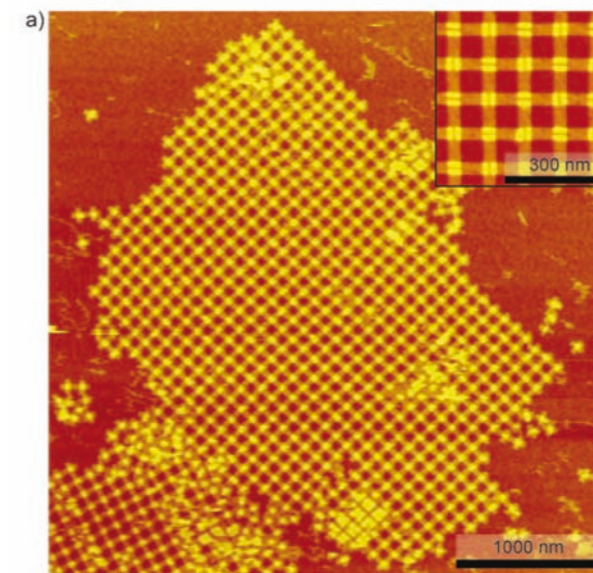
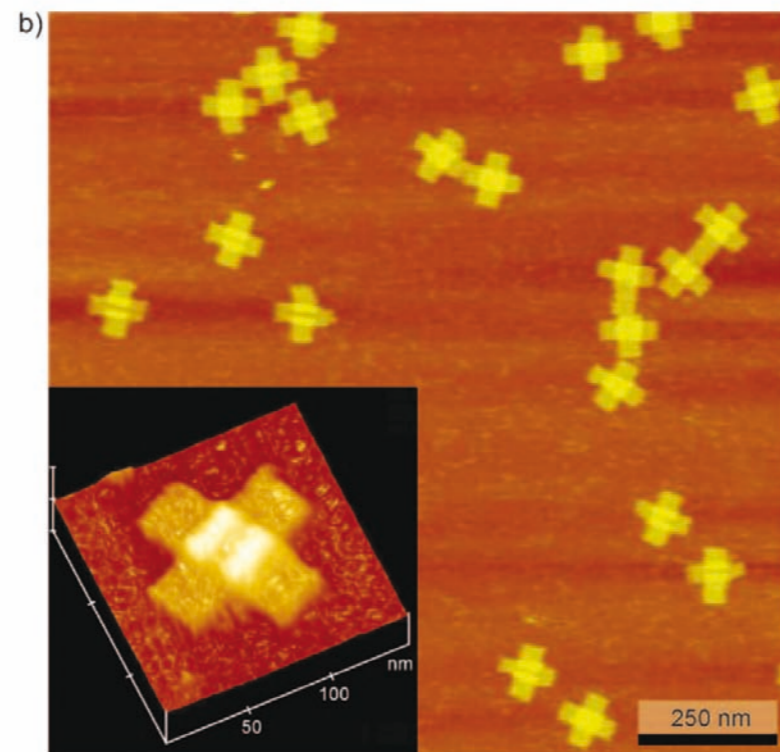
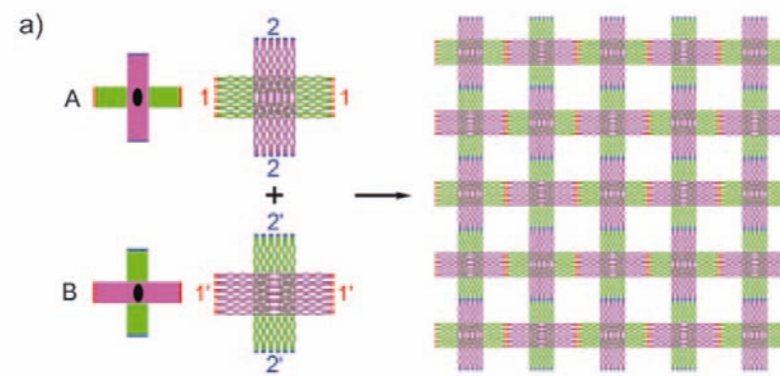
# Cross-shaped DNA origami tiles

VIP DNA Nanotechnology

DOI: 10.1002/anie.201005911

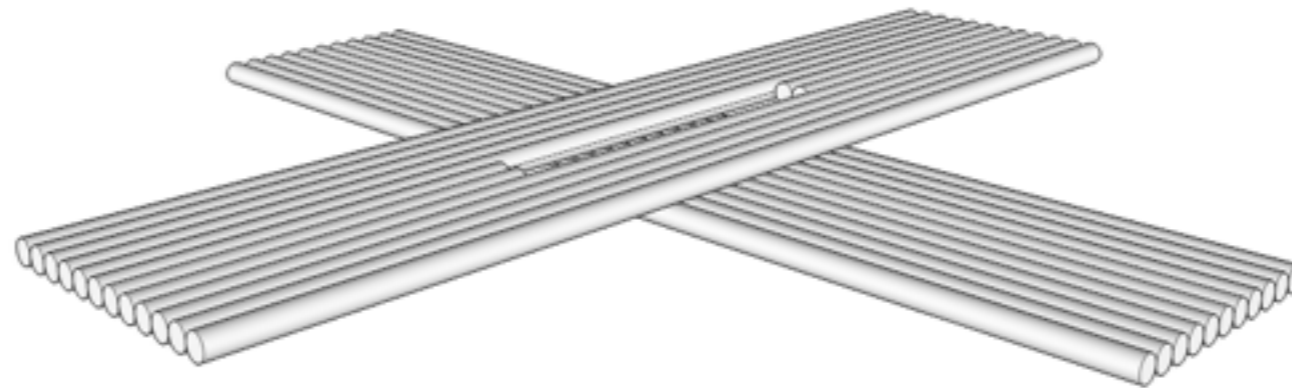
## Crystalline Two-Dimensional DNA-Origami Arrays\*\*

Wenyan Liu, Hong Zhong, Risheng Wang, and Nadrian C. Seeman\*

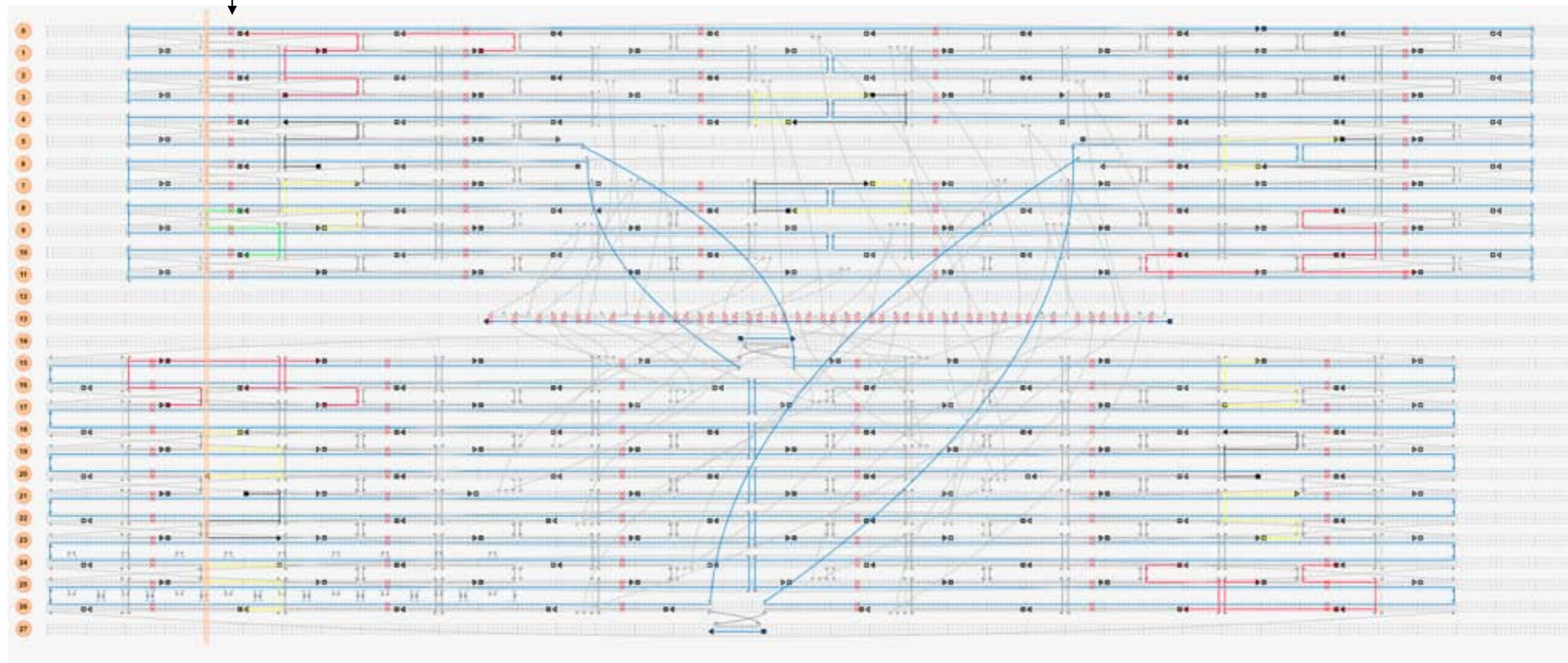


# A DNA origami cross tile

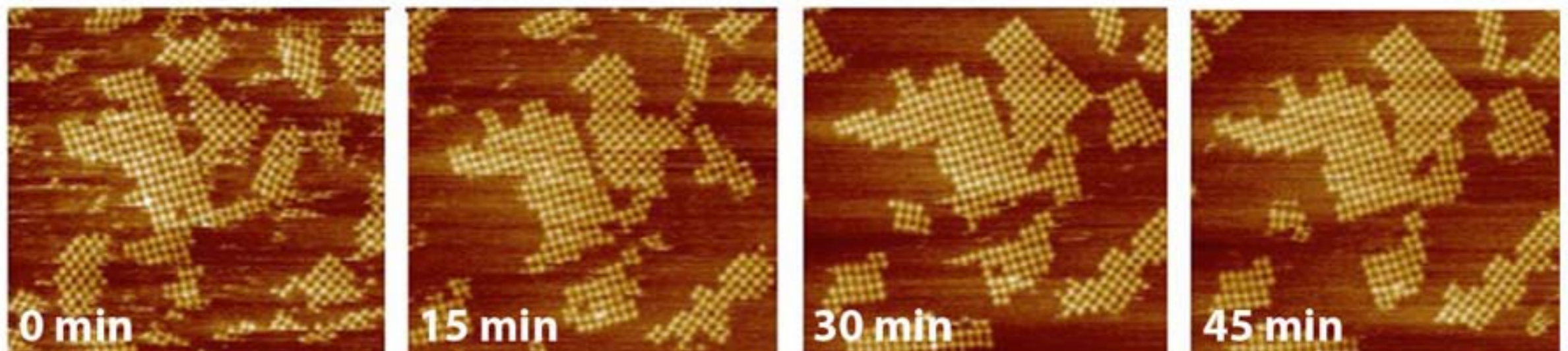
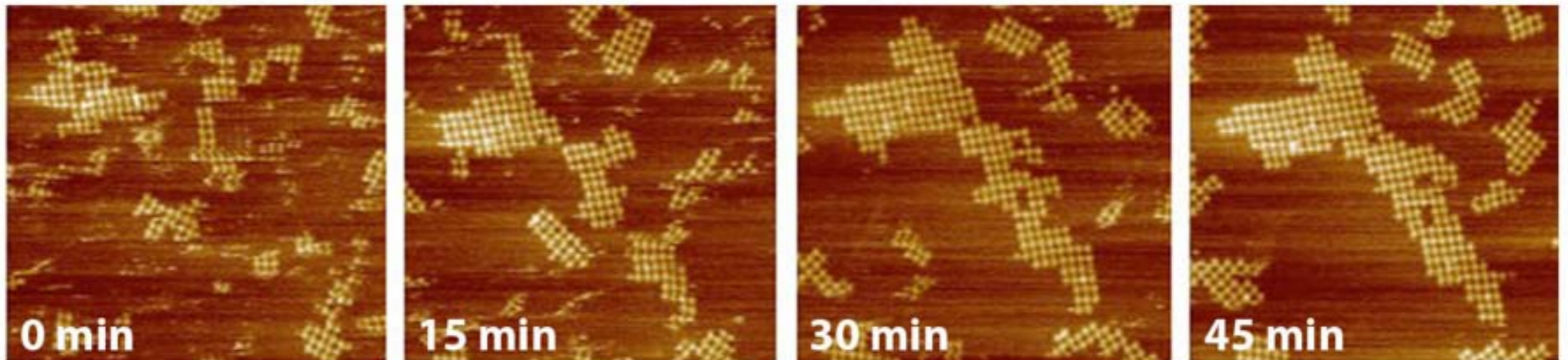
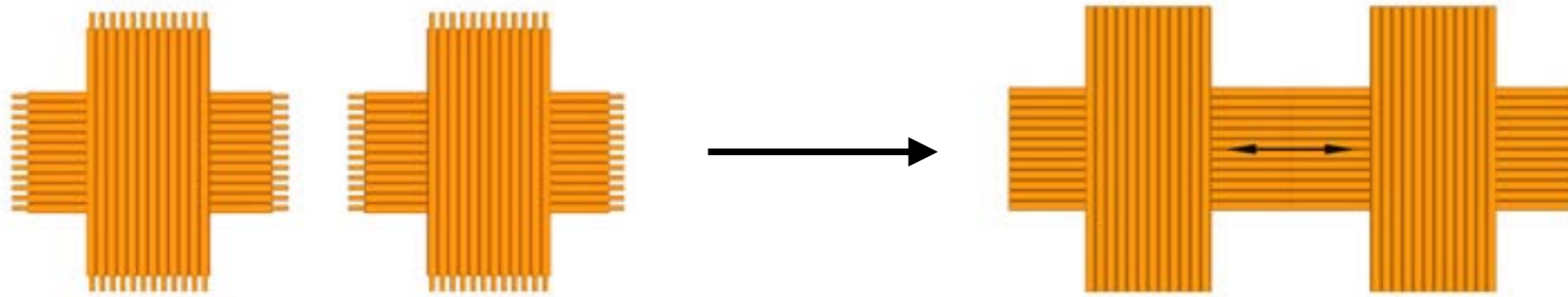
(cf. W. Liu, H. Zhong, R. Wang, N. C. Seeman, *Angewandte Chemie* 2011, 123, 278-281)



twist corrected and with adjusted staple crossover positions

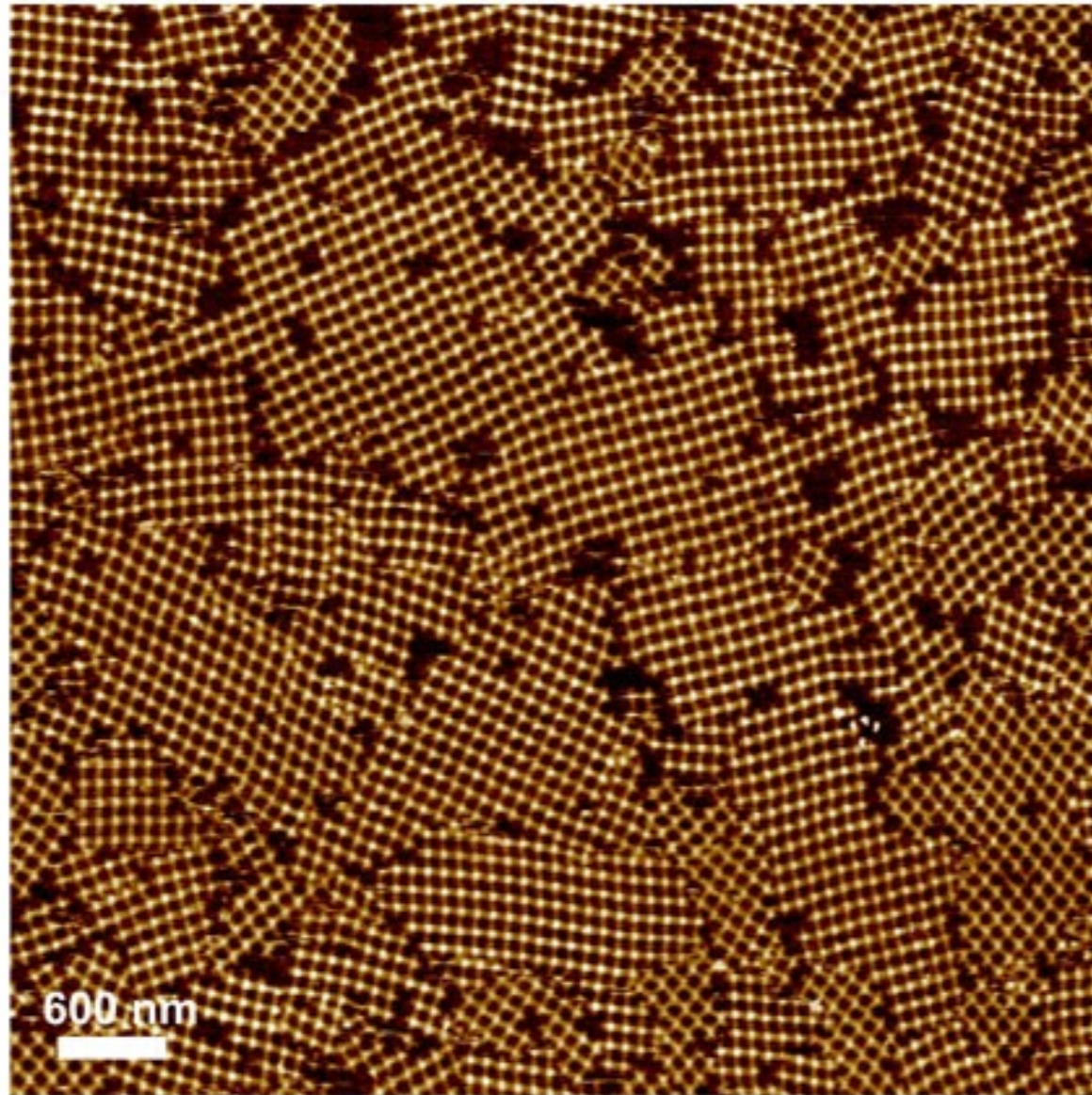


# Utilizing stacking interactions

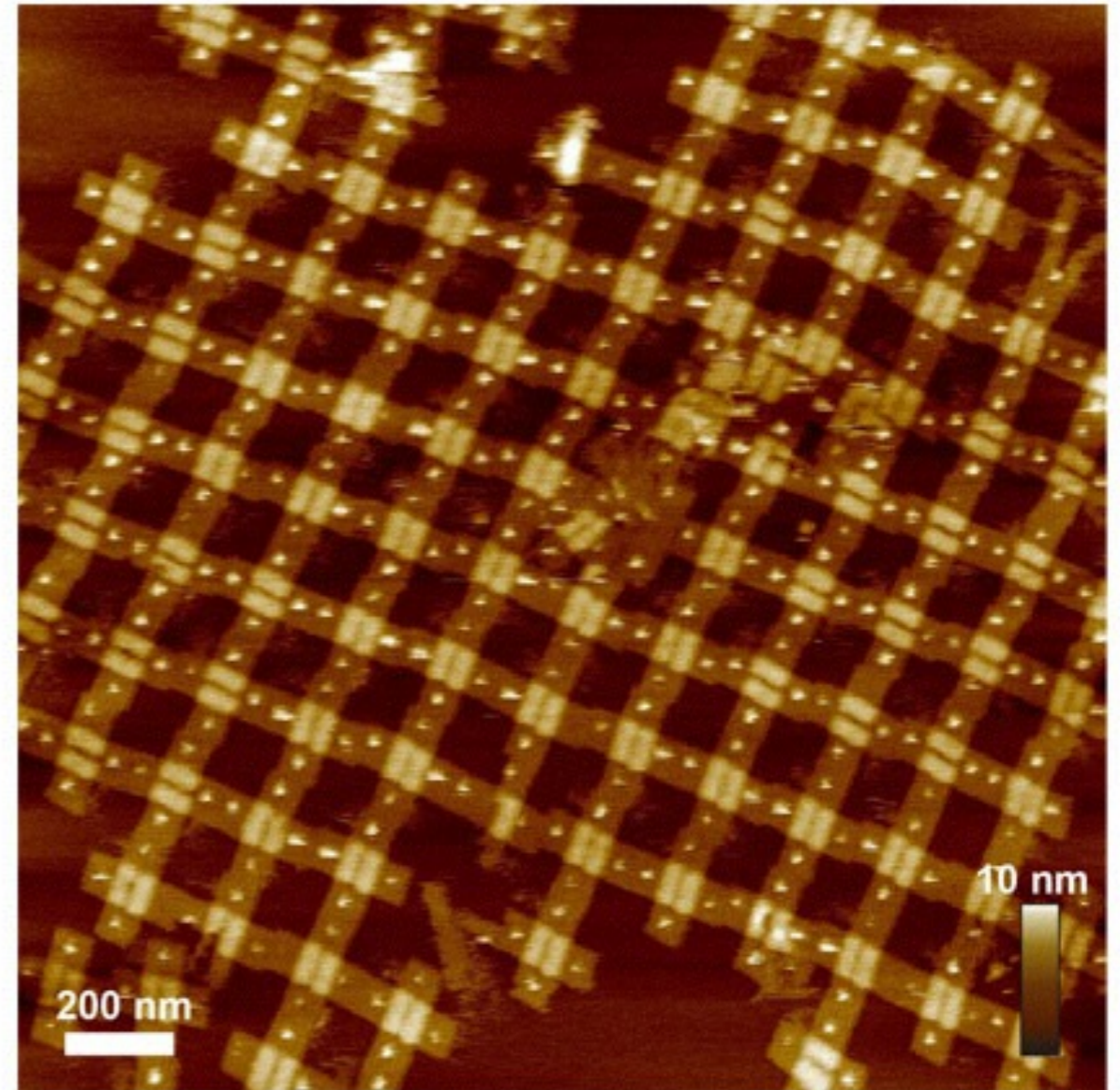


# Utilizing stacking interactions

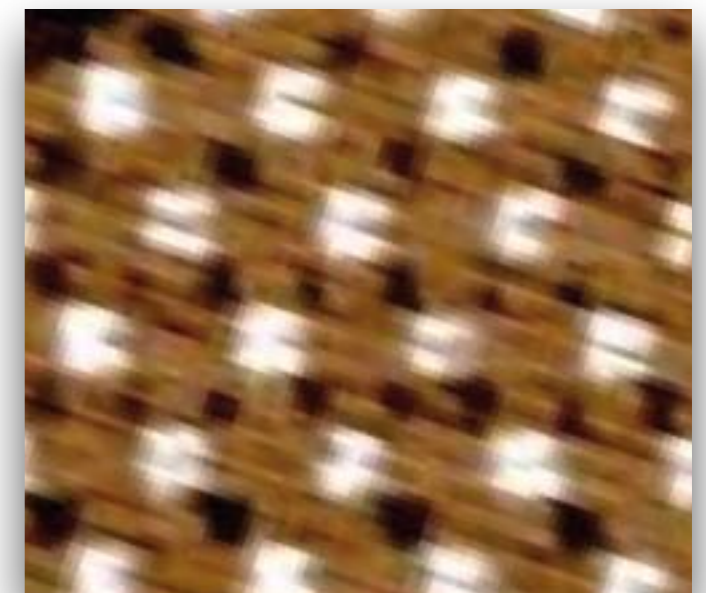
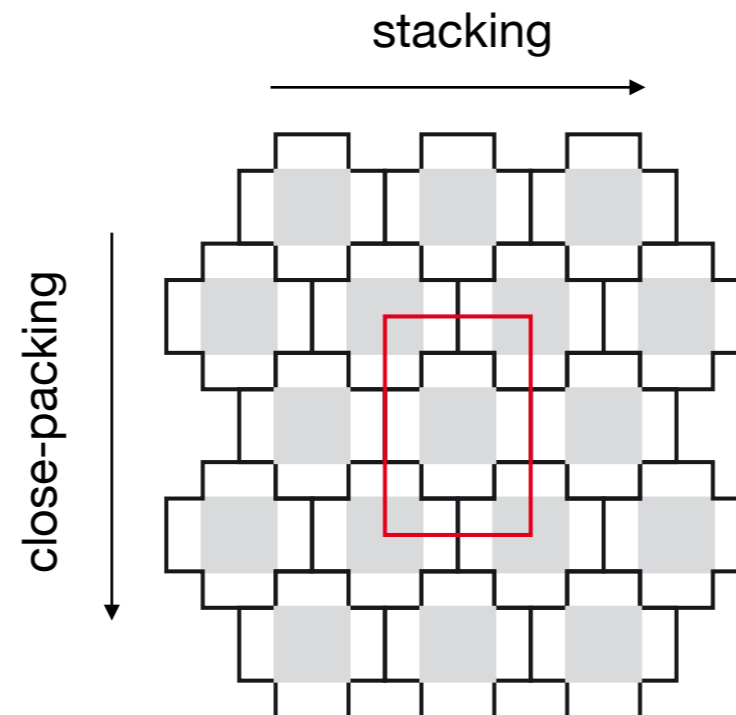
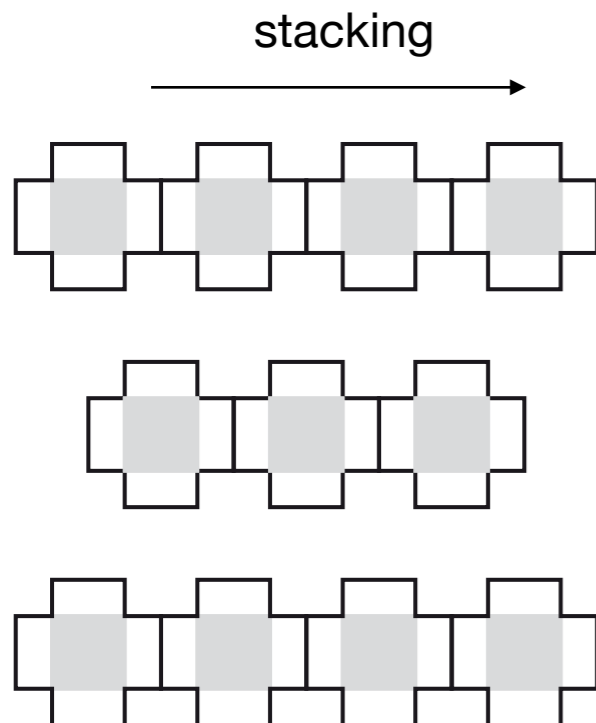
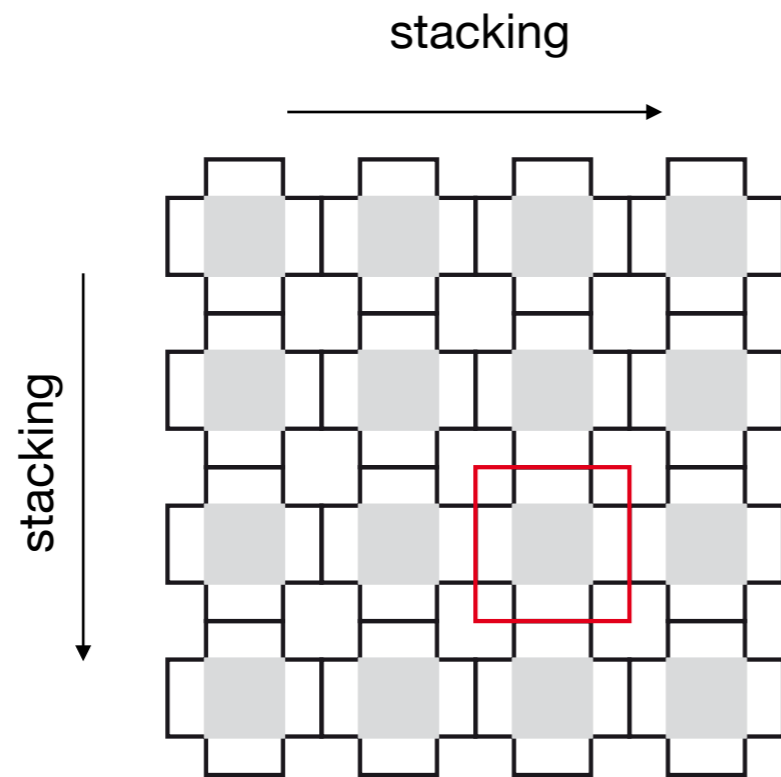
A



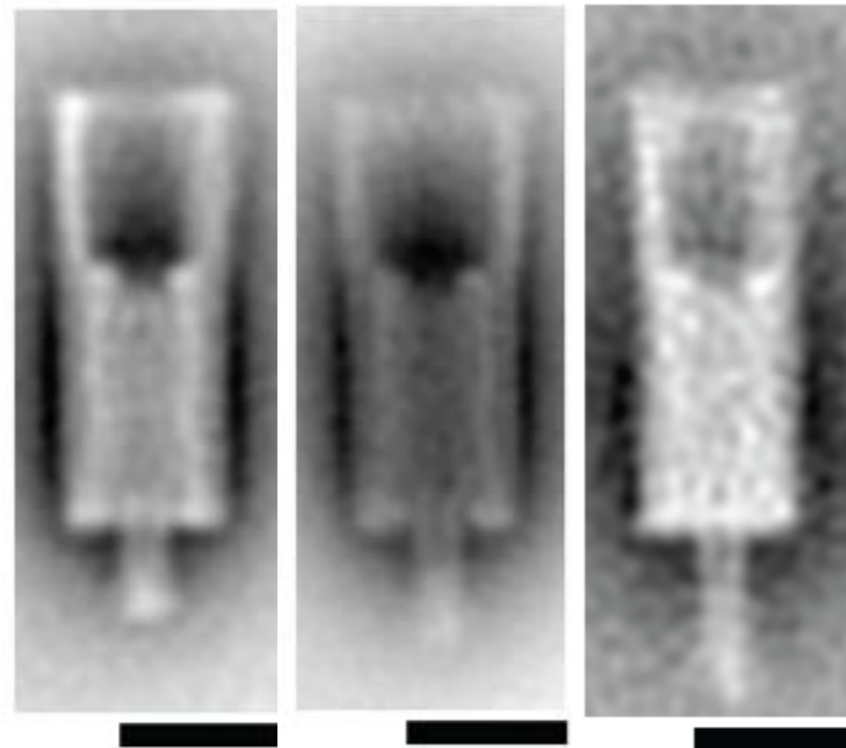
B



# Surface-assisted arrangement of DNA origami tiles



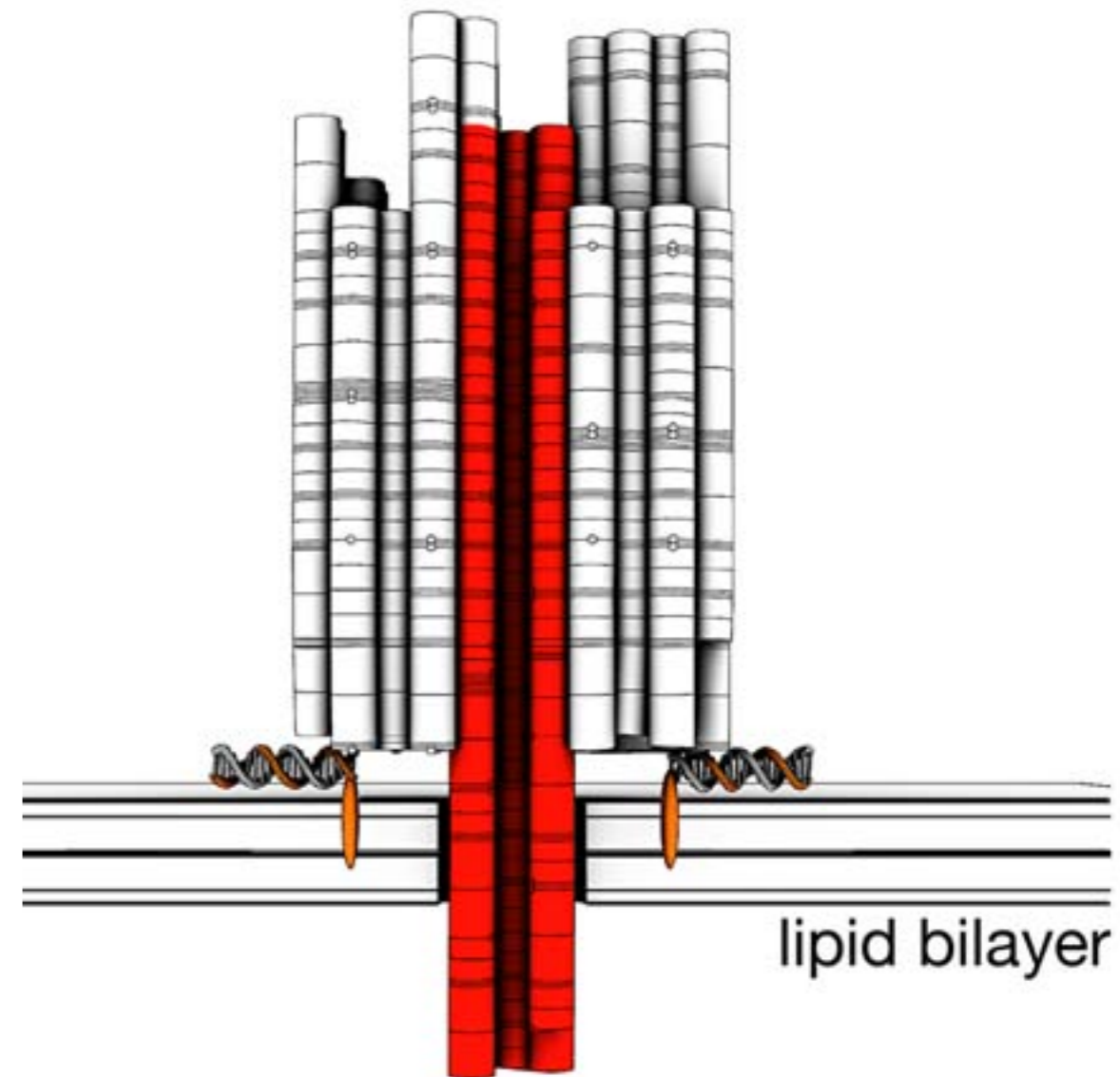
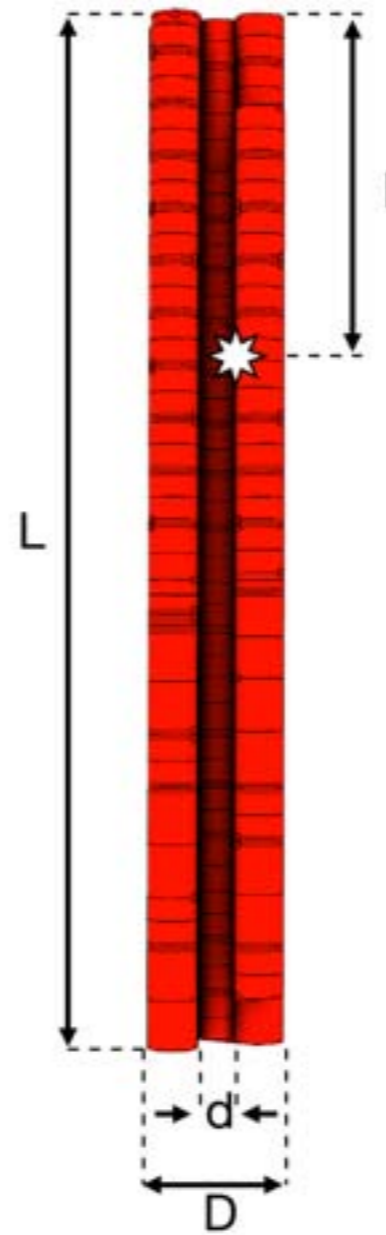
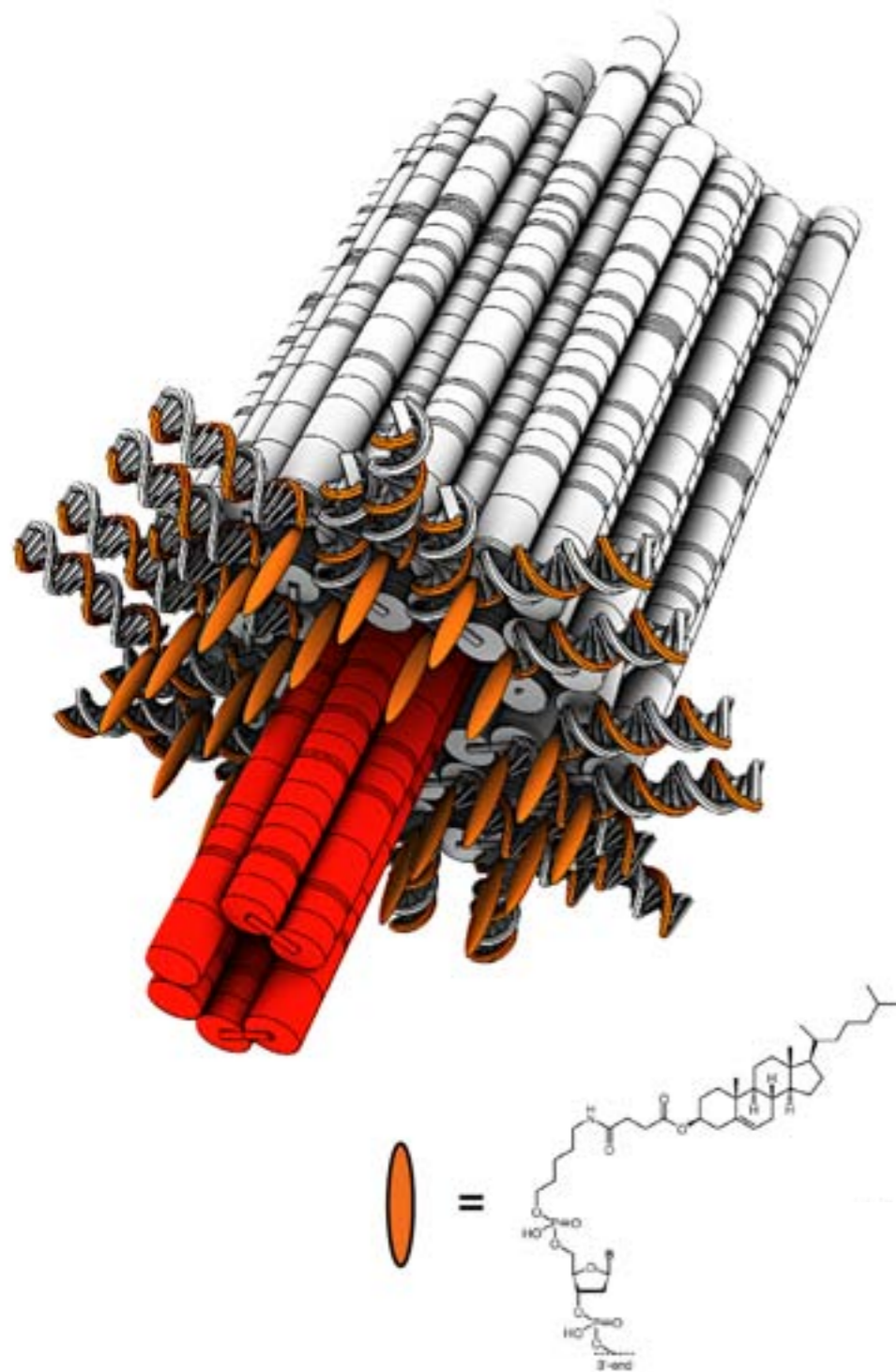
# DNA-based membrane channels



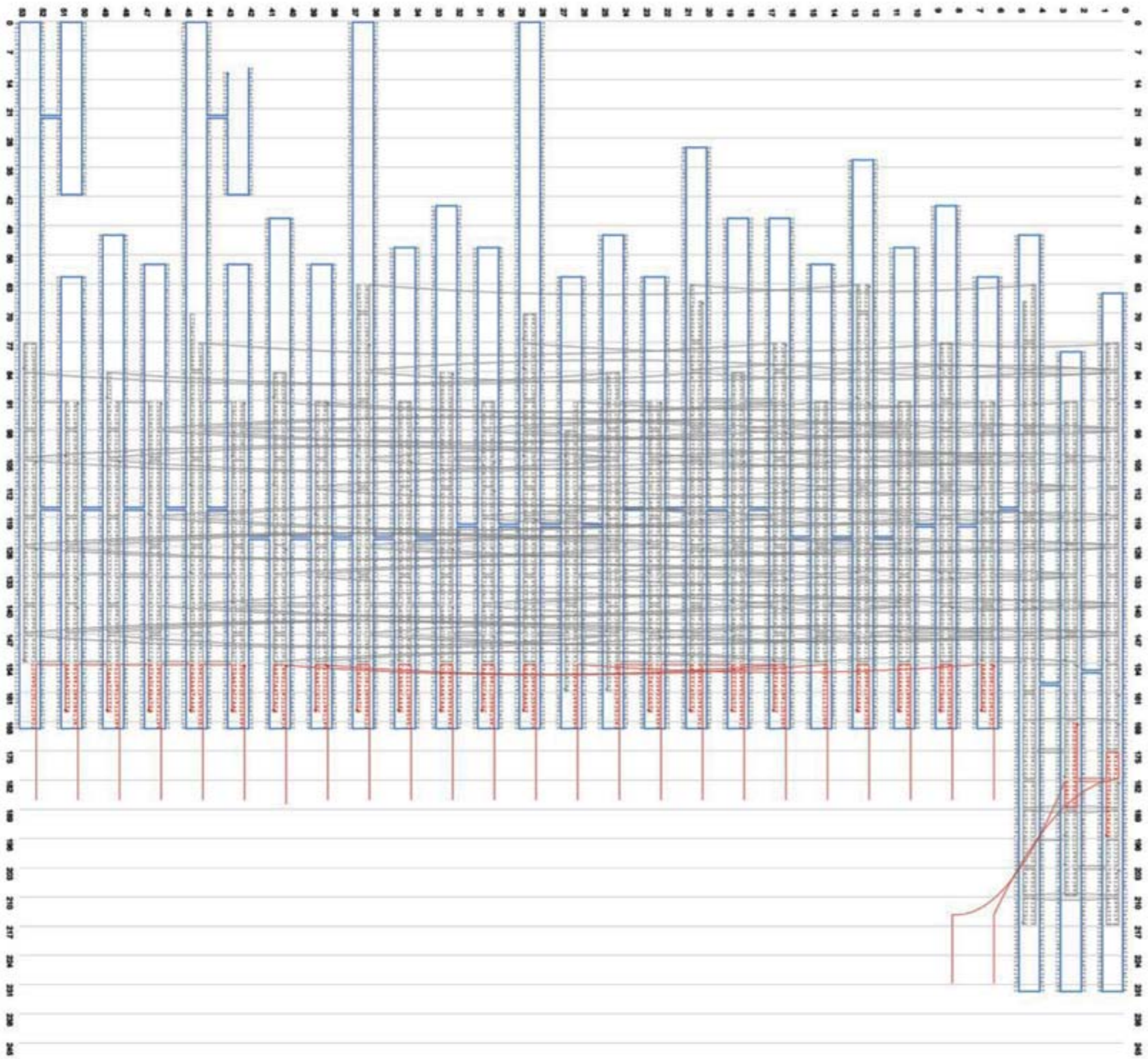
Langecker, M., Arnaut, V., Martin, T. G., List, J., Renner, S., Mayer, M., Dietz, H., Simmel, F. C.,  
Synthetic Lipid Membrane Channels Formed by Designed DNA Nanostructures, *Science* **338**, 932-936 (2012).



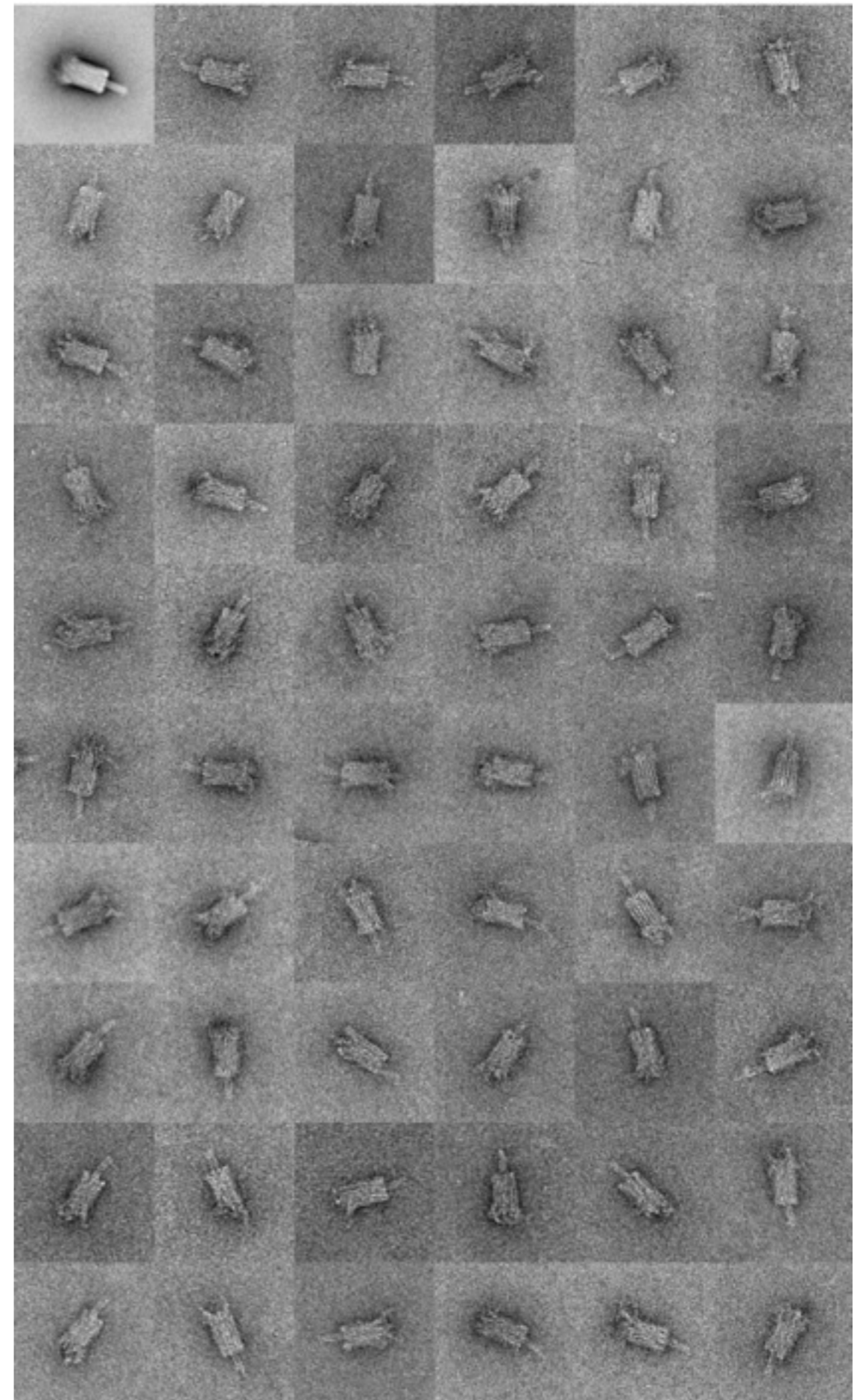
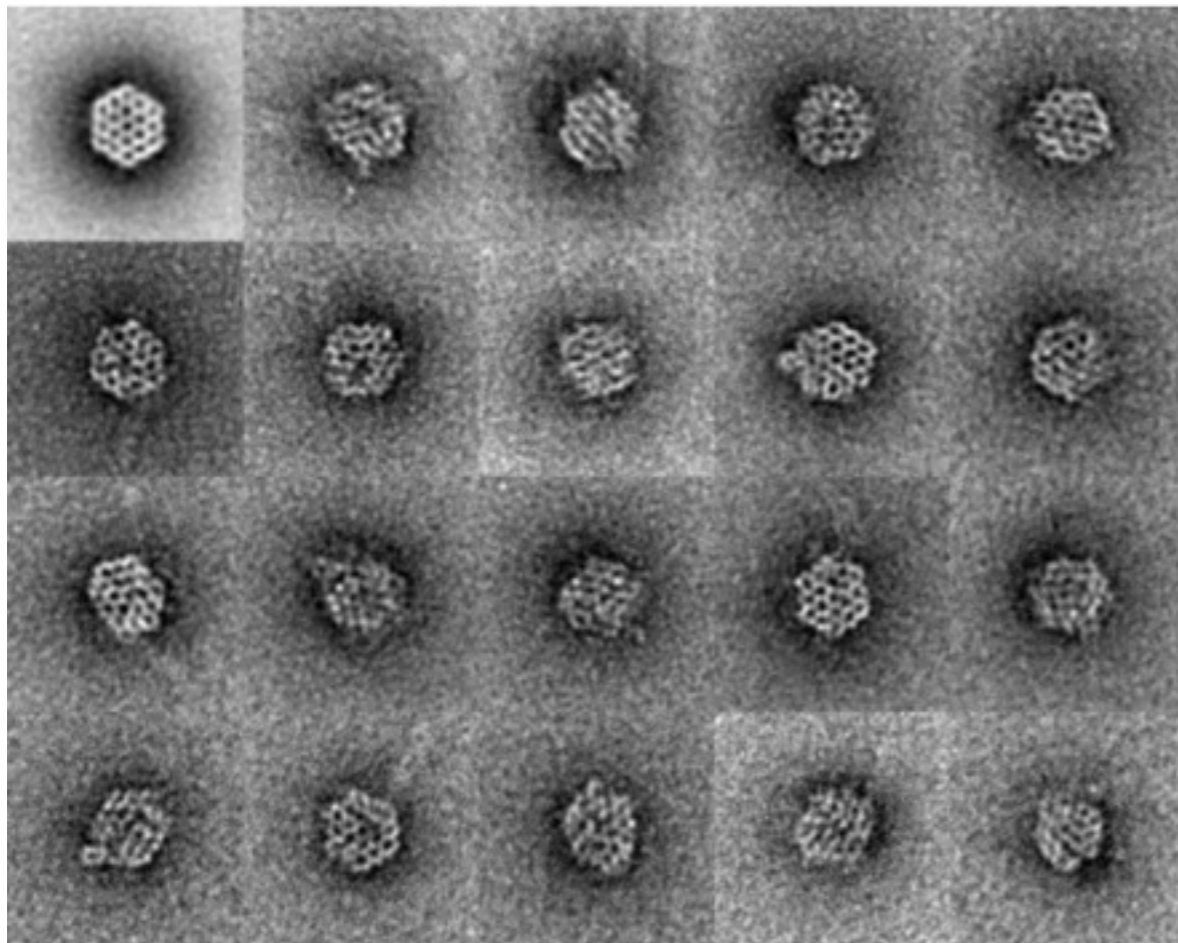
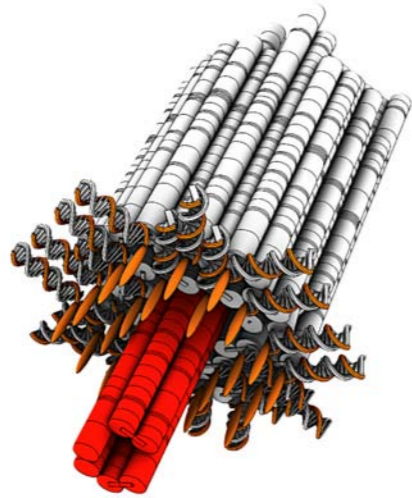
# A synthetic membrane channel made from DNA



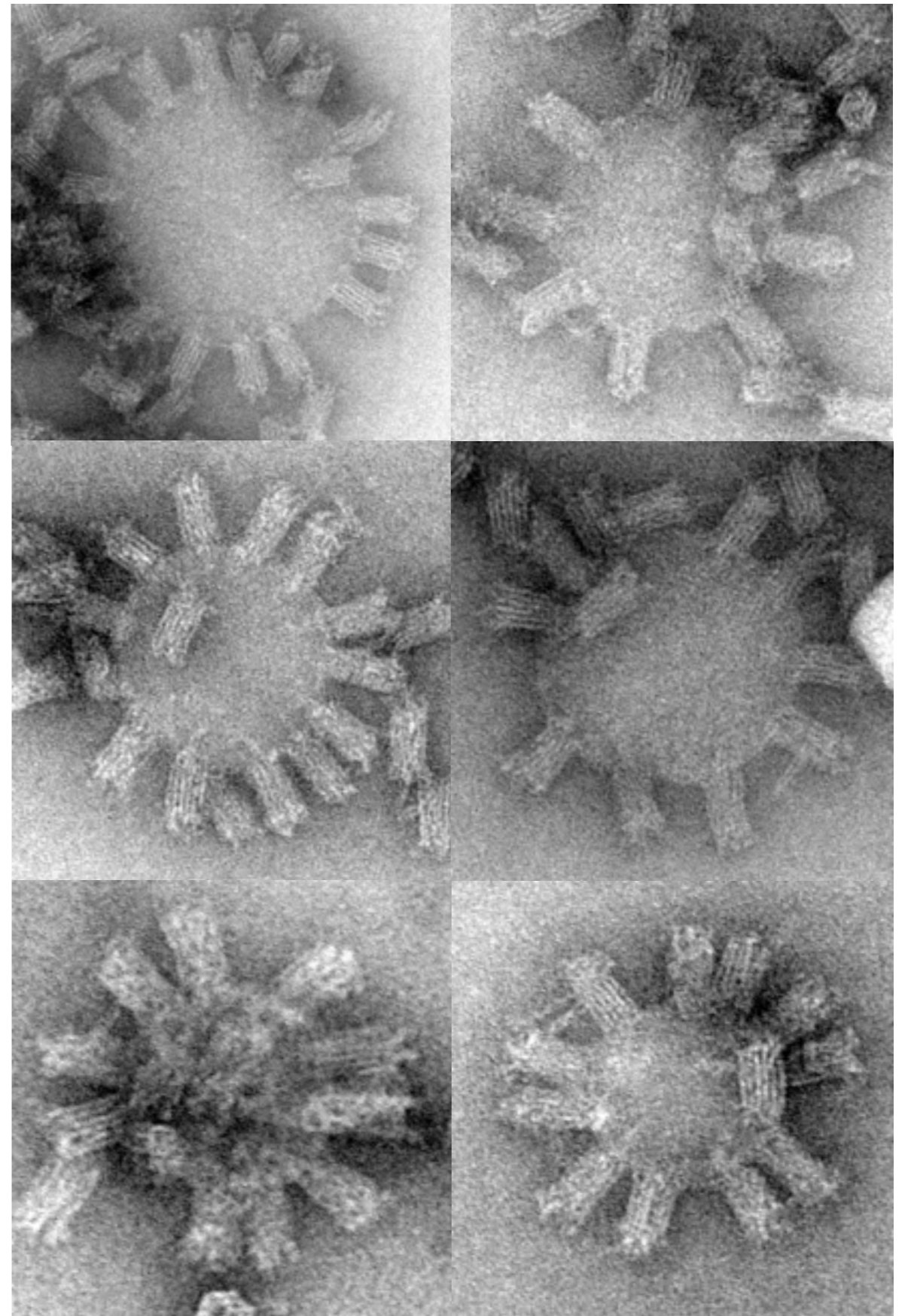
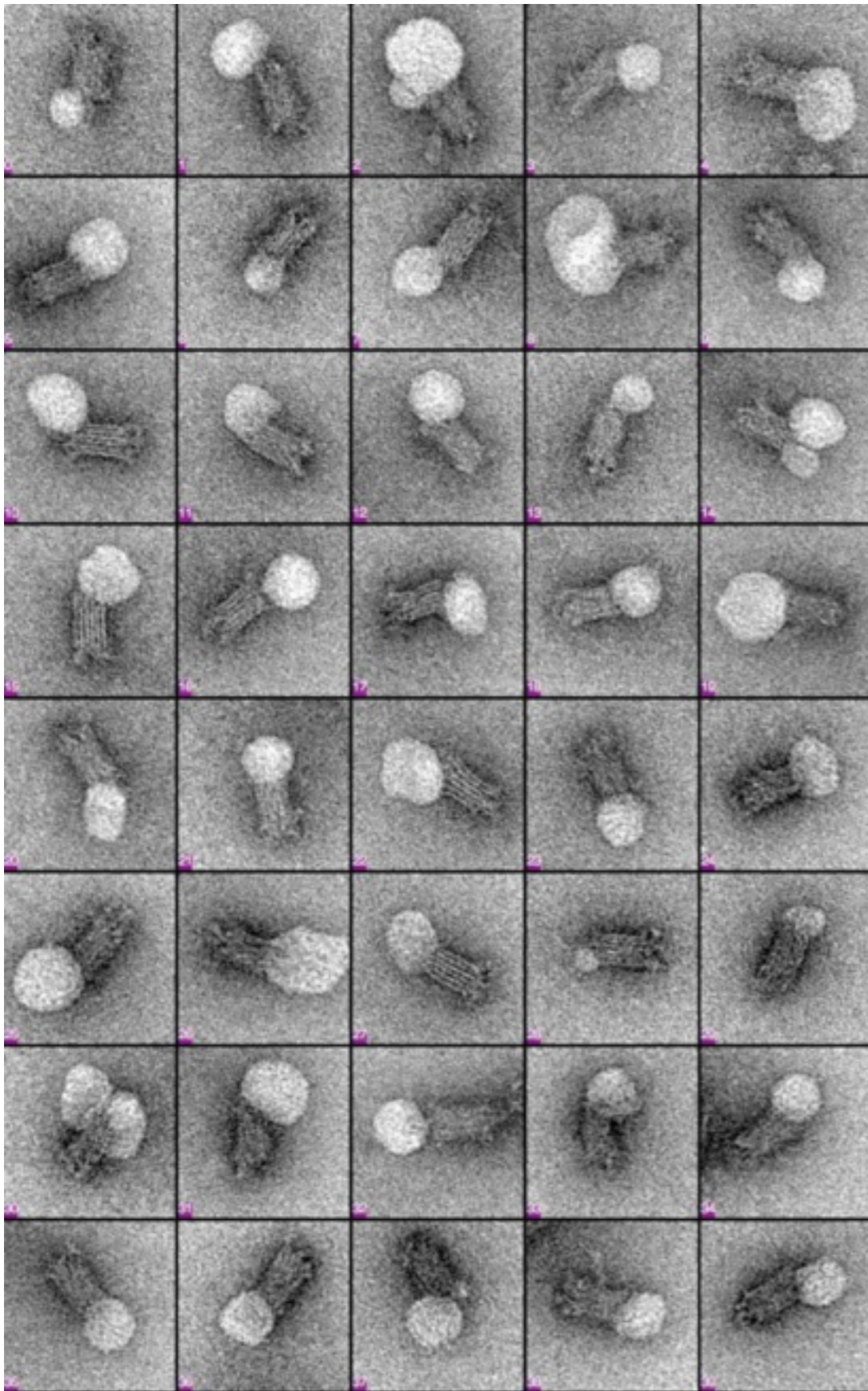
$L=47\text{ nm}, D=6\text{ nm}, d=2\text{ nm}$



# Quality control

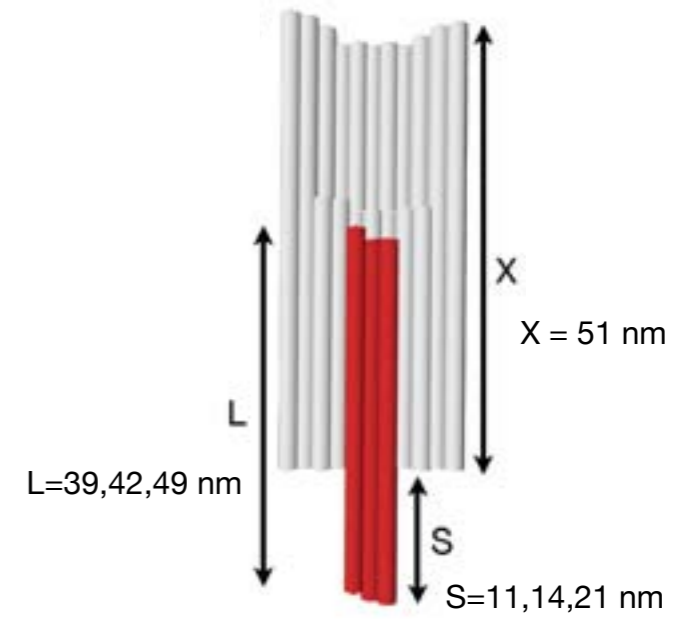
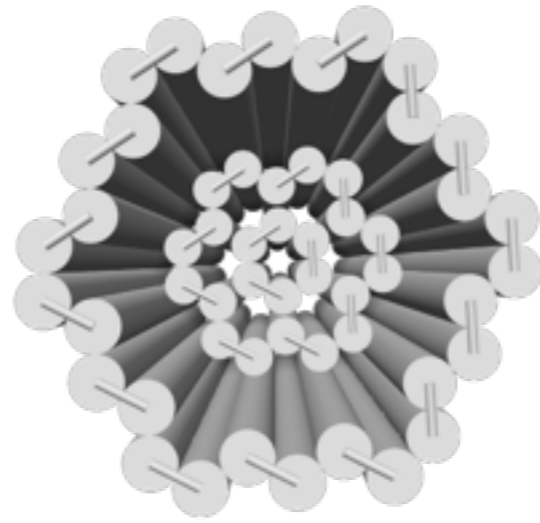


# Binding of DNA channels to small vesicles (POPC)

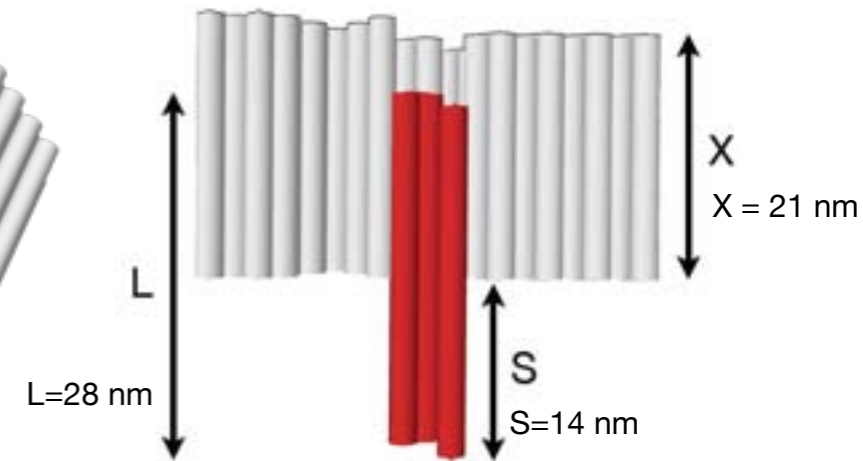
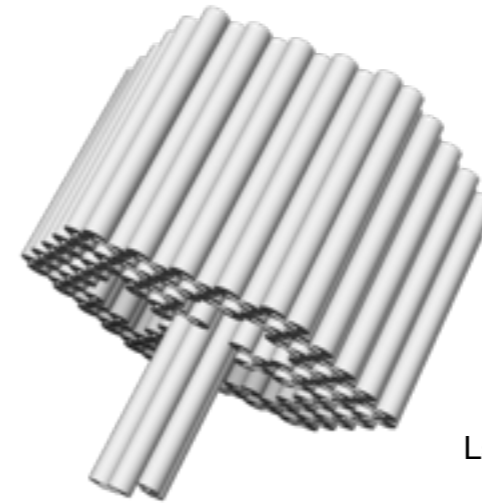
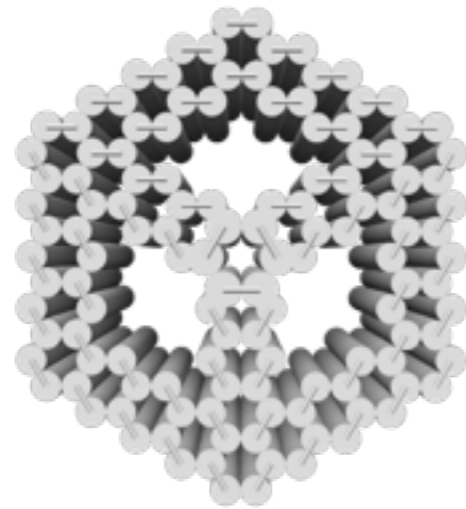


# Various pore geometries

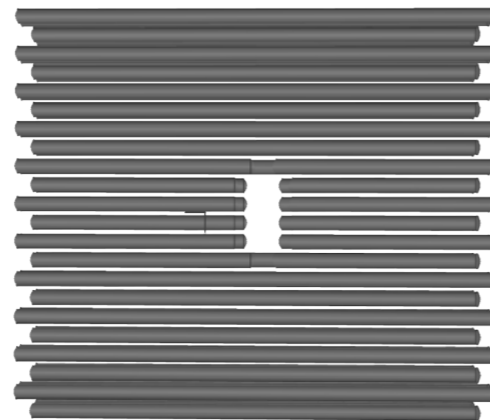
Pore with vestibule



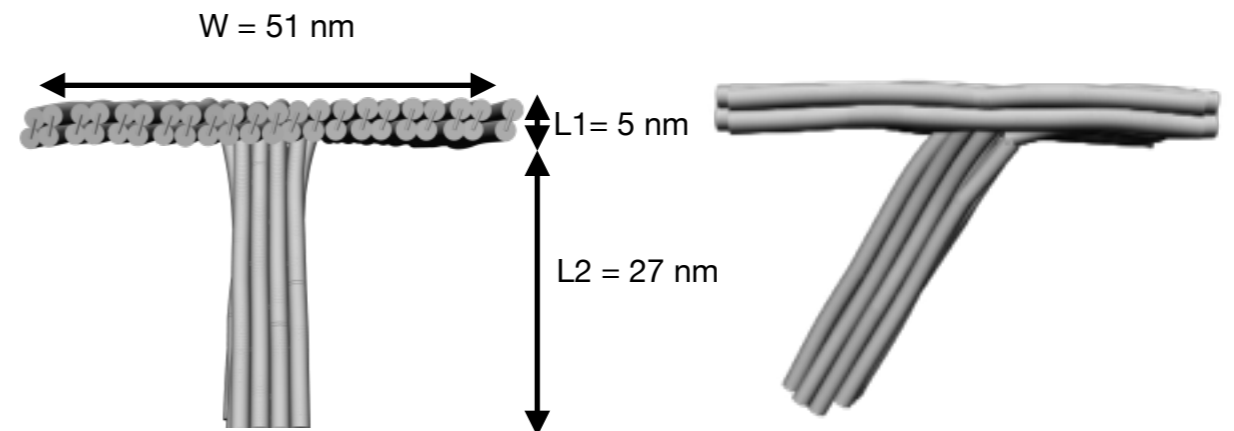
Flat & fat pore



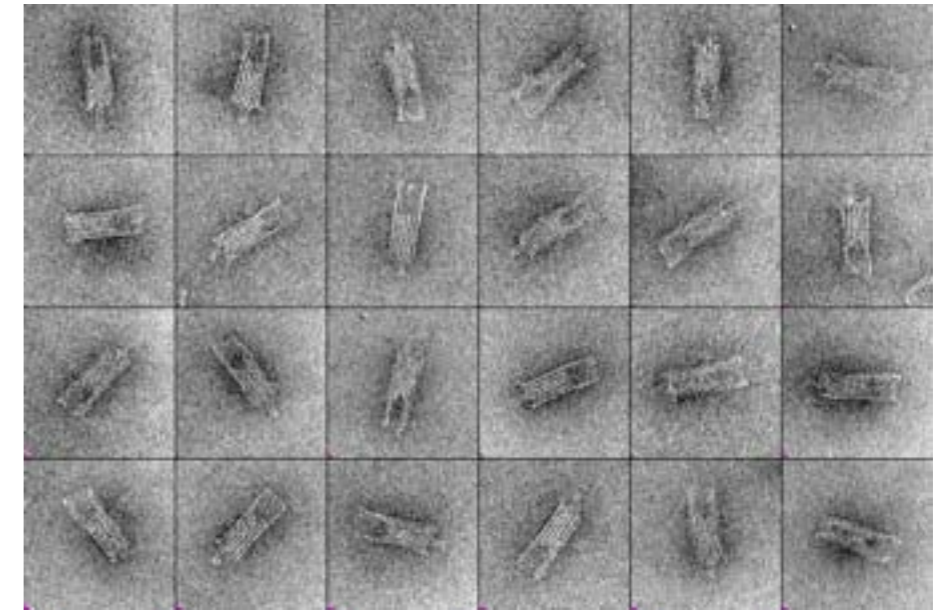
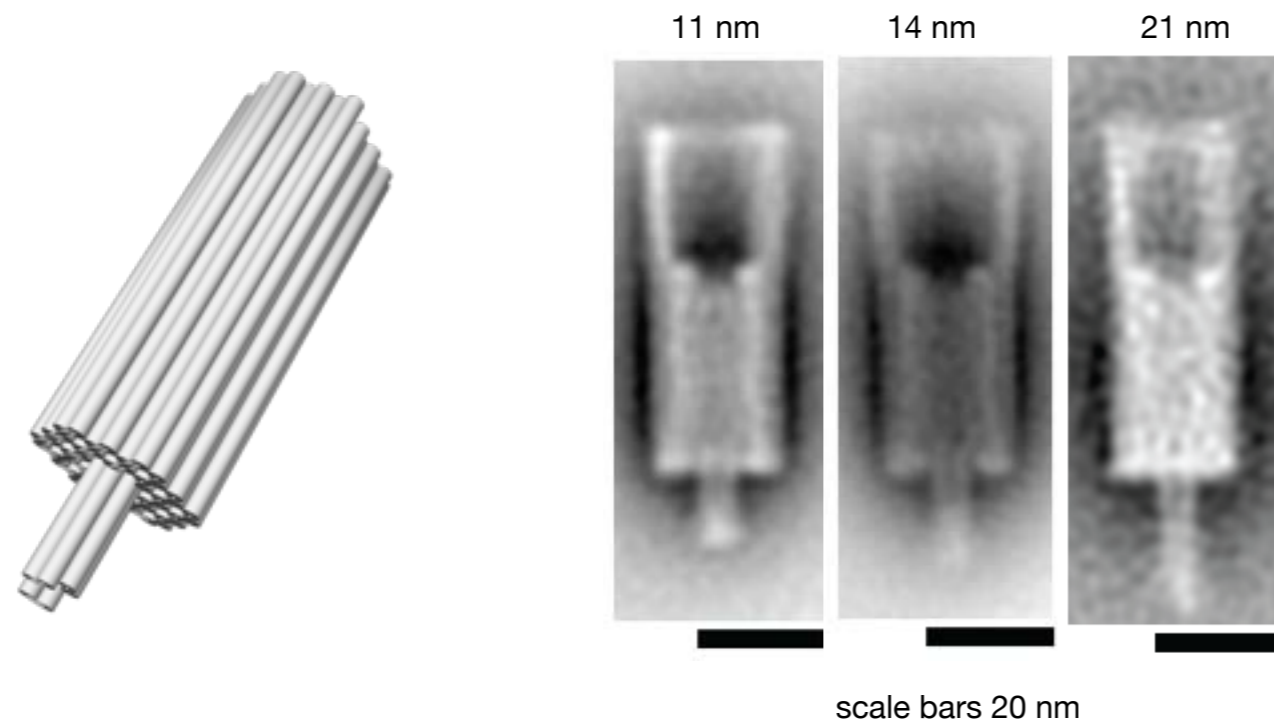
Square pore



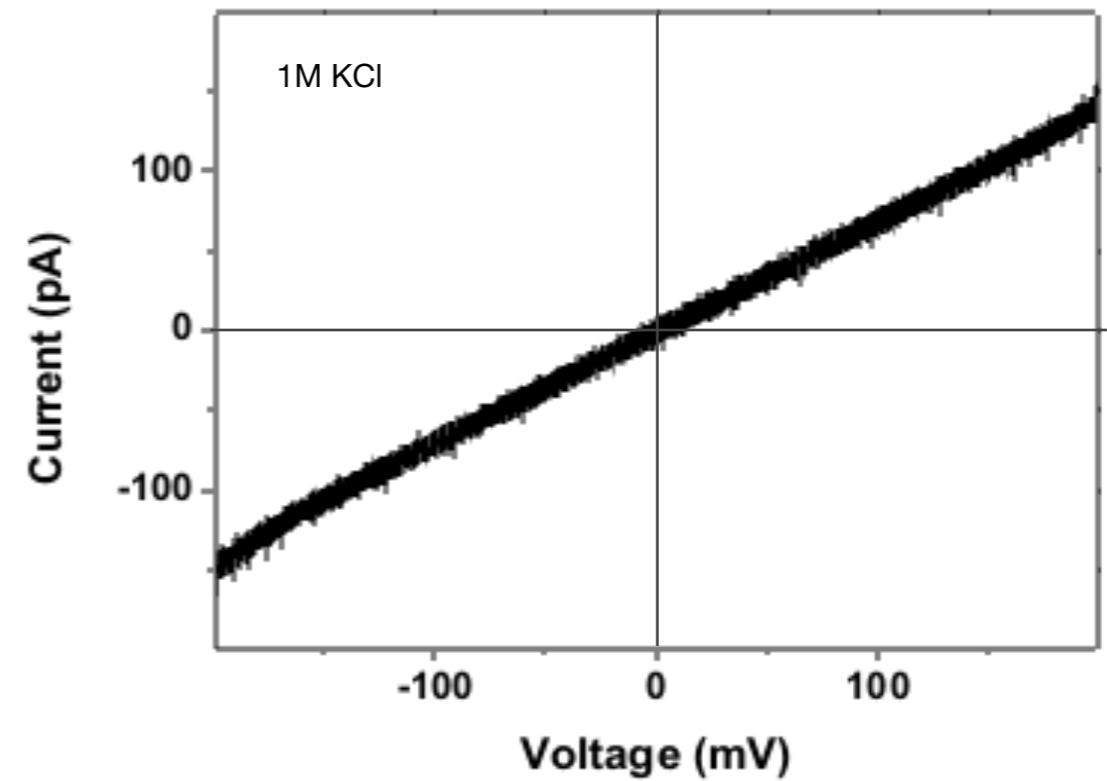
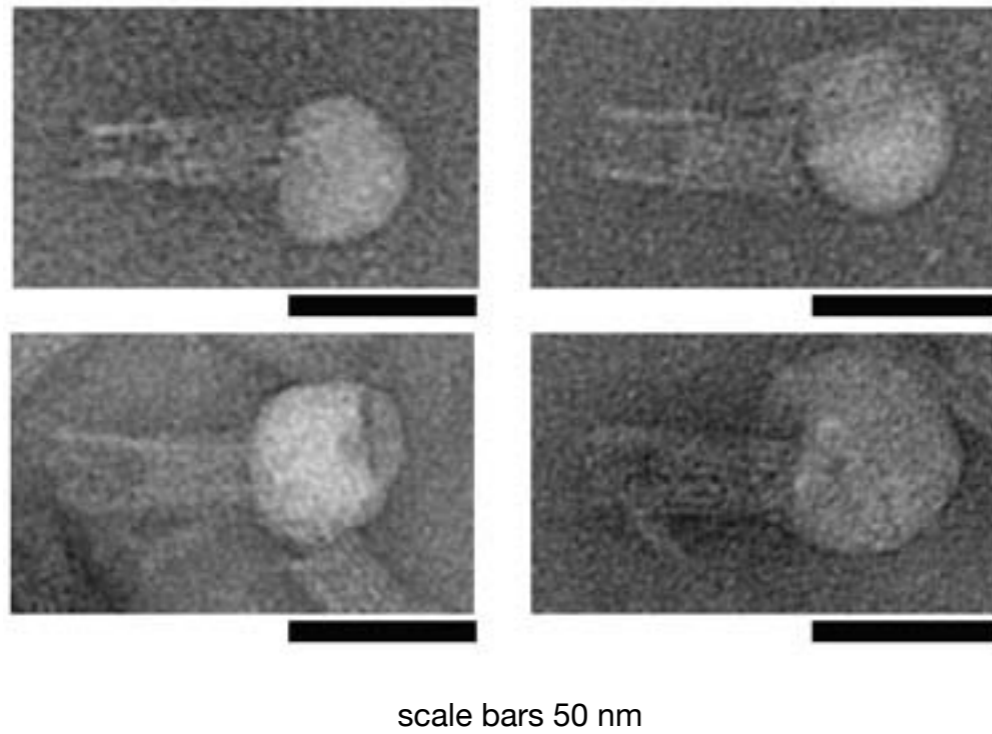
hole = 5 nm x 5 nm



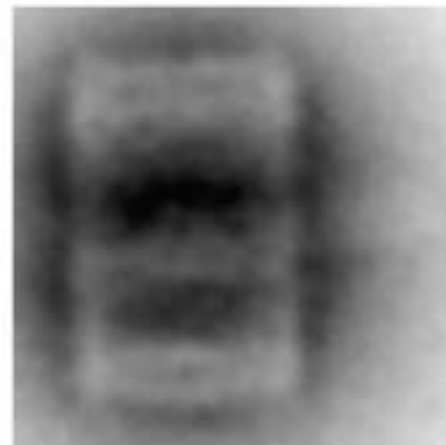
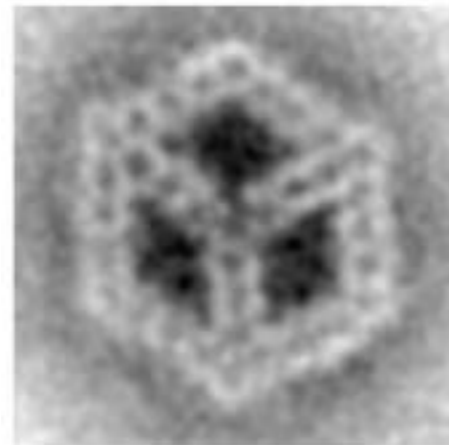
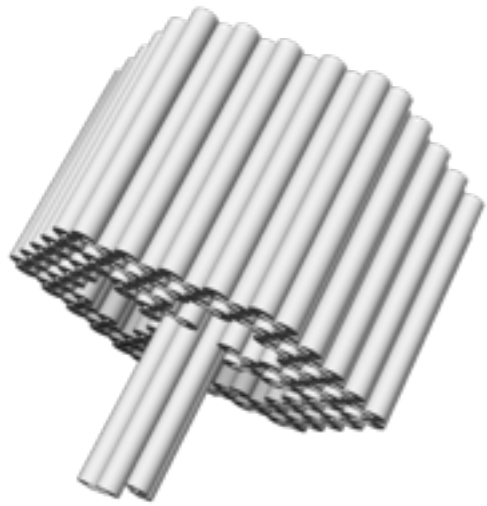
# Pore with vestibule



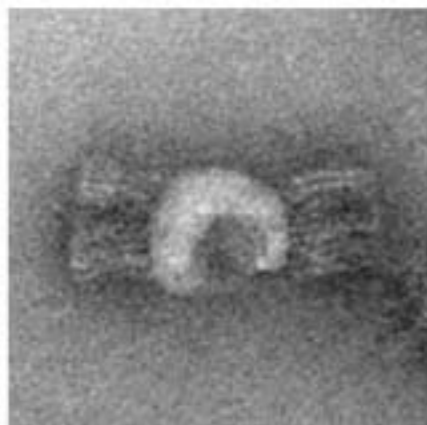
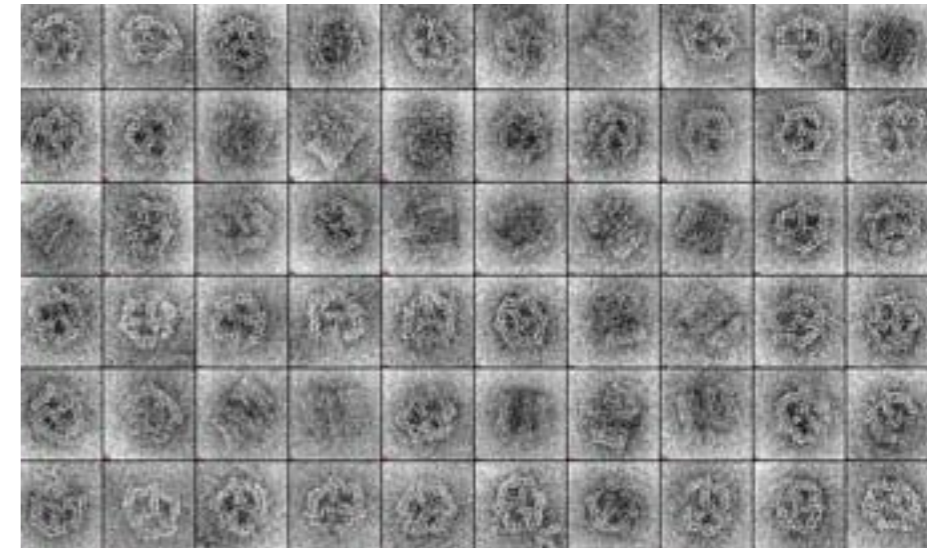
## I-V characteristics



# Flat & fat pore

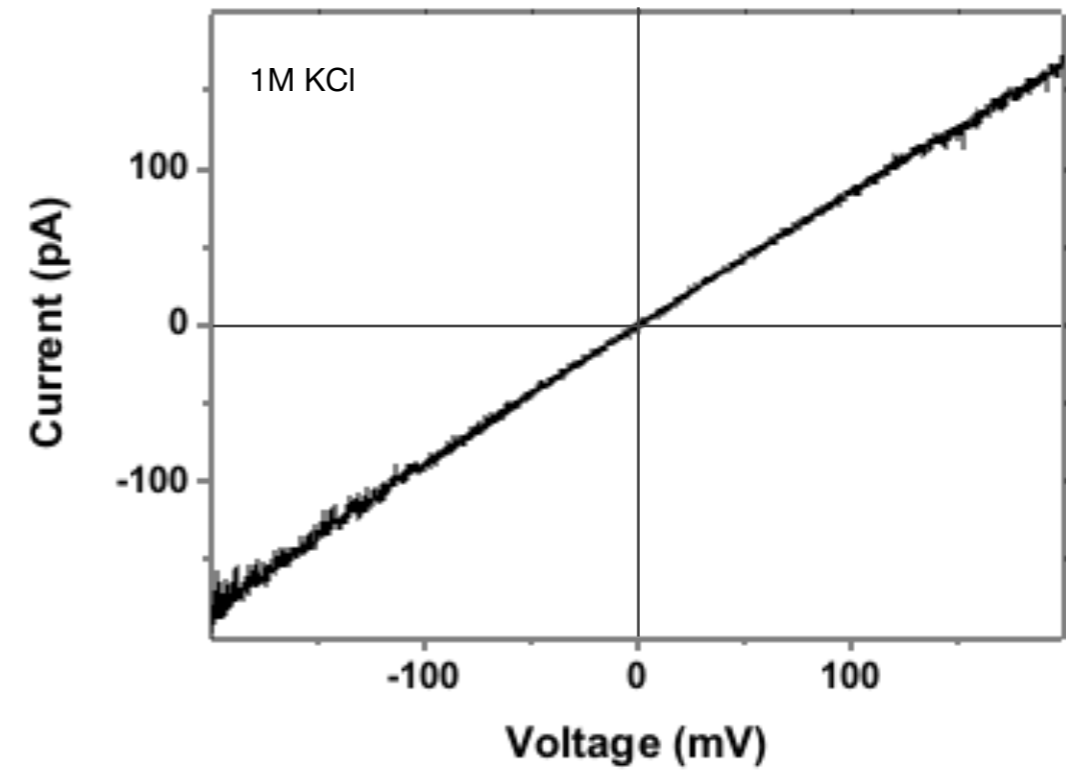


scale bars 20 nm

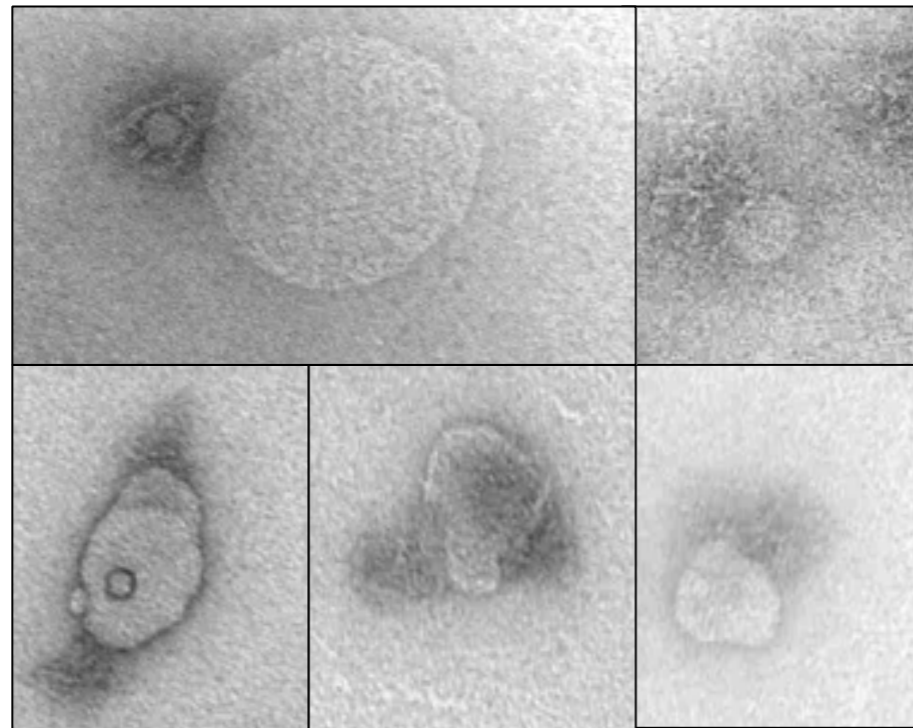
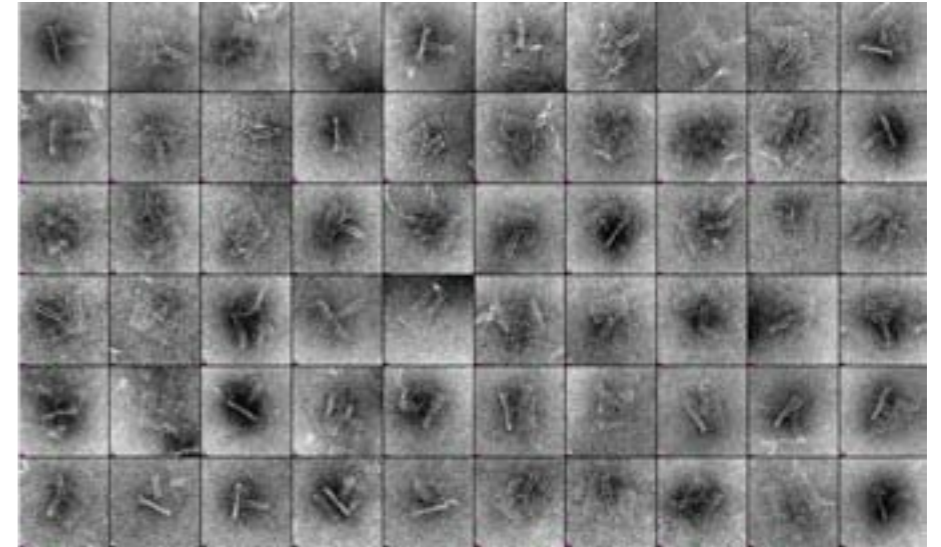
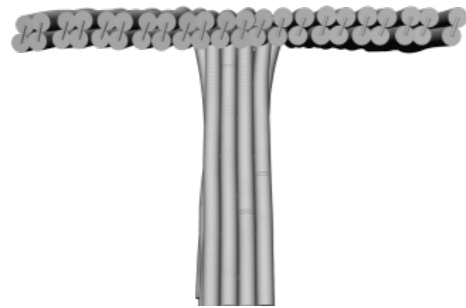


scale bars 50 nm

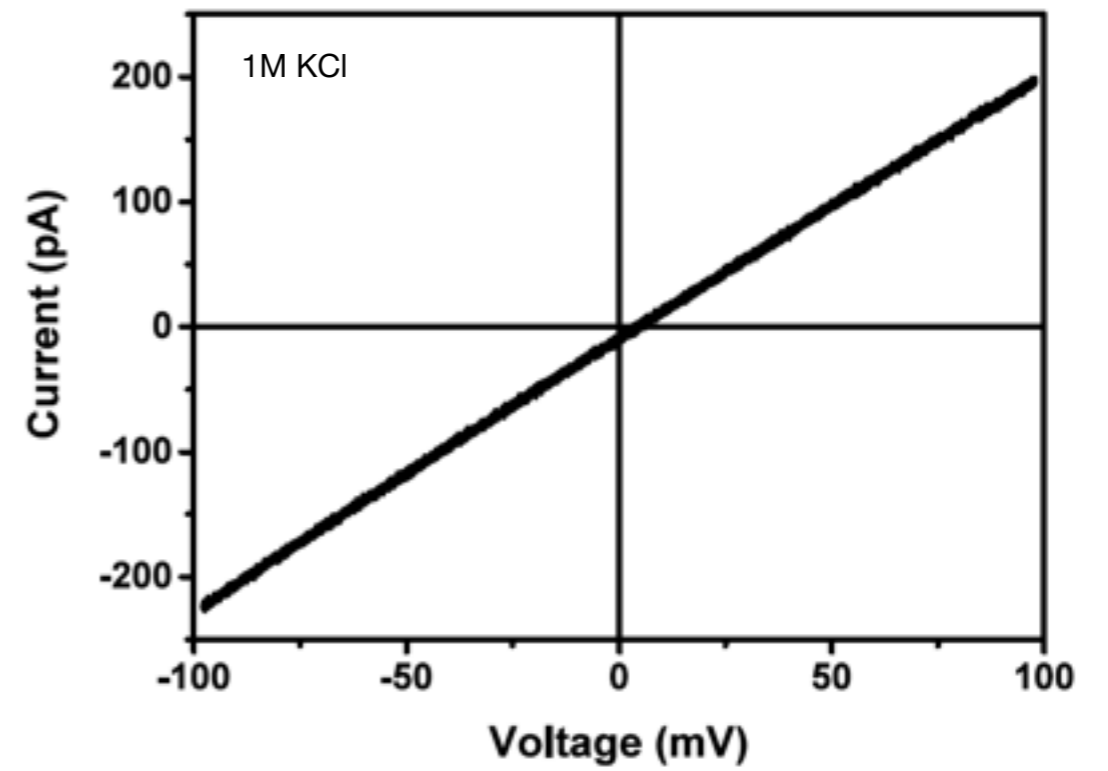
## I-V characteristics



# Square pore

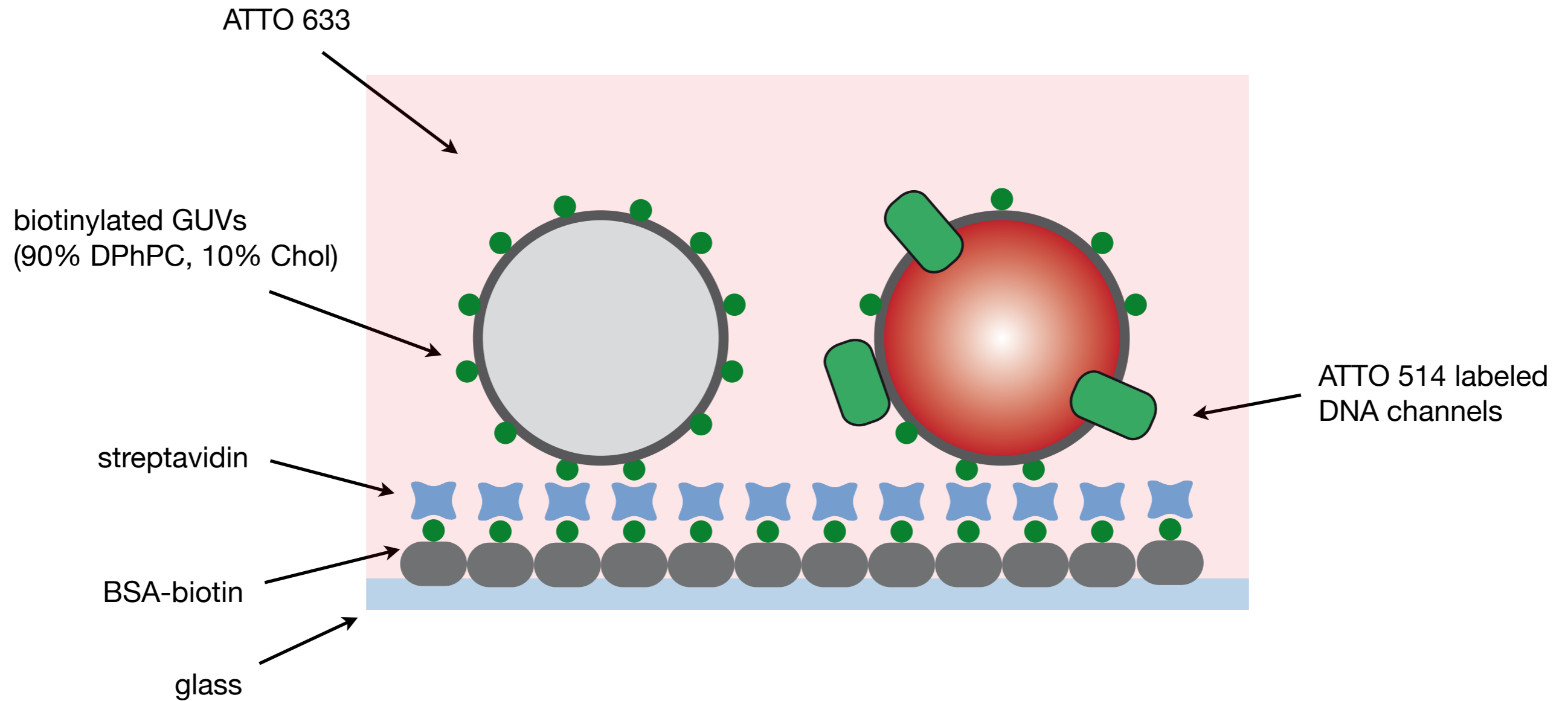


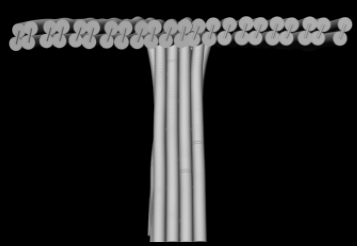
## I-V characteristics





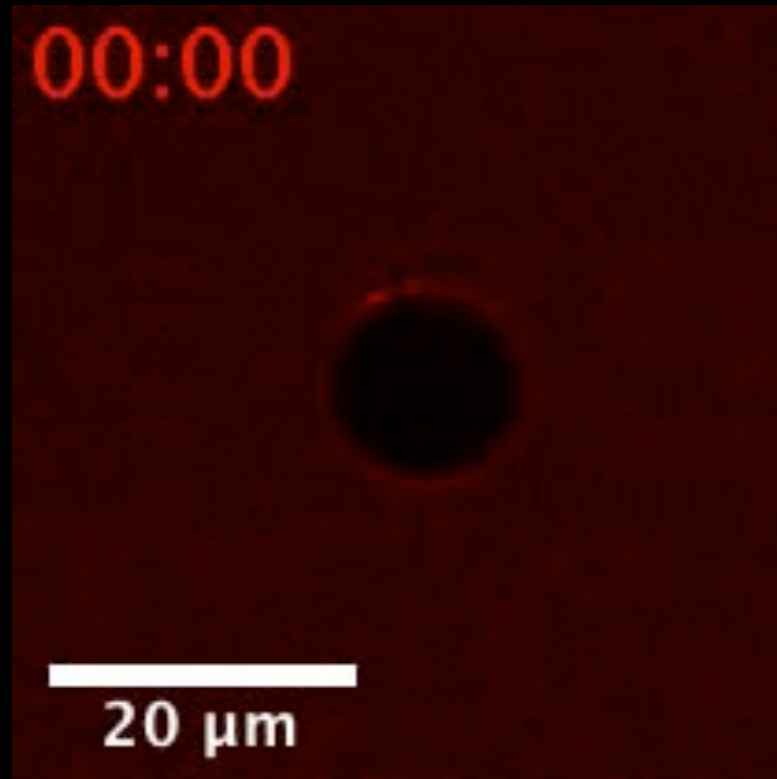
# Dye influx experiments with immobilized vesicles



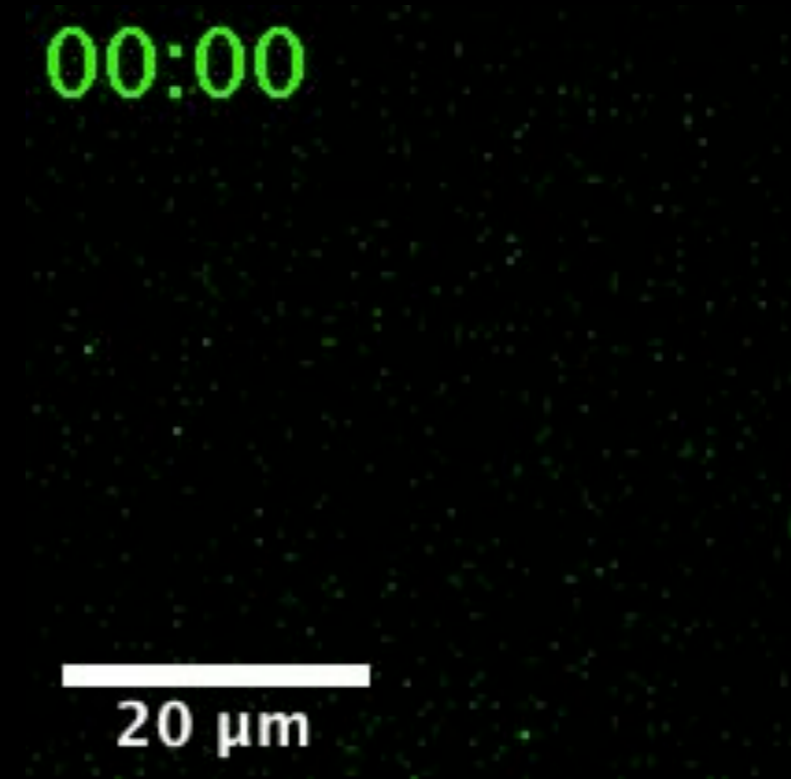


# Dye influx experiments with immobilized vesicles

ATTO 633

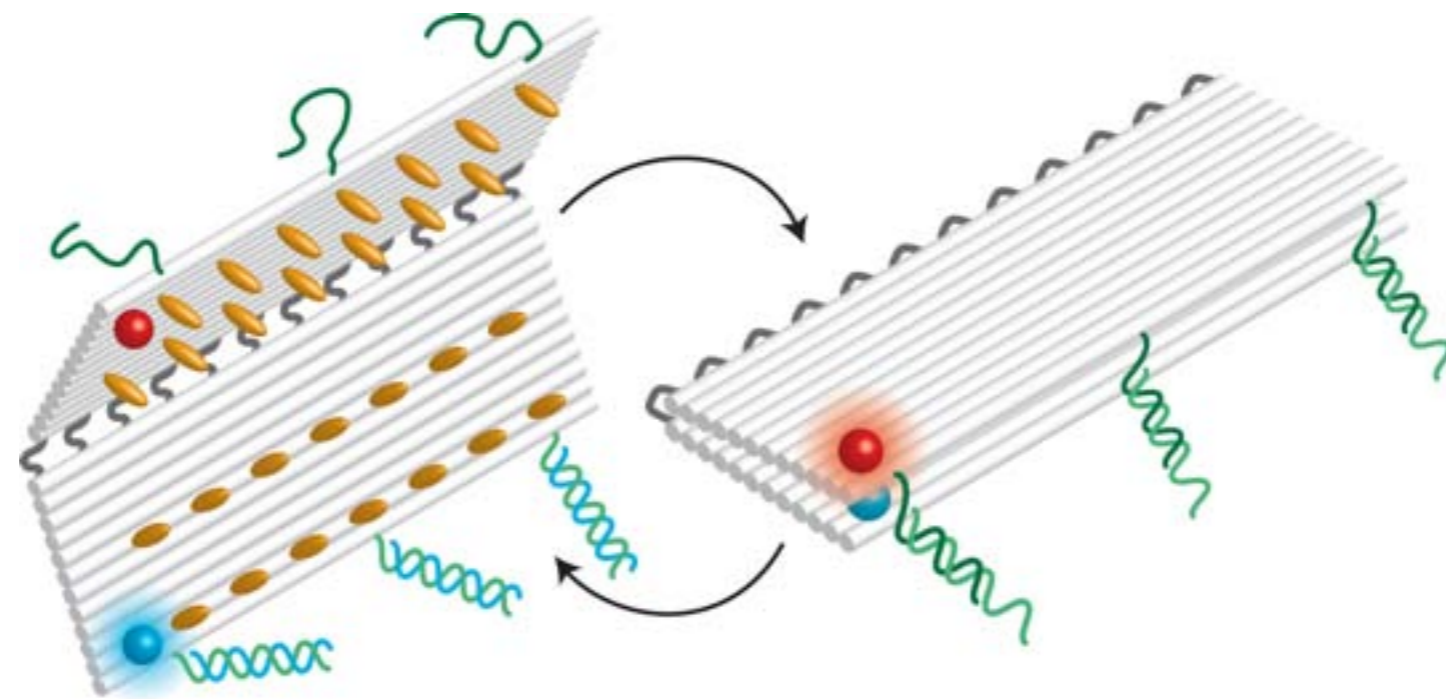


ATTO 514 labeled DNA channels



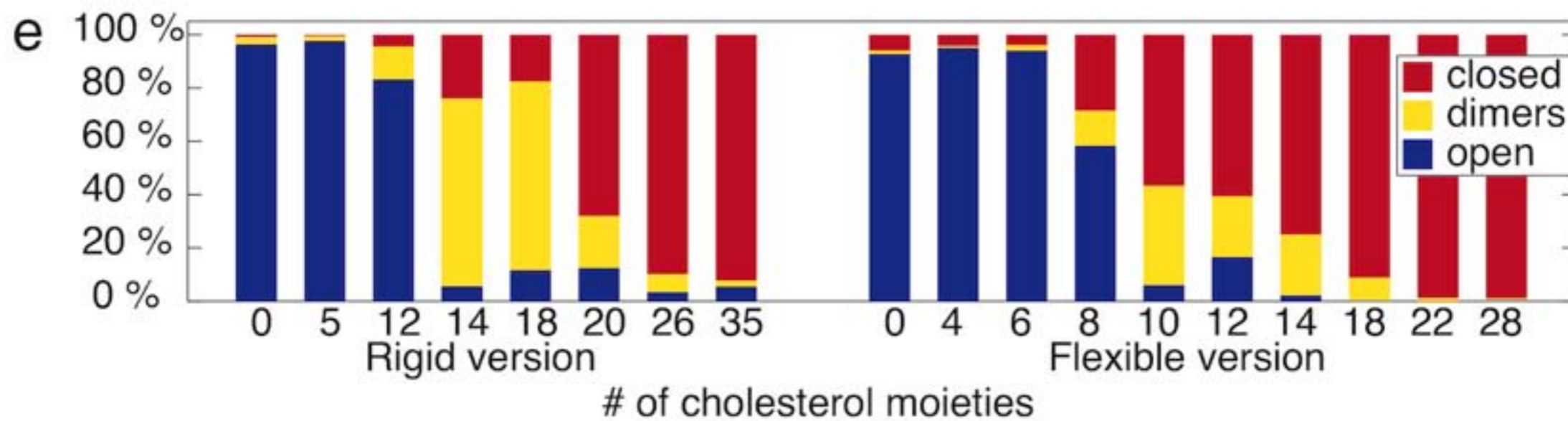
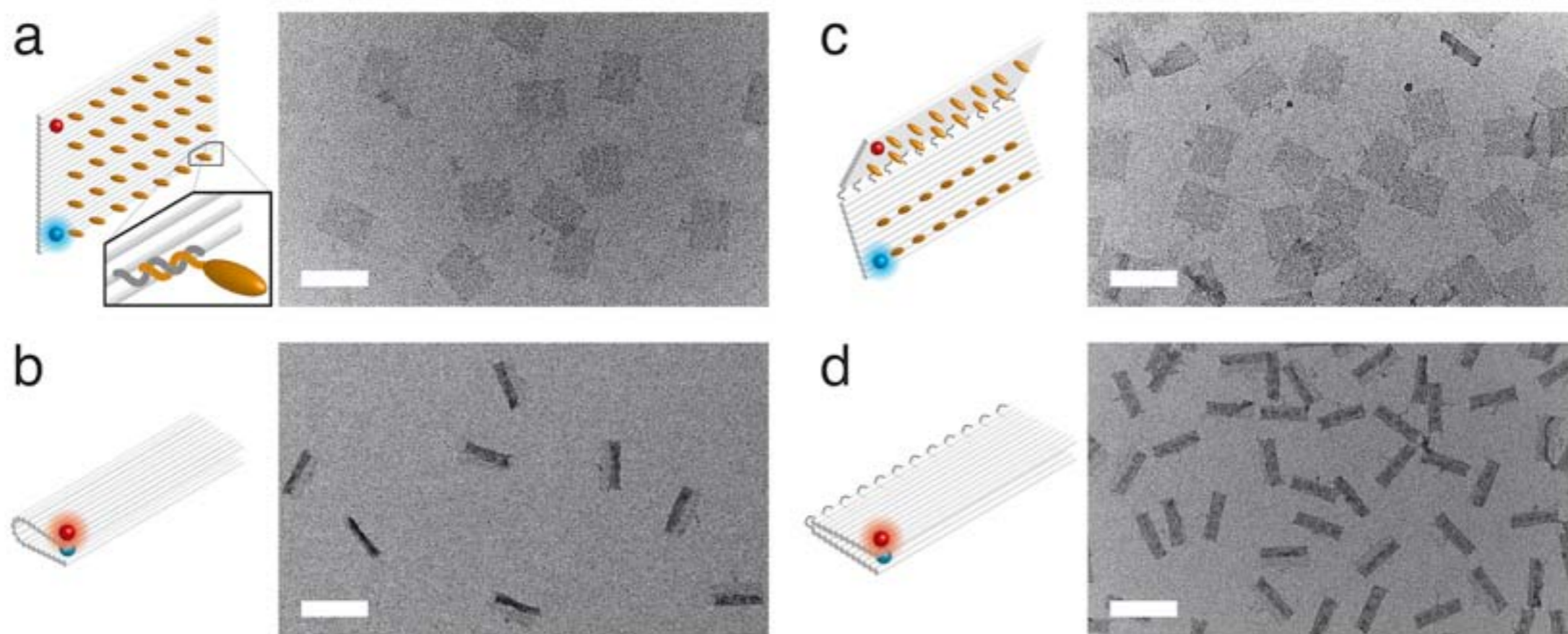
300 mM KCl, 10mM Tris, 1mM EDTA

# A switchable DNA bilayer structure

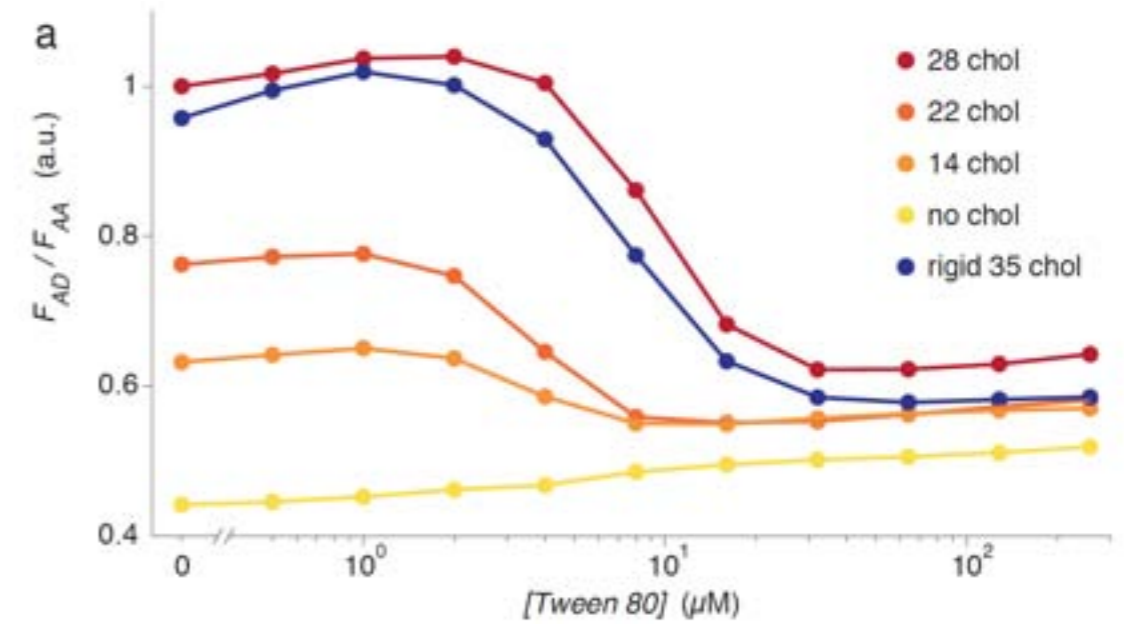
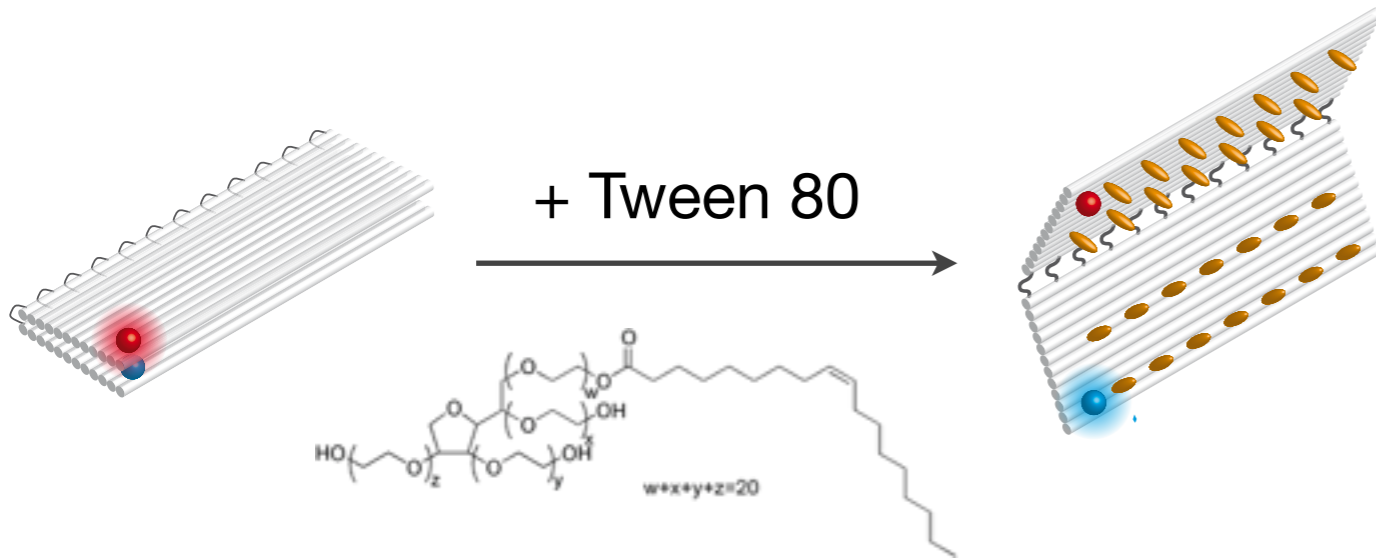


List, J., Weber, M. & Simmel, F. C. Hydrophobic Actuation of a DNA Origami Bilayer Structure. *Angew. Chem. Int. Ed.* **53**, 4236-4239 (2014).

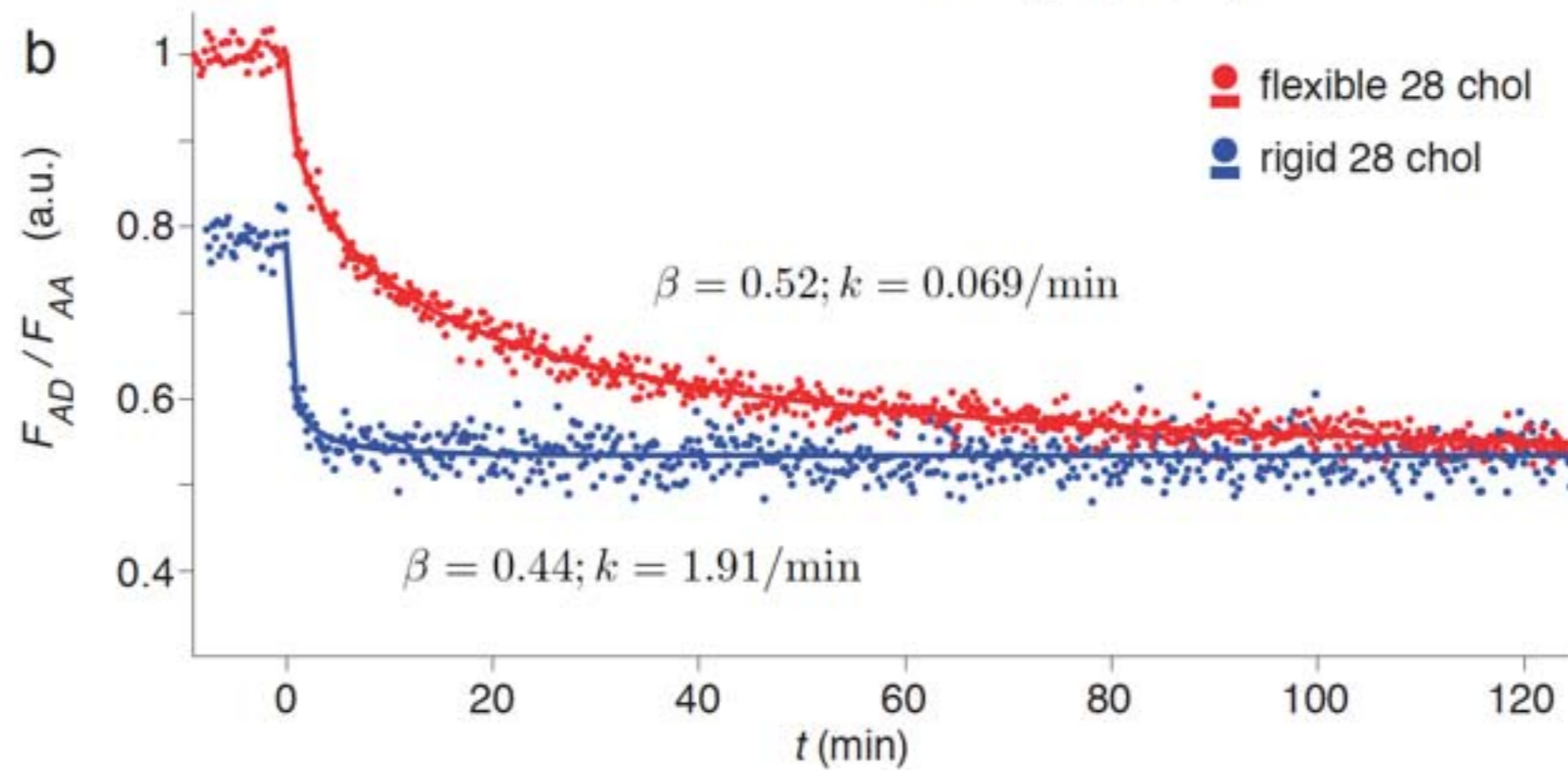
# Hydrophobic interactions bend DNA origami sheets



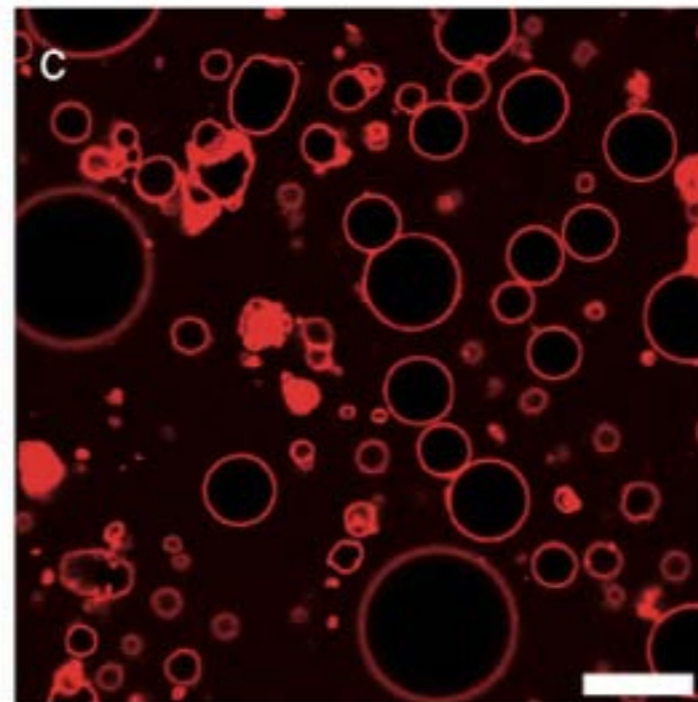
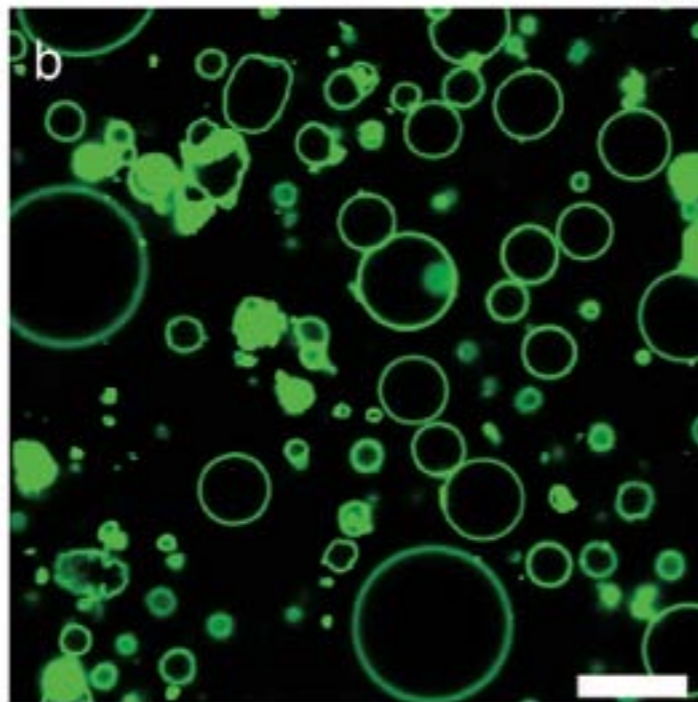
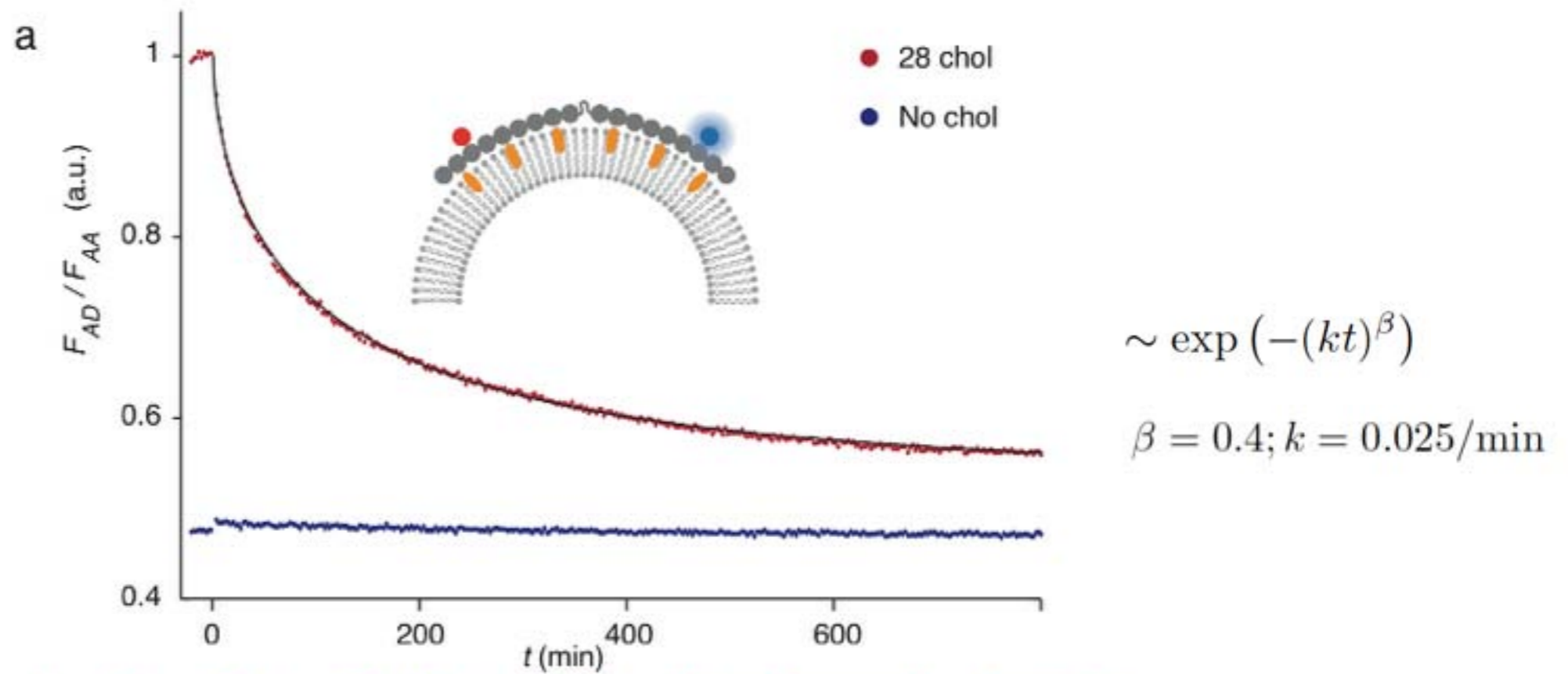
# Opening the DNA bilayers with a surfactant



opening kinetics  $\sim \exp(-(kt)^\beta)$



# Opening in the presence of SUVs and GUVs



# A conditional hydrophobic switch ...

