

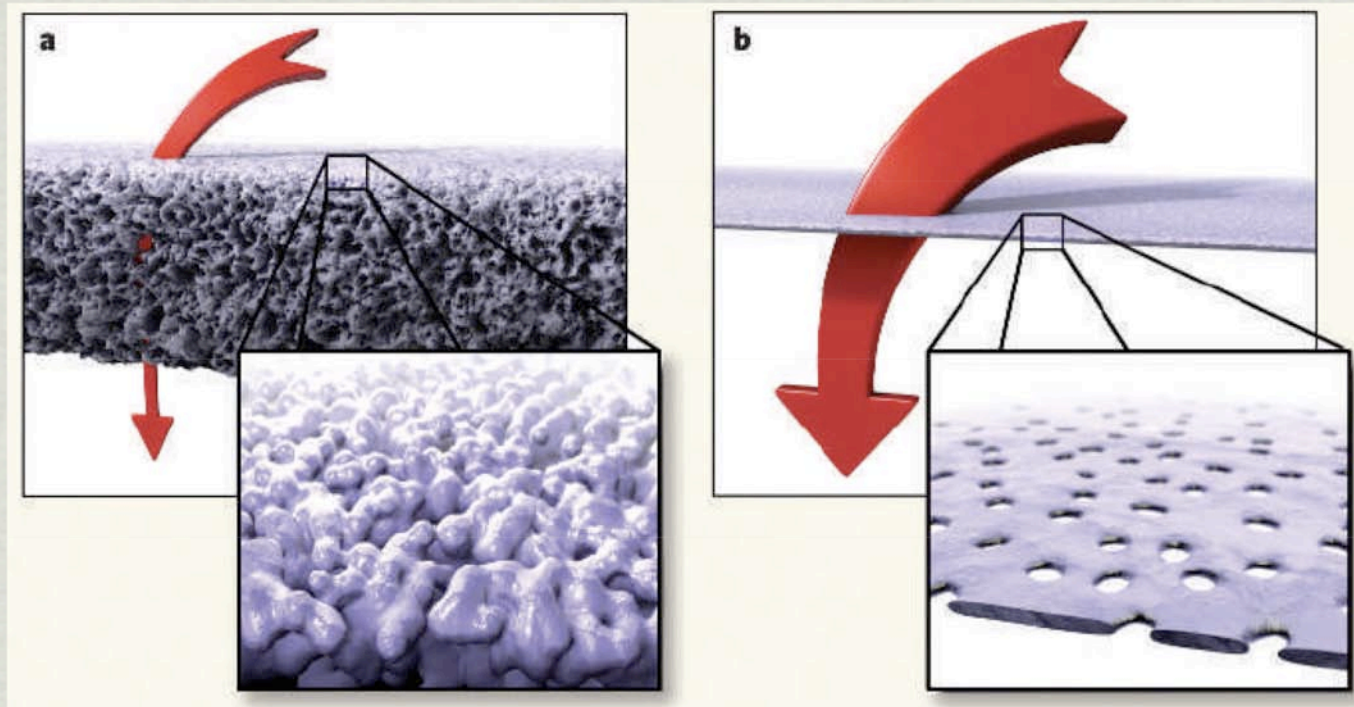
Silicon nanomembranes (pnc-Si): Molecular and Cellular applications

James L. McGrath
Biomedical Engineering
University of Rochester
Nanomembrane Research Group

*EMBS @ SYRACUSE UNIVERSITY
APRIL 23, 2009*

Silicon Nanomembranes

(pnc-Si - porous nanocrystalline silicon)



15nm thick
5-80 nm pores

Thinner = faster transport and less sample loss

A. van den Berg and M. Wessling, Nature 445, 726, 2007

Nanomembrane Research Group

Bioengineering Group

Jim McGrath

Tom Gaborski*

Jess Snyder (*Protein separations*)

Anant Agrawal (*Cellular studies*)

Barrett Nehilla (*Cell applications*)

Henry Chung (*Microfluidics*)

Crowe, Hoffman, Summers (*undergraduates*)

Materials Group

Philippe Fauchet

Chris Striemer*

Dave Fang (*Material development*)

Maryna Kavalenka (*Air permeability*)

SiMPore Inc.*

Rick Richmond

Jamie Roussie

JP Desormeaux

Karl Reisig

Nakul Nataraj

Collaborations

Shigeru Amemiya, PITT

Bill Bernhard, URMC

RIT: SMFL

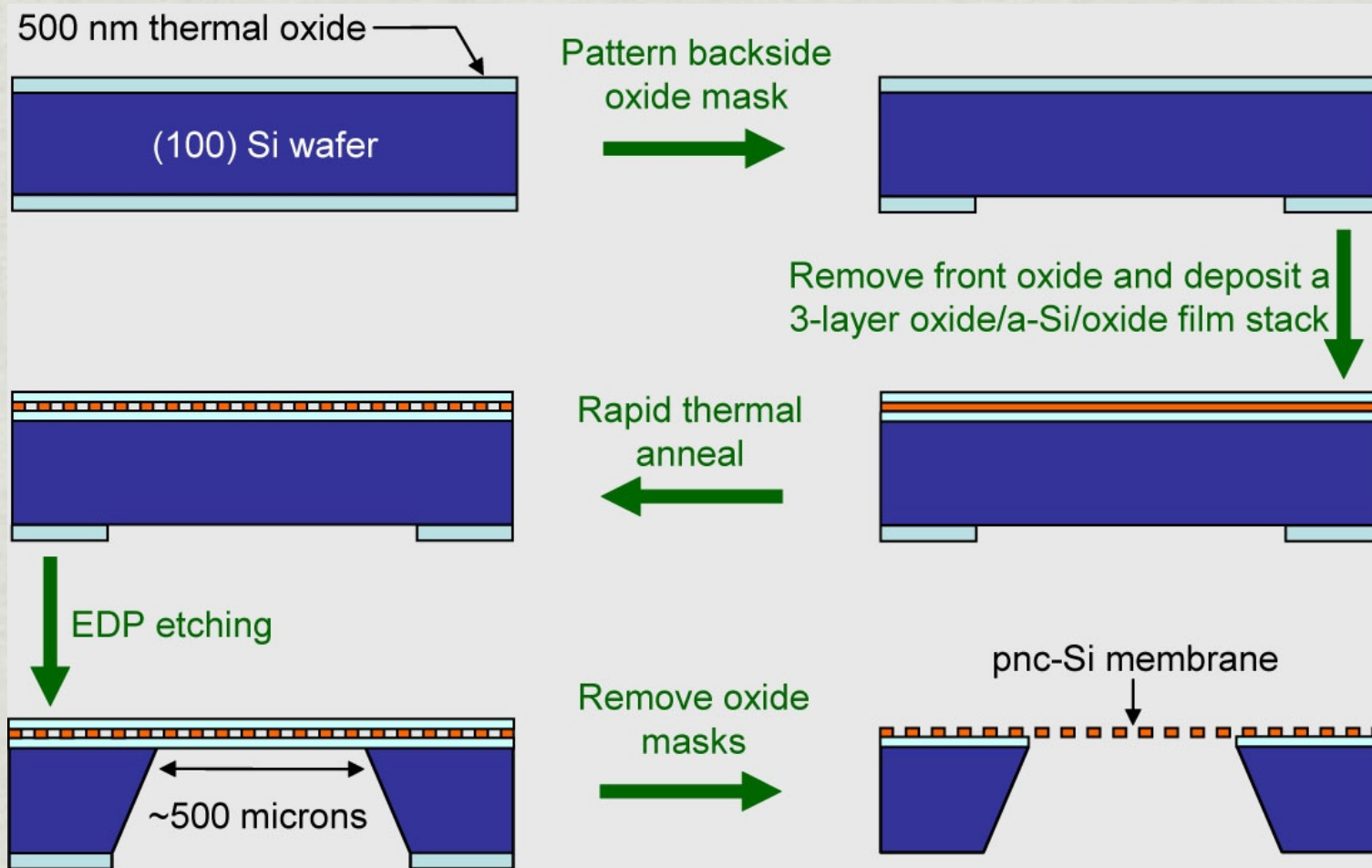
Funding

NSF, NIH, CSTI,
J&J, NYSTAR, CEIS

Outline

- ✱ **Overview of pnc-Si**
- ✱ Transport
- ✱ Separations
- ✱ Cell Culture
- ✱ Other Applications

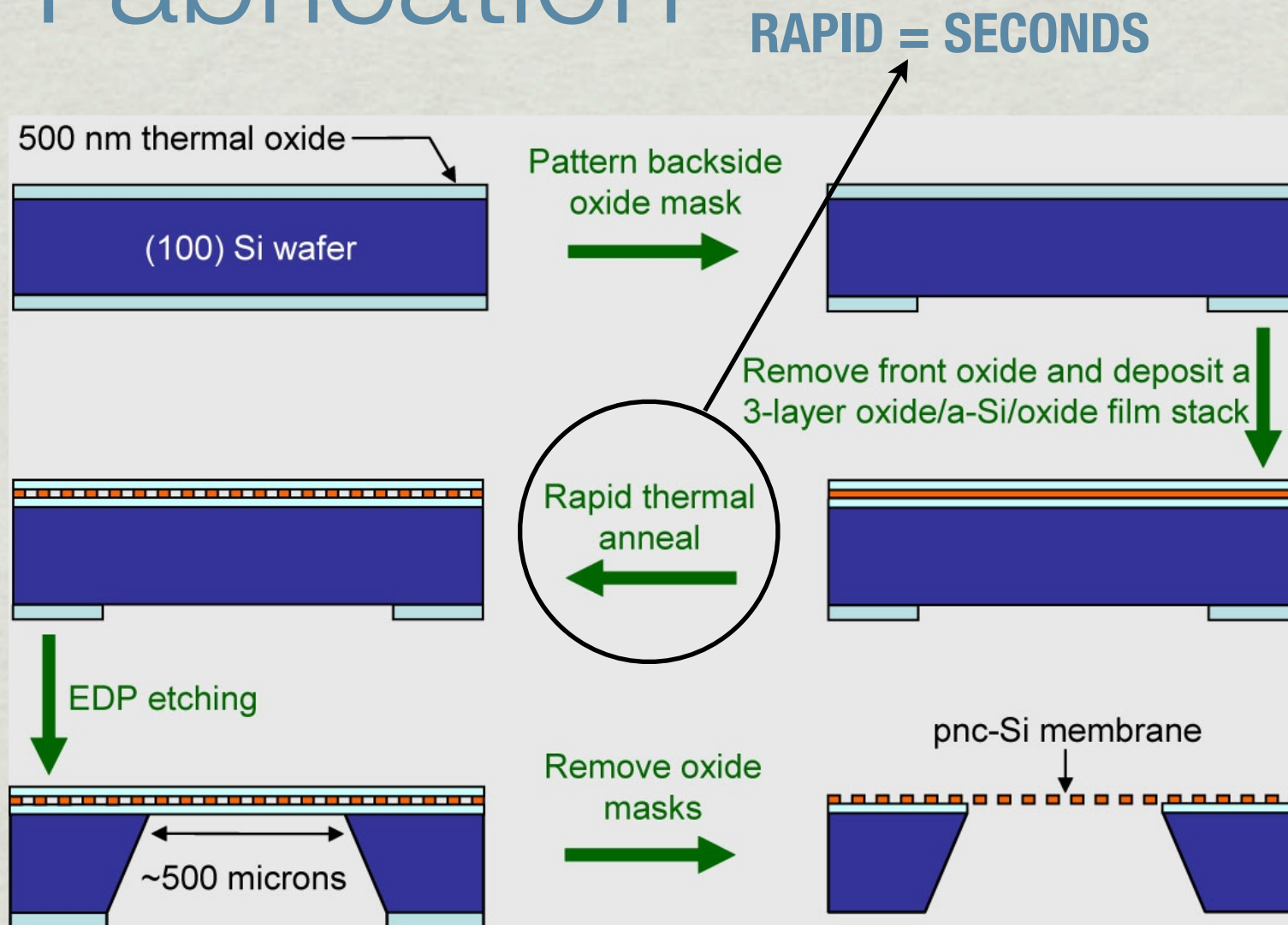
Fabrication



Striemer, C.S, et al. 2007, Nature, 445:749-53



Fabrication

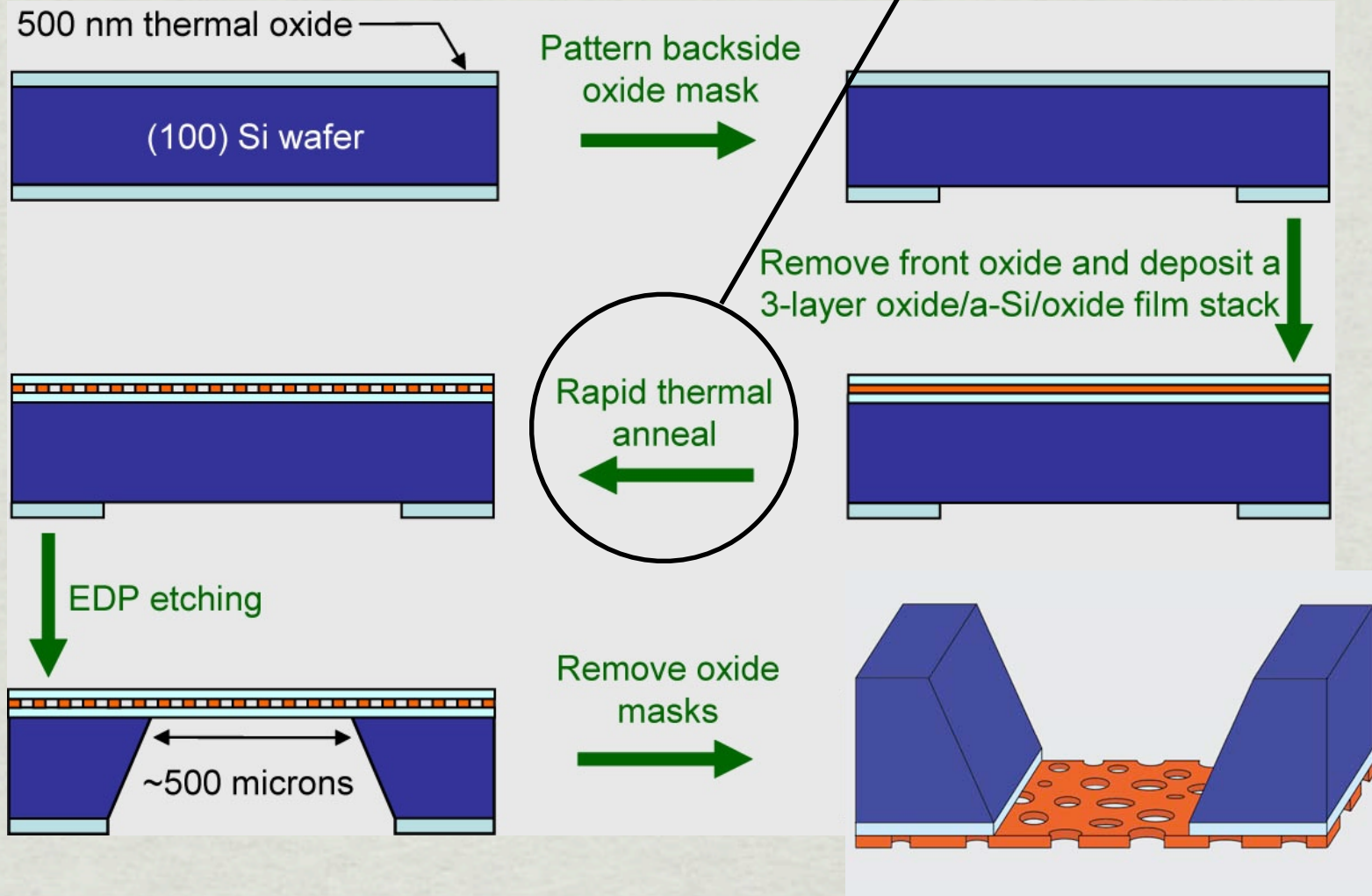


Striemer, C.S, et al. 2007, Nature, 445:749-53



Fabrication

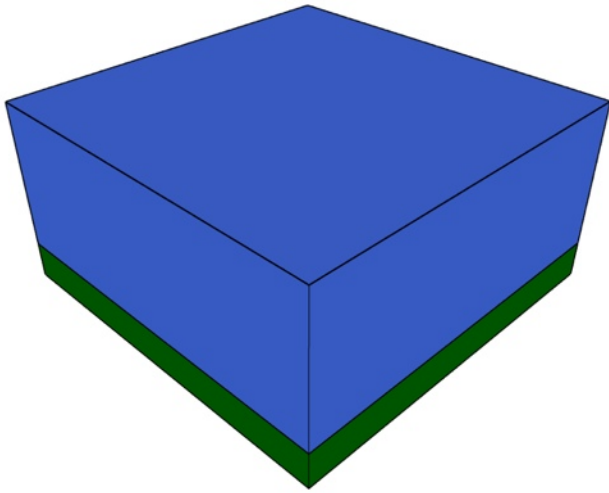
RAPID = SECONDS



Striemer, C.S, et al. 2007, Nature, 445:749-53

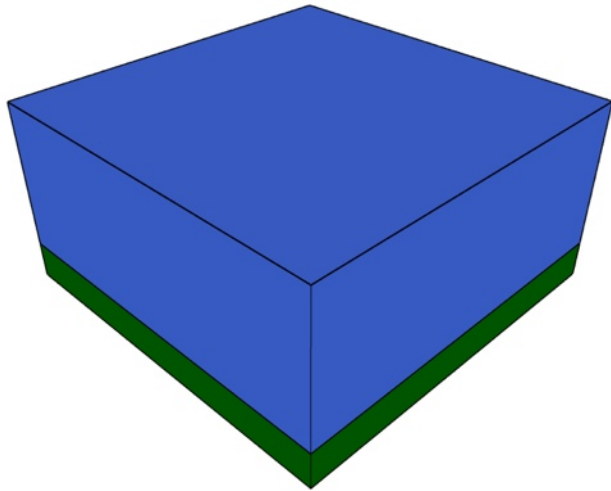
physical vapor deposition

frontside

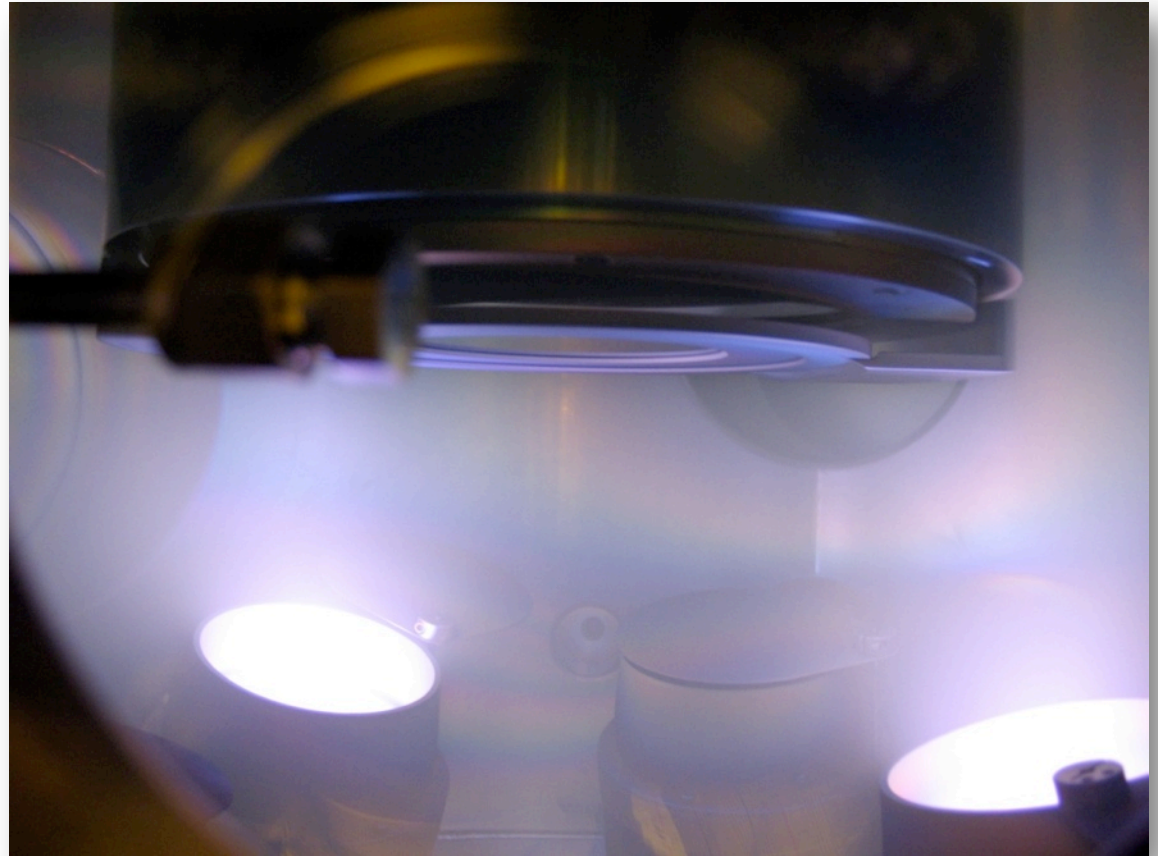


physical vapor deposition

frontside

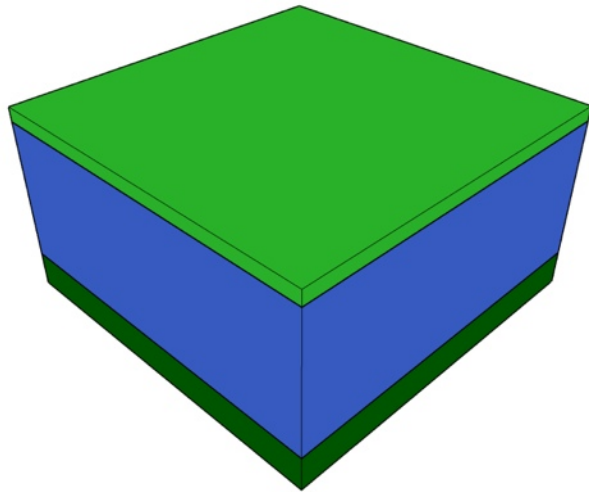


rf magnetron sputtering



physical vapor deposition

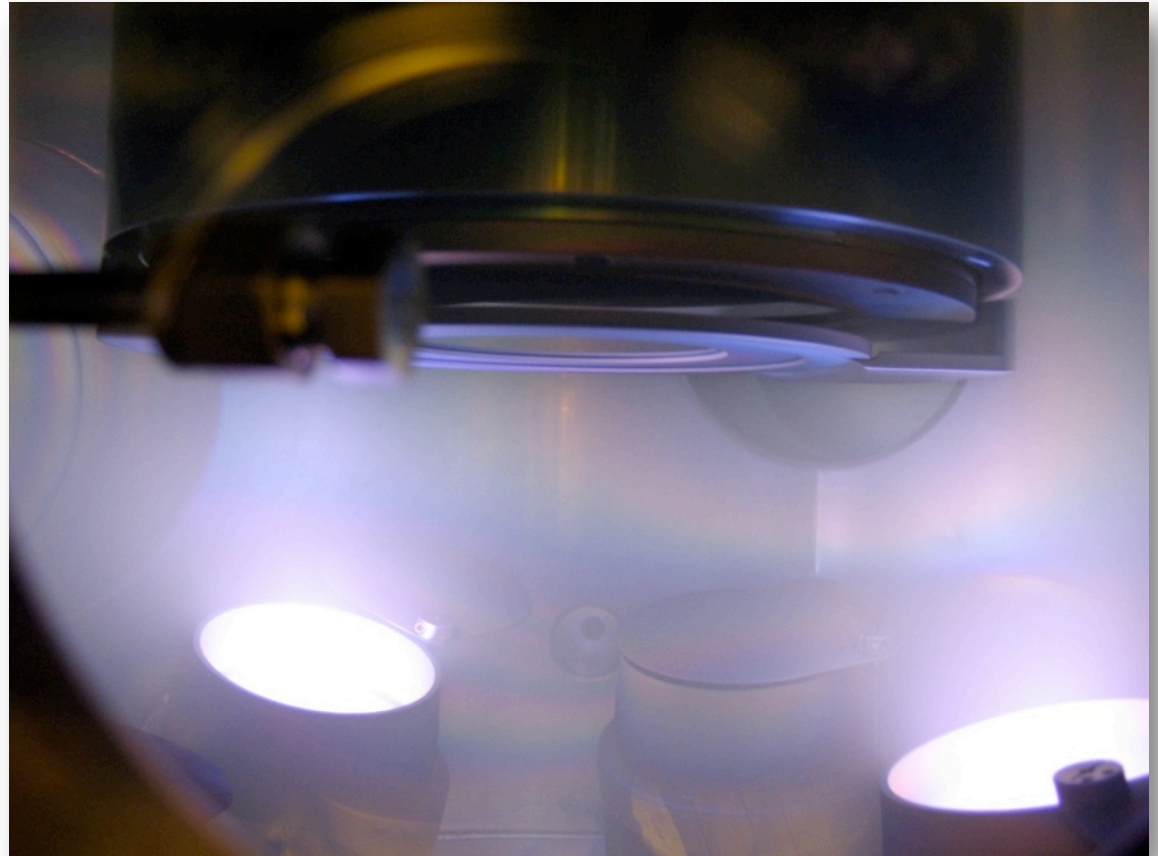
frontside



 20 nm sputtered SiO_2

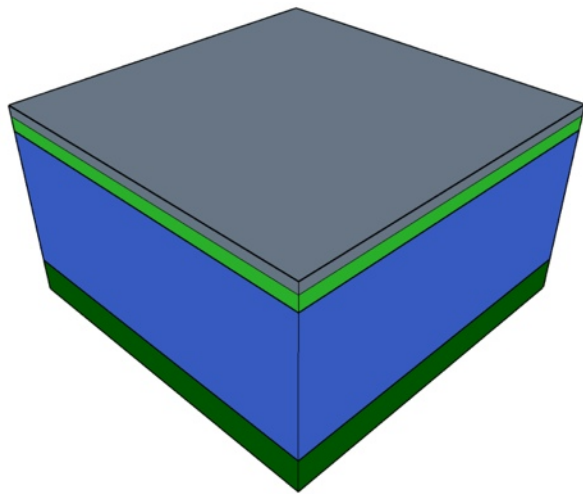
 15 nm $\alpha\text{-Si}$

rf magnetron sputtering



physical vapor deposition

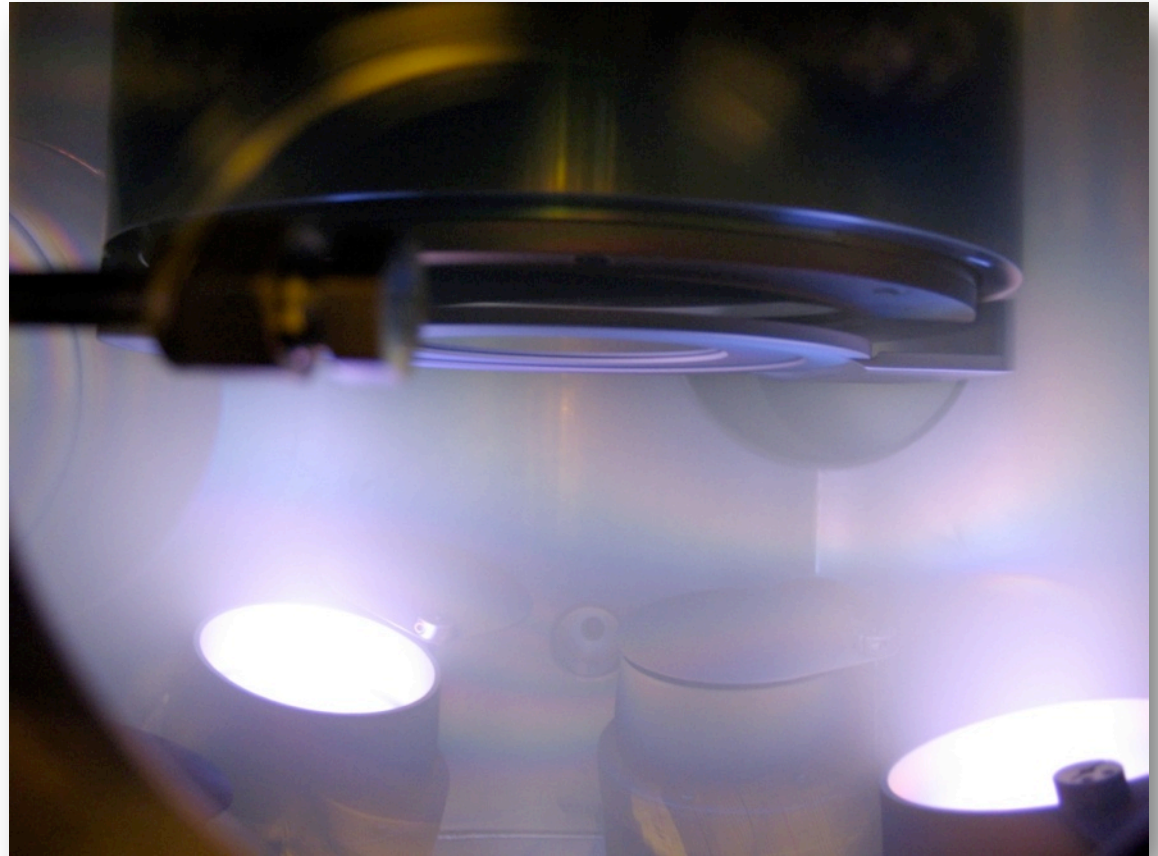
frontside



 20 nm sputtered SiO_2

 15 nm $\alpha\text{-Si}$

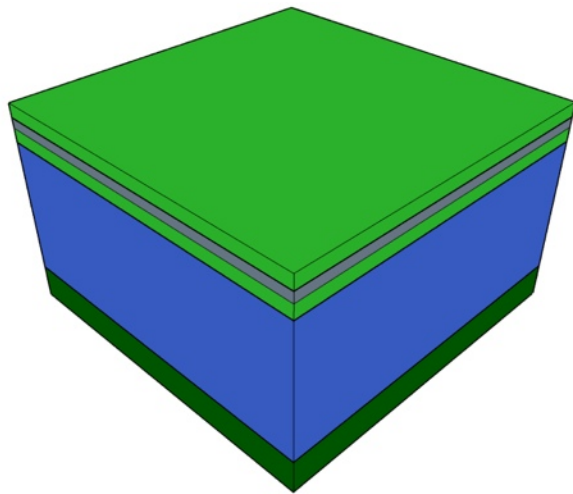
rf magnetron sputtering



physical vapor deposition

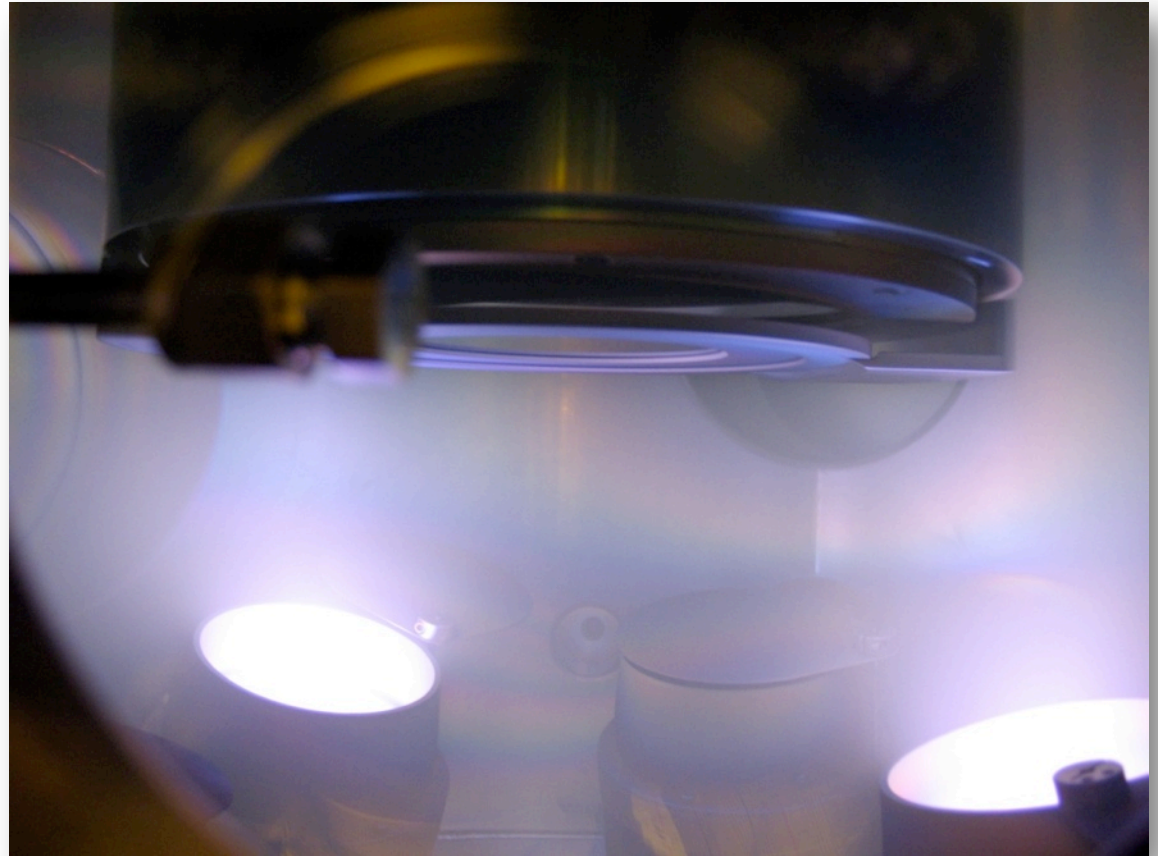
frontside

rf magnetron sputtering



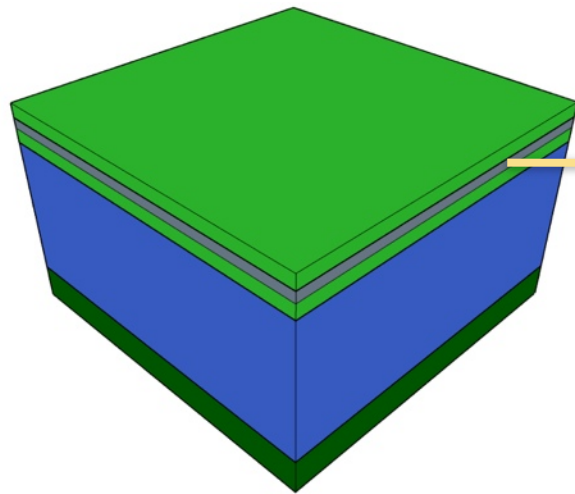
 20 nm sputtered SiO_2

 15 nm $\alpha\text{-Si}$

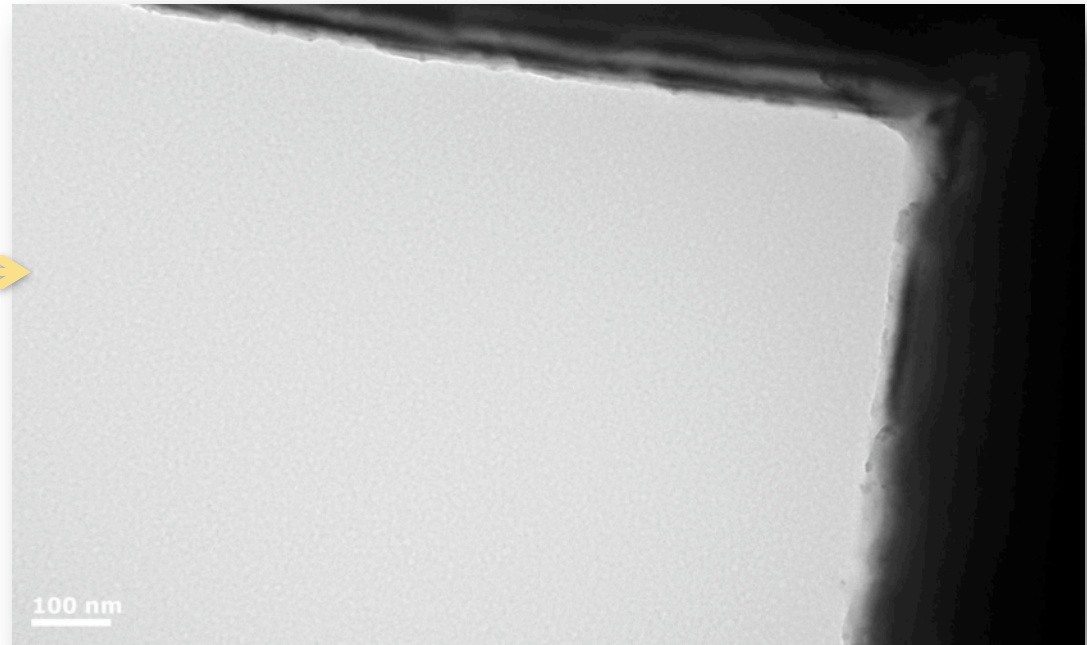


physical vapor deposition


frontside



amorphous Si

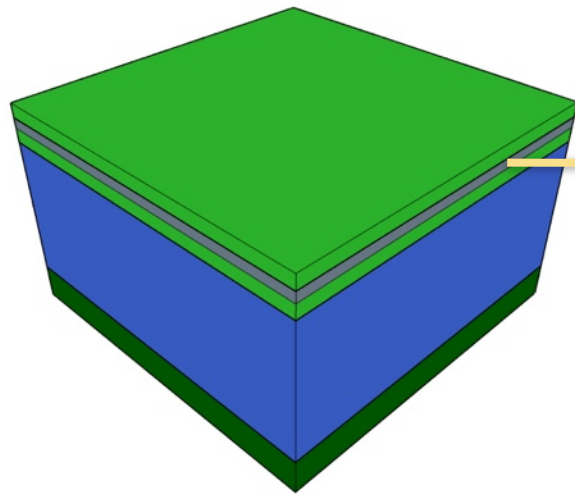


 20 nm sputtered SiO₂

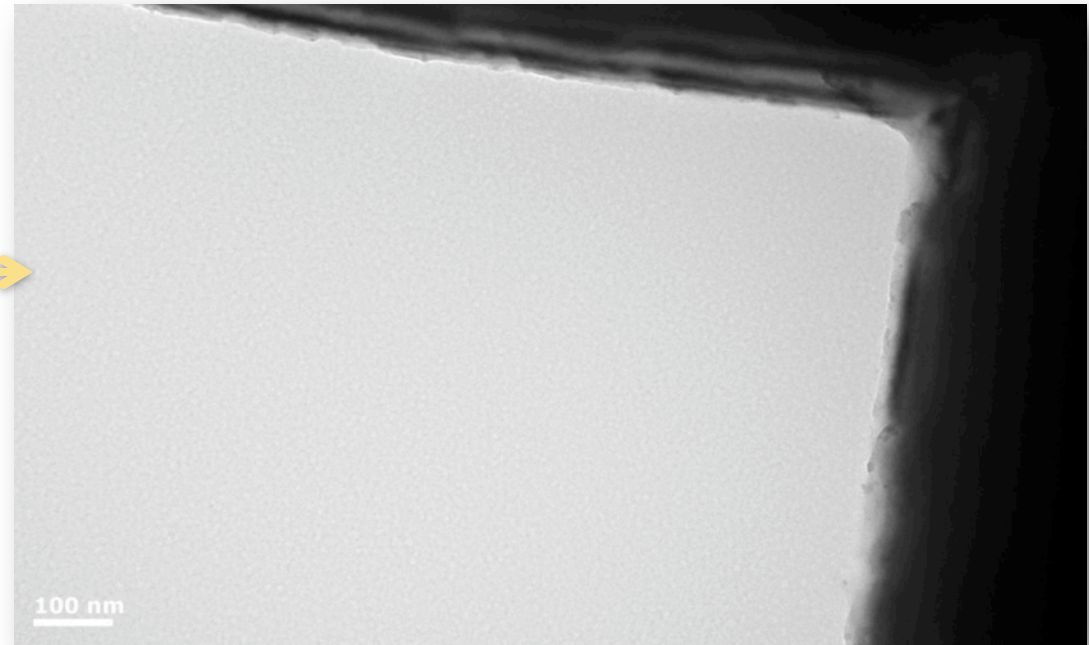
 15 nm α -Si

rapid thermal anneal

frontside



amorphous Si



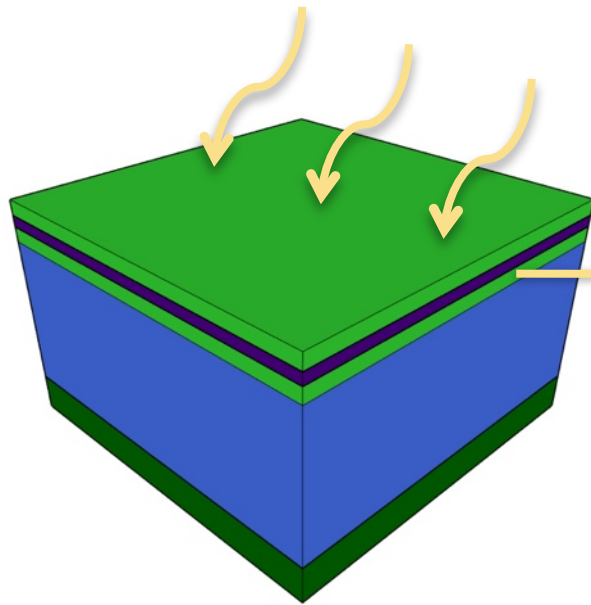
 20 nm sputtered SiO₂

 15 nm α -Si

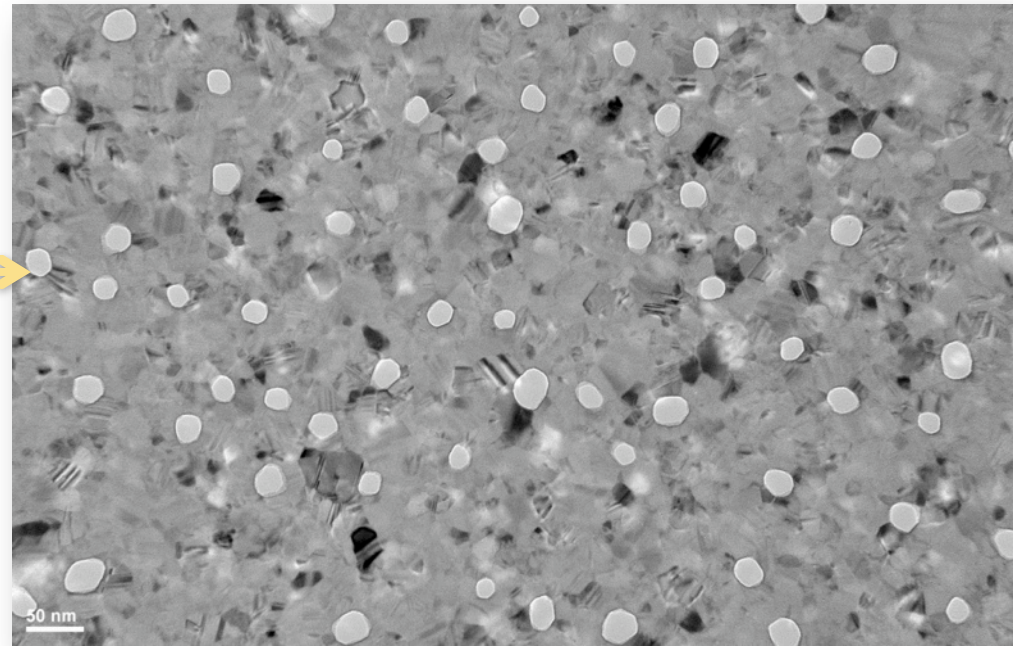
rapid thermal anneal

frontside

700 °C – 1000 °C < 5 min



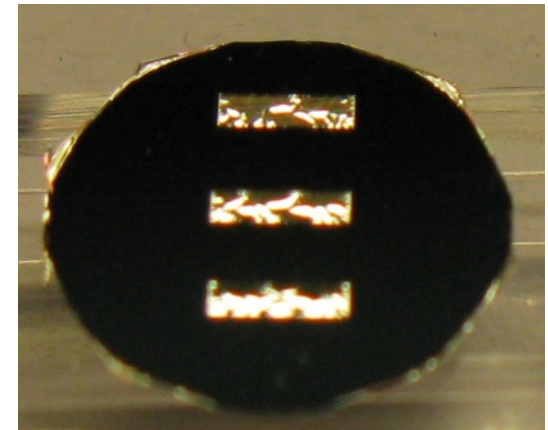
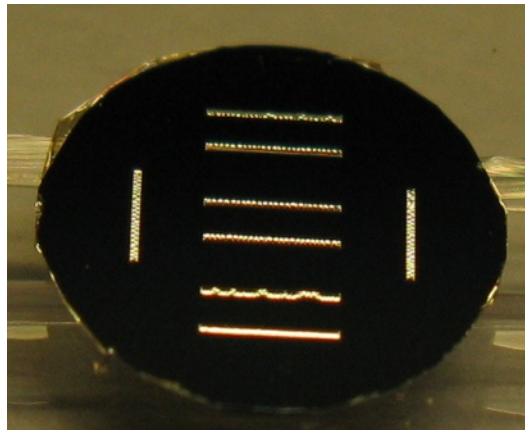
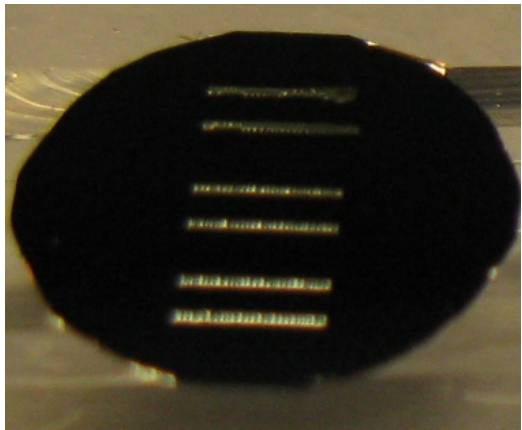
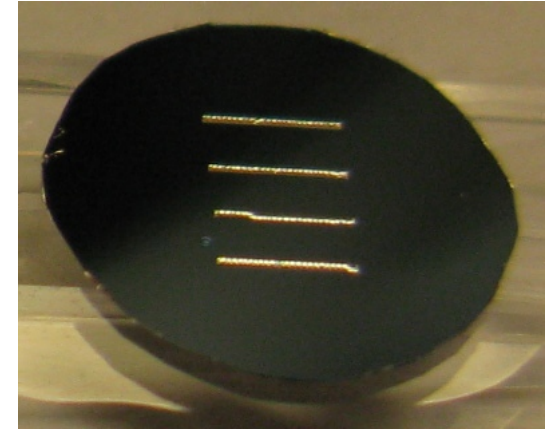
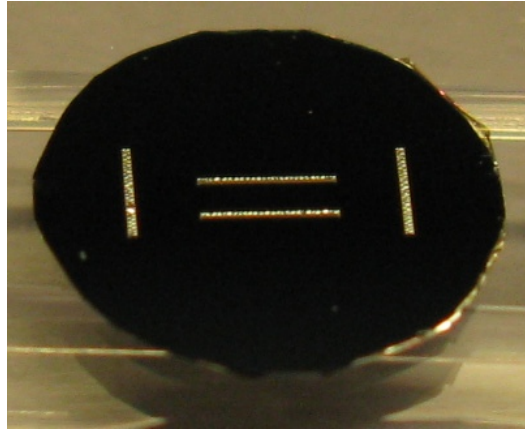
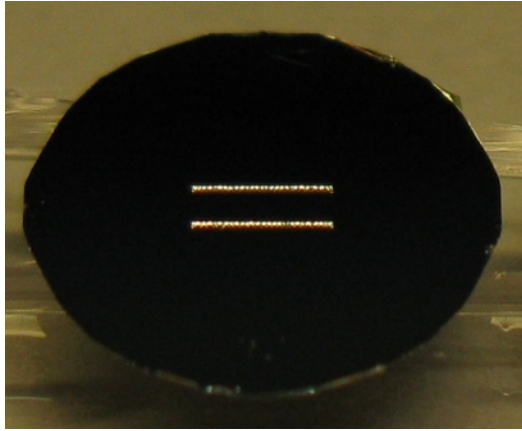
pnc-Si



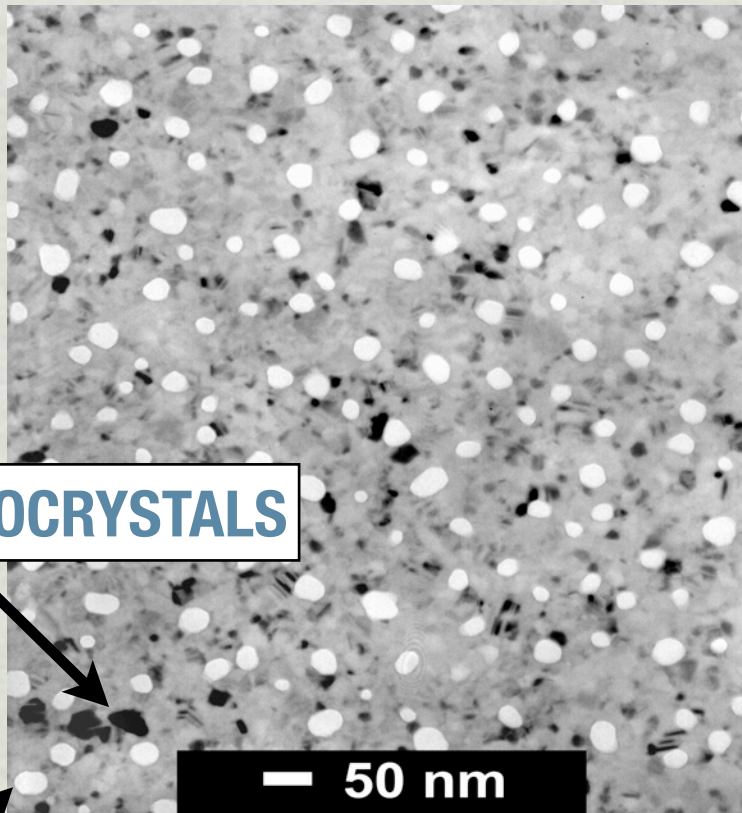
20 nm sputtered SiO₂

15 nm pnc-Si

variety of formats



Sharp Cut-offs and Tunable Pore Sizes

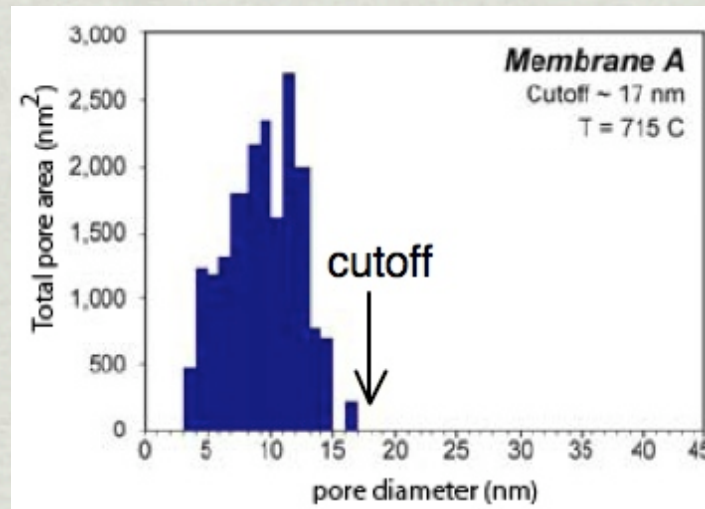


NANOCRYSTALS

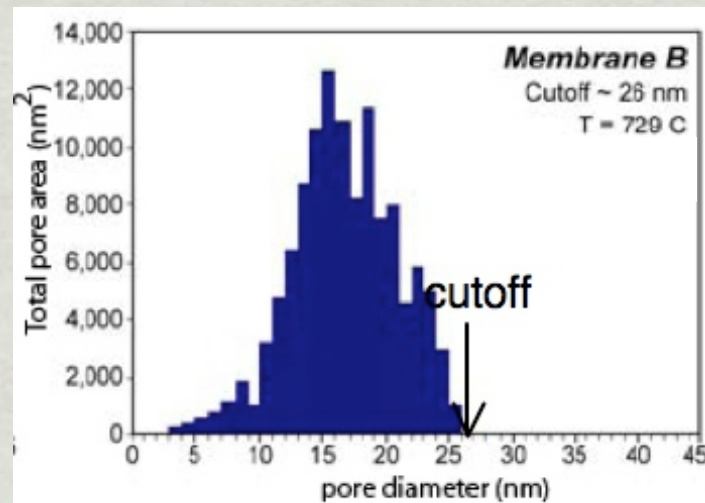
NANOPORES

— 50 nm

Striemer, C.S, et al. 2007, Nature, 445:749-53



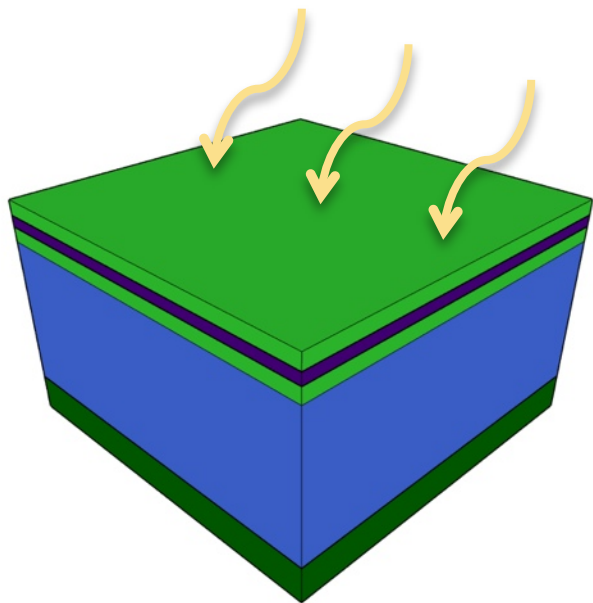
T = 715C



T = 729C

NRG 2009

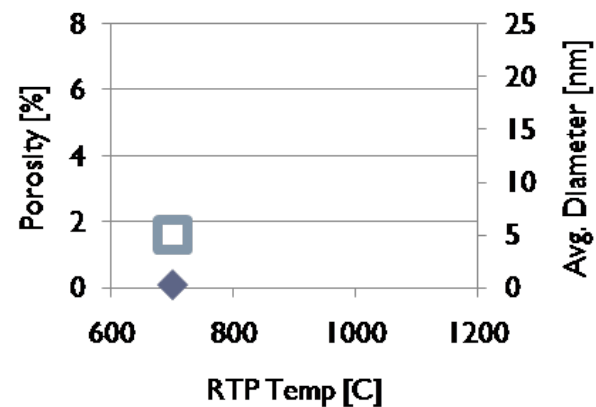
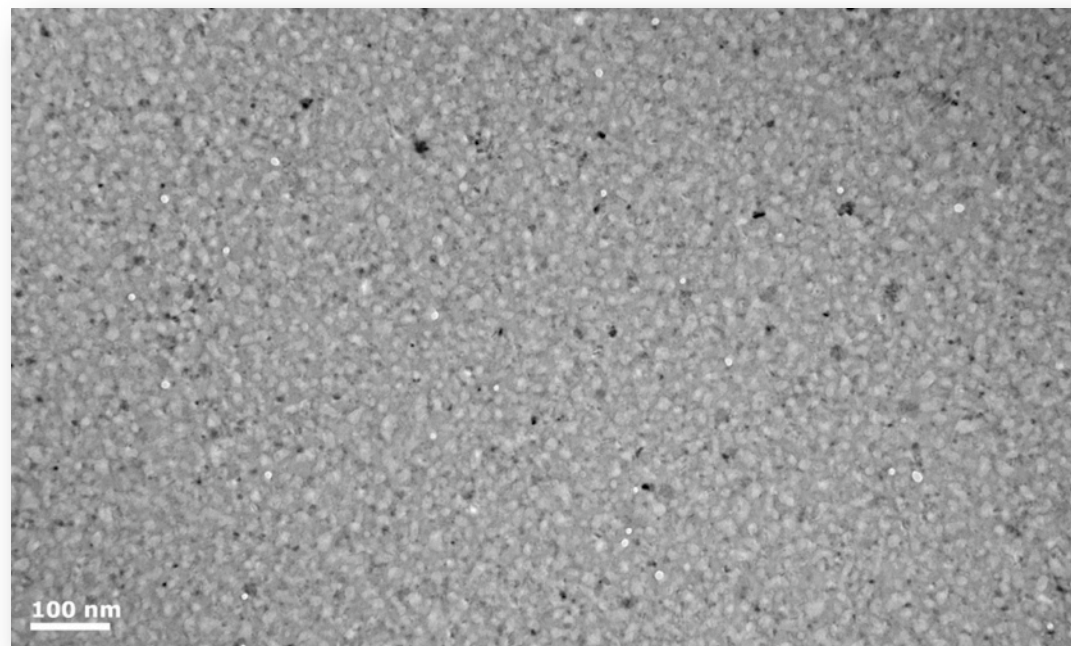
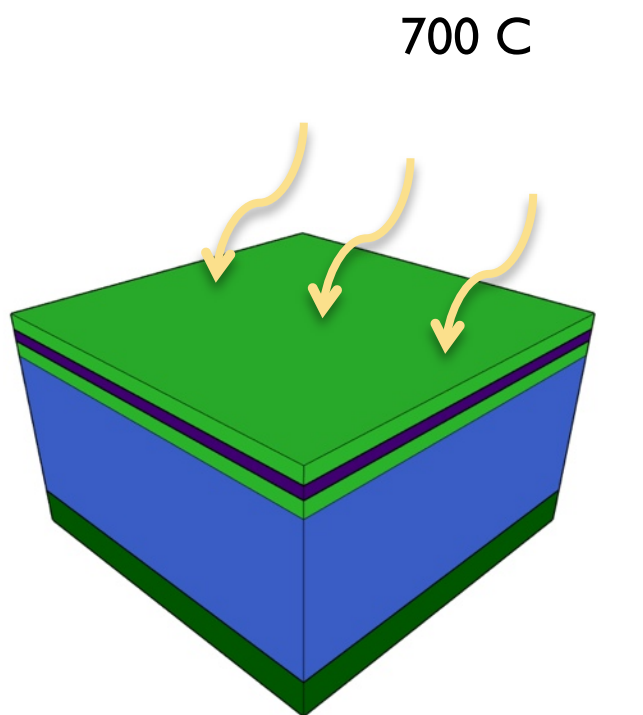
pore size & density – rtp temperature



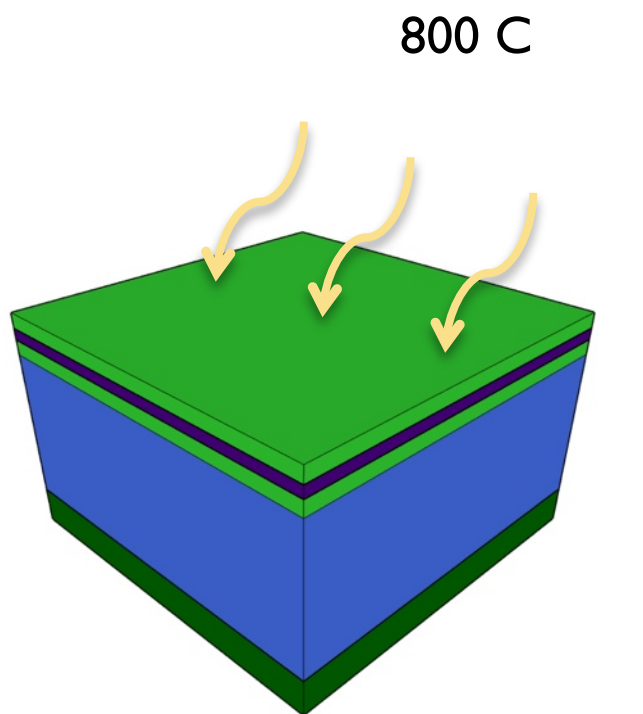
 20 nm sputtered SiO₂

 15 nm pnc-Si

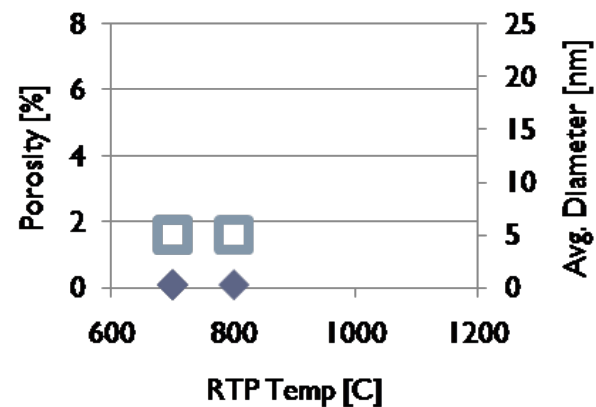
pore size & density – rtp temperature



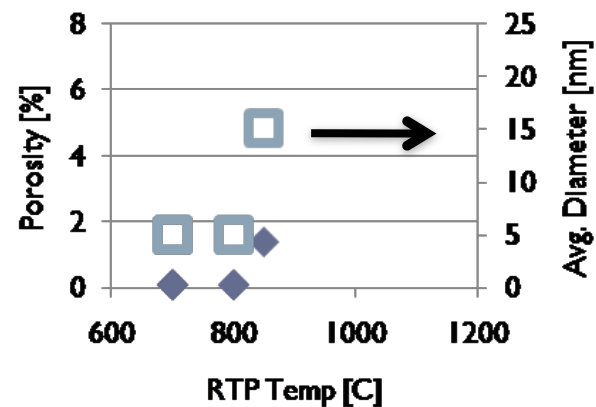
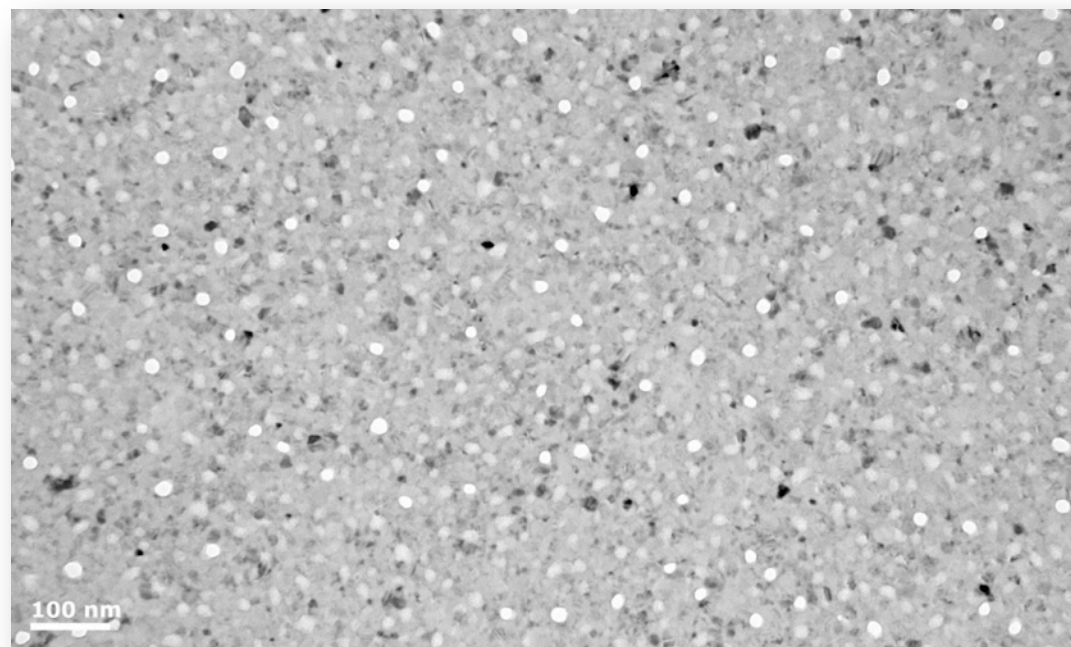
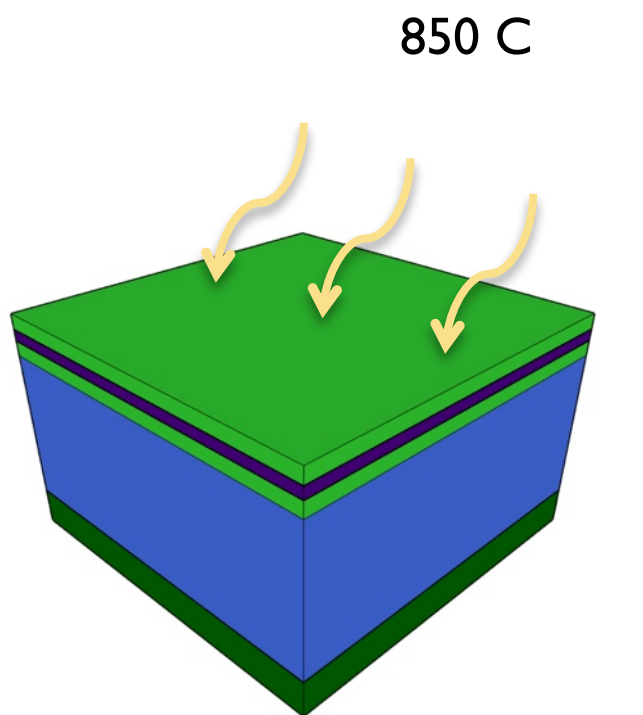
pore size & density – rtp temperature



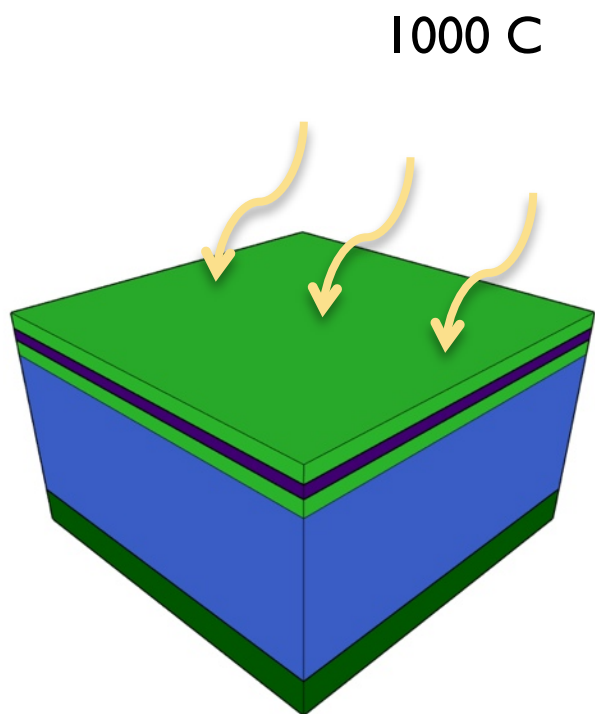
- 20 nm sputtered SiO₂
- 15 nm pnc-Si



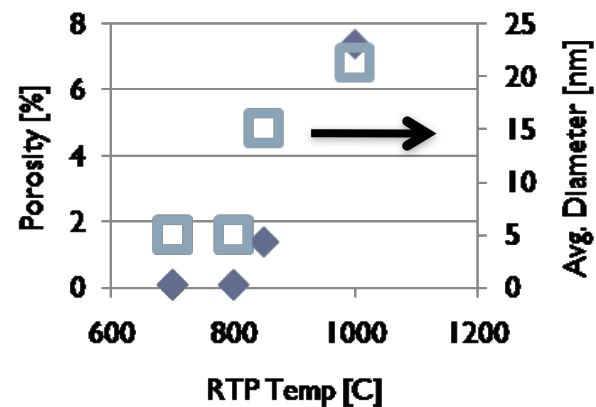
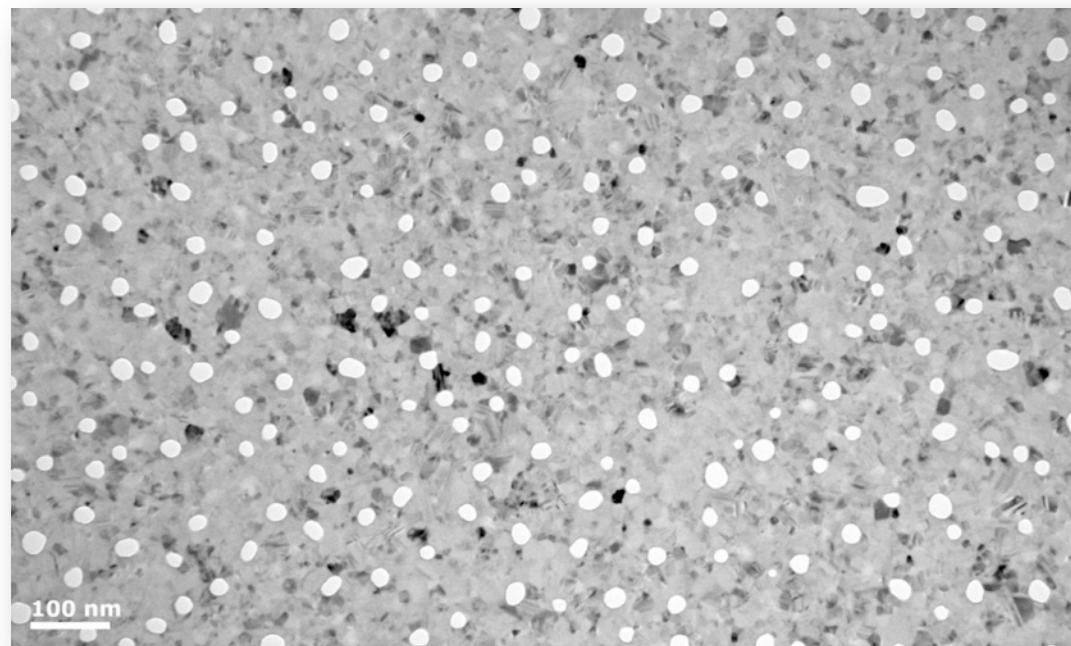
pore size & density – rtp temperature



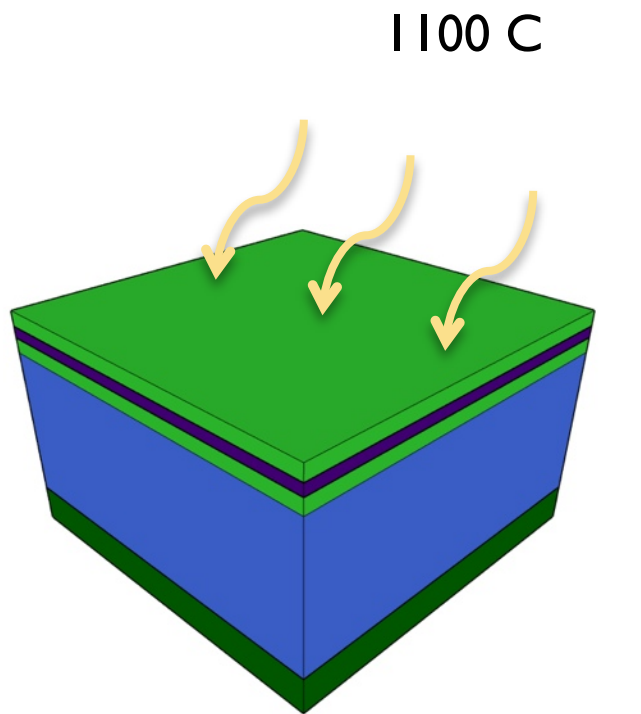
pore size & density – rtp temperature



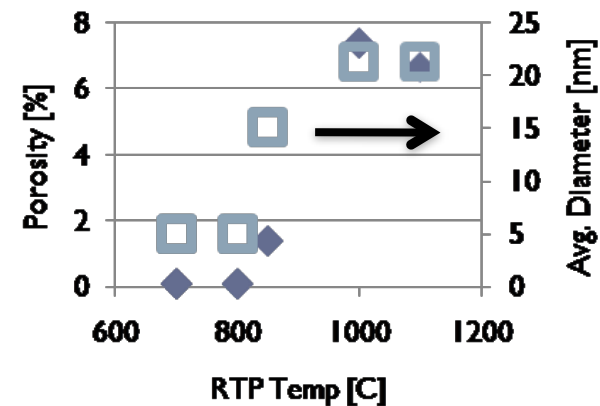
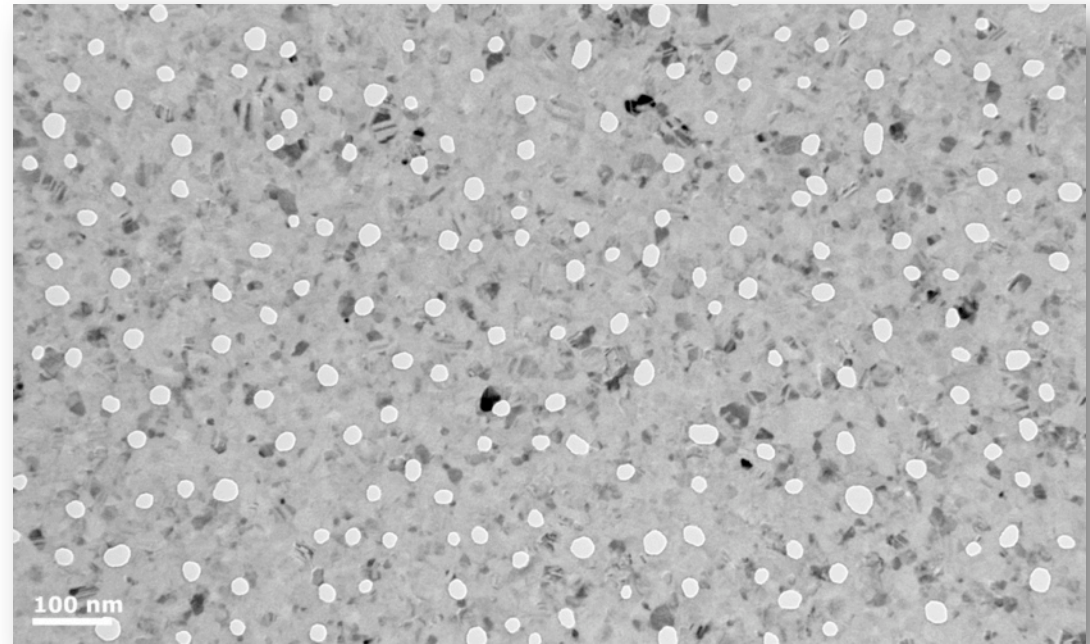
- 20 nm sputtered SiO₂
- 15 nm pnc-Si



pore size & density – rtp temperature

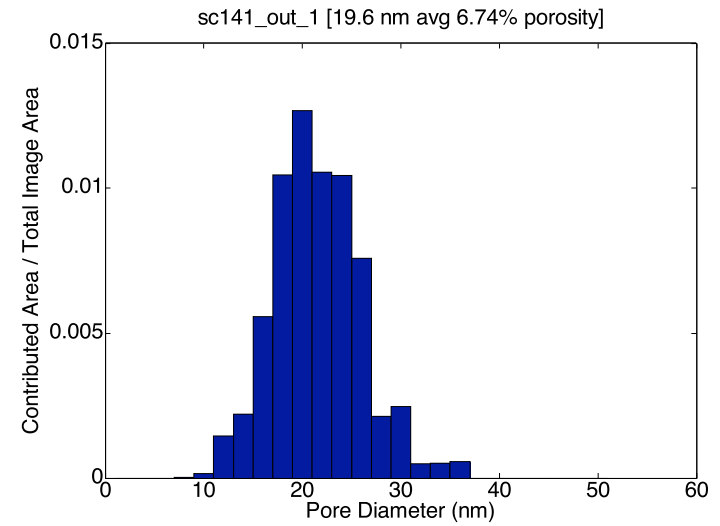
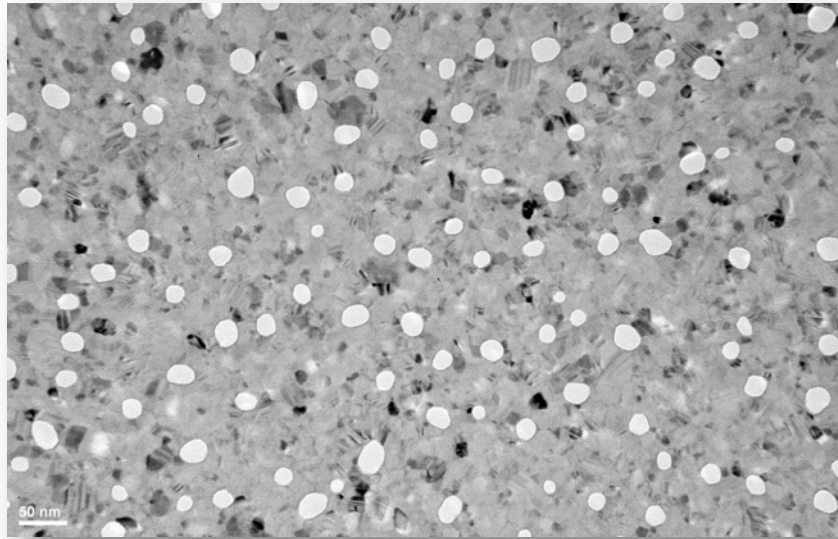


- 20 nm sputtered SiO₂
- 15 nm pnc-Si



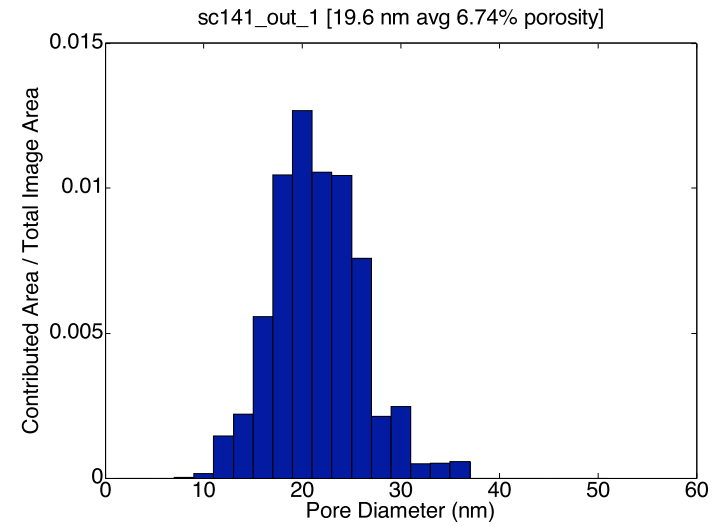
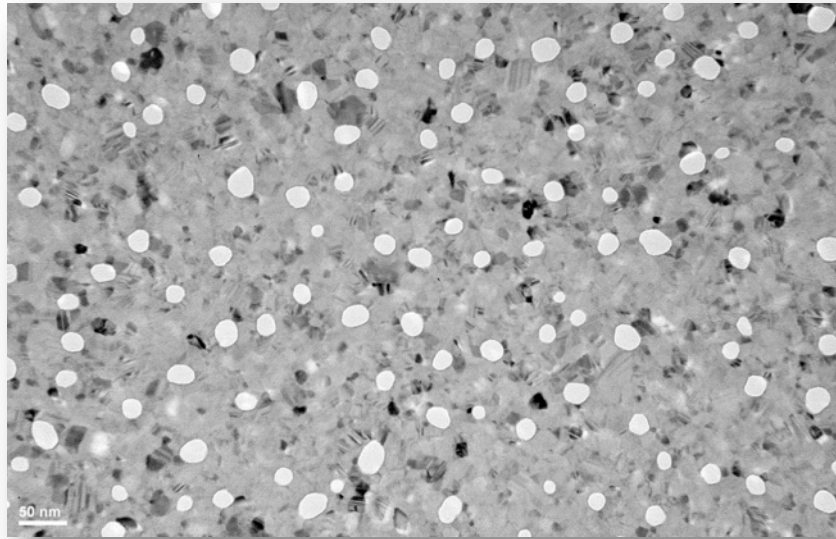
controlling morphology – thickness

15 nm pnc-Si

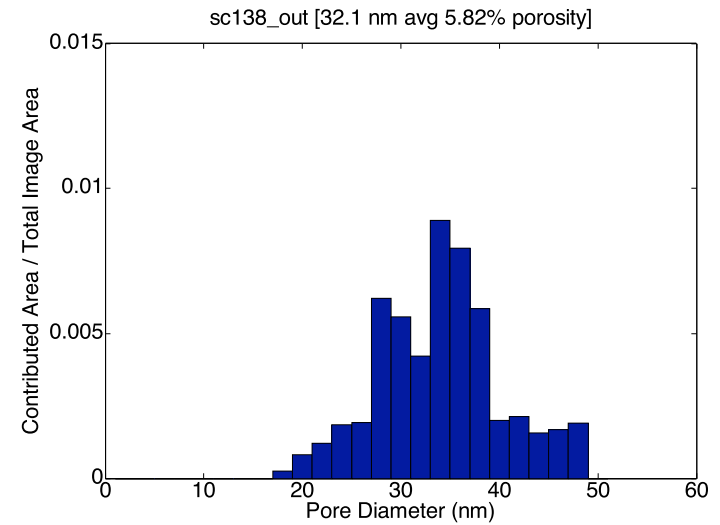
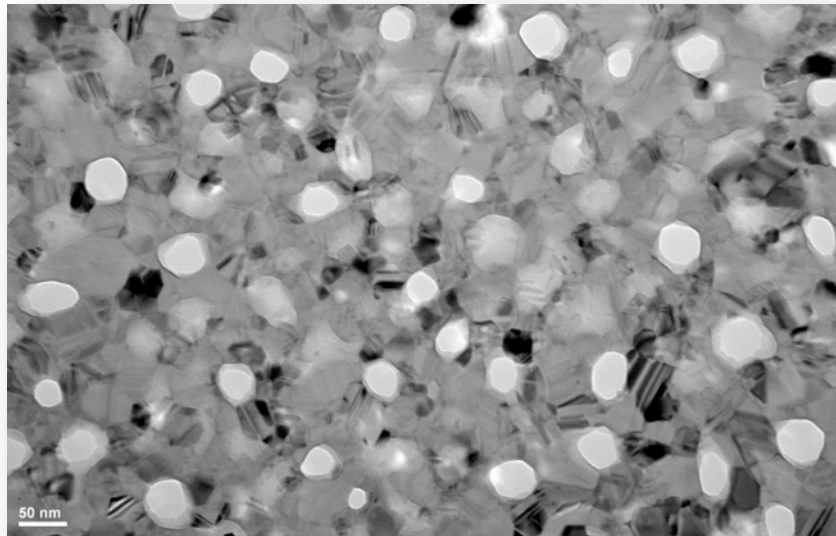


controlling morphology – thickness

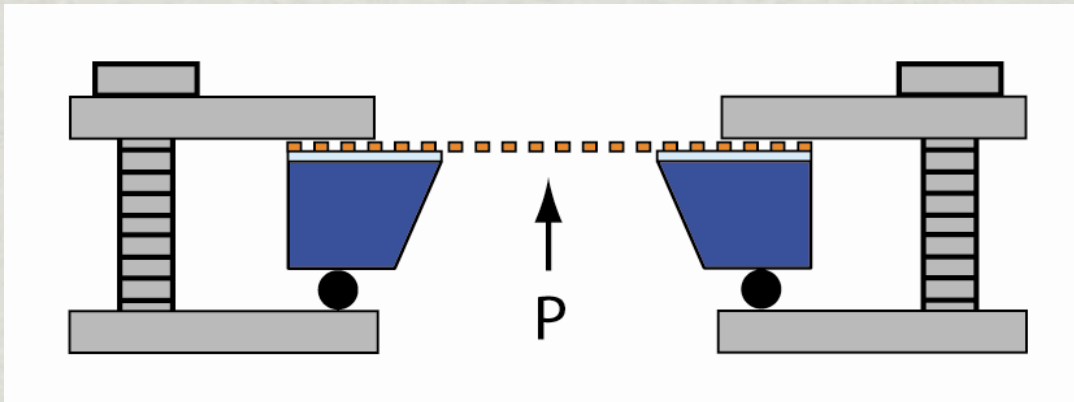
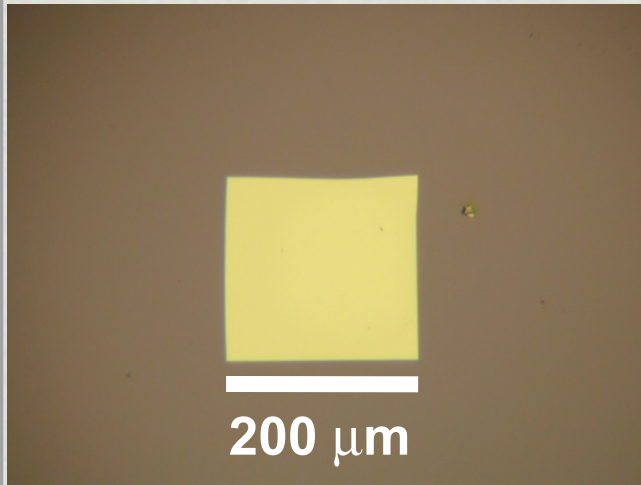
15 nm pnc-Si



30 nm pnc-Si



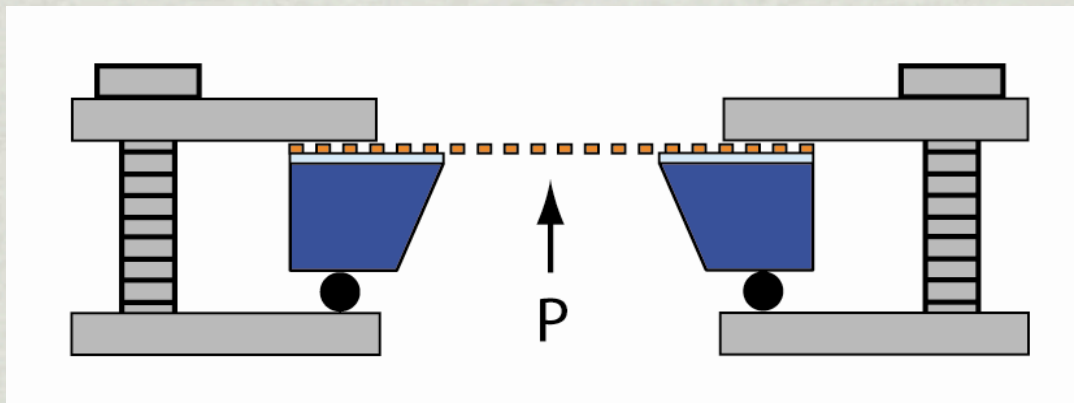
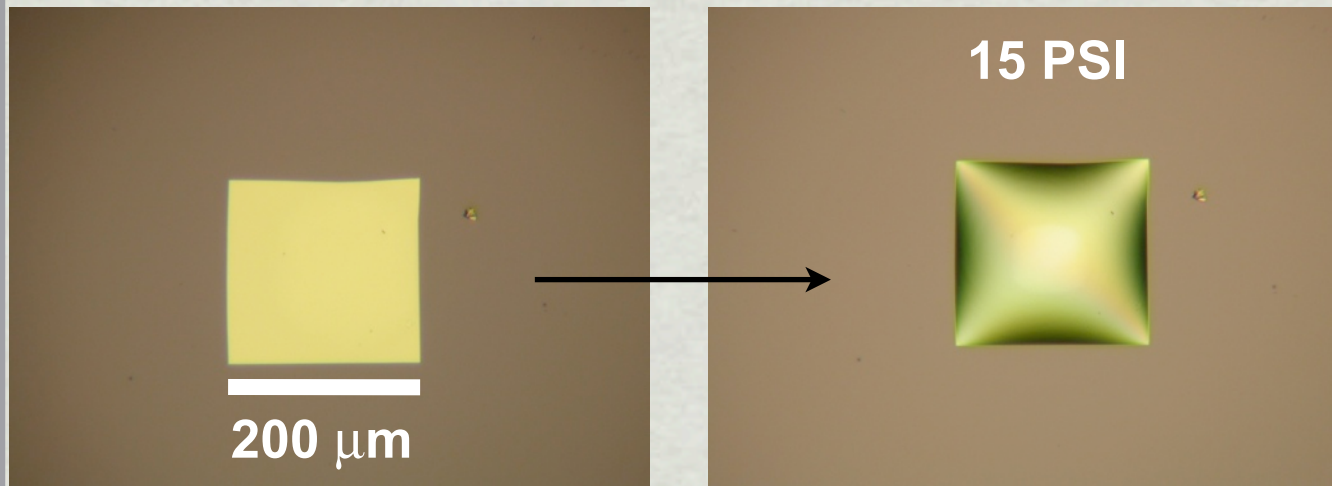
Mechanically Robust



Striener, C.S, et al. 2007, Nature, 445:749-53

 NRG 2009

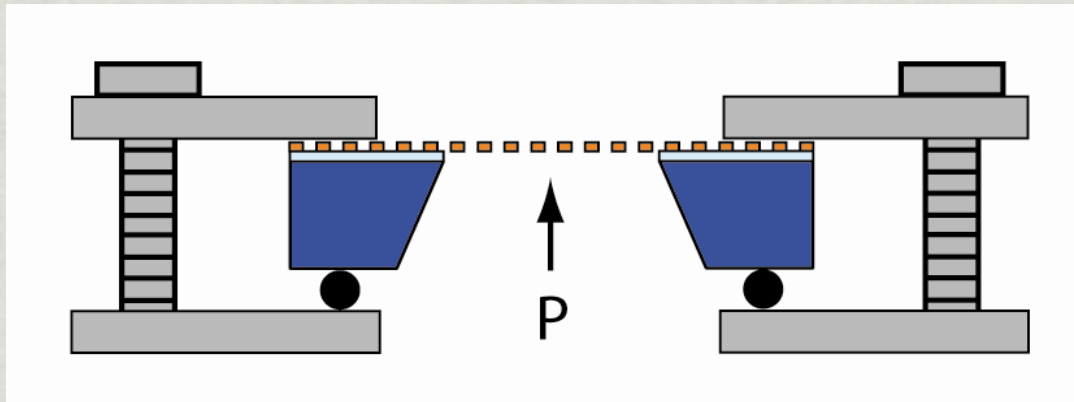
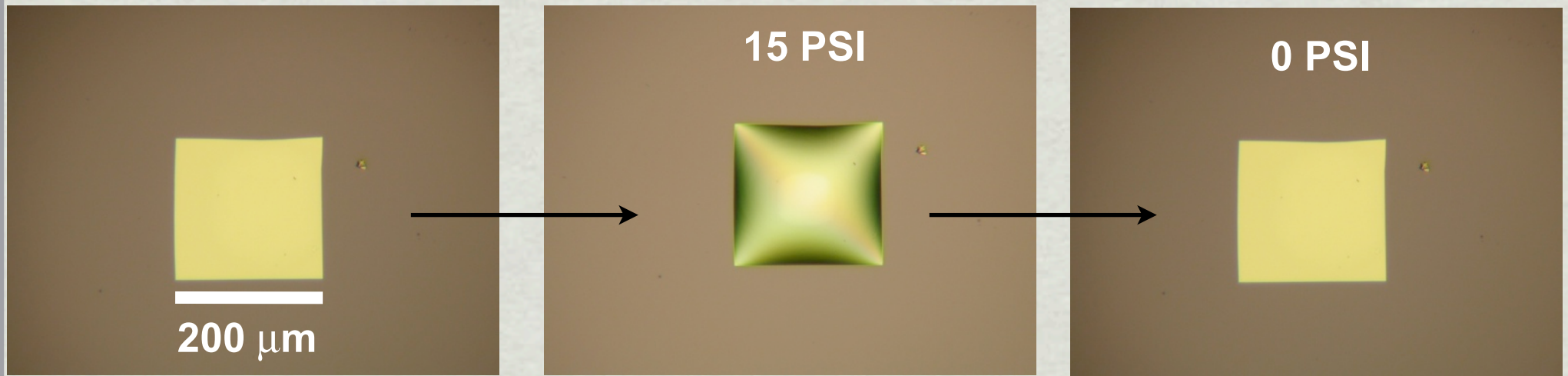
Mechanically Robust



Striemer, C.S, et al. 2007, Nature, 445:749-53

 NRG 2009

Mechanically Robust



Striemer, C.S, et al. 2007, Nature, 445:749-53

 NRG 2009

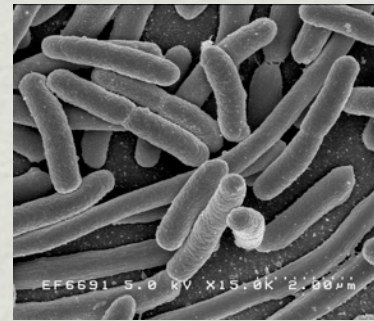
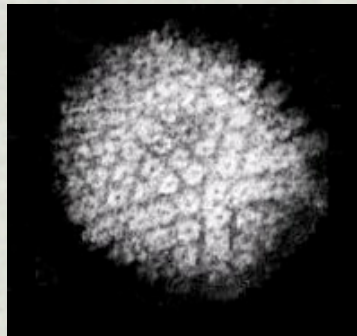
Key Biological Sizes

proteins

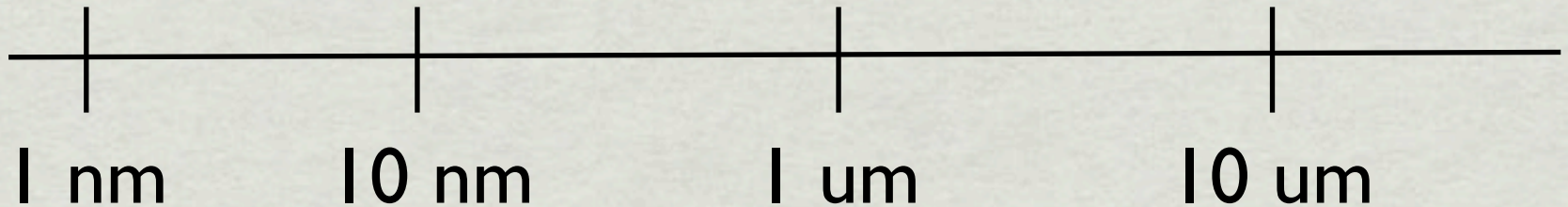
virus

bacteria

eukaryotic
cells



small
solutes



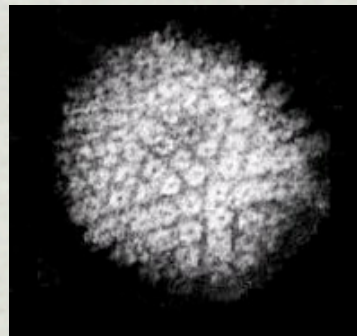
Key Biological Sizes

proteins

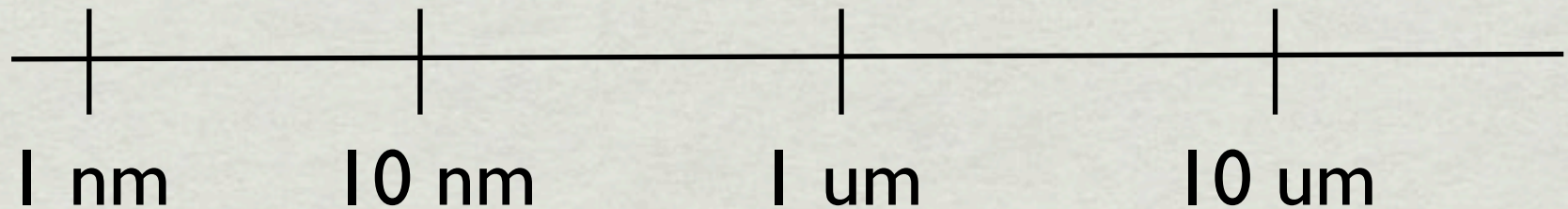
virus

bacteria

eukaryotic
cells



small
solute



PNC-SI (5-80 NM)

 NRG 2009

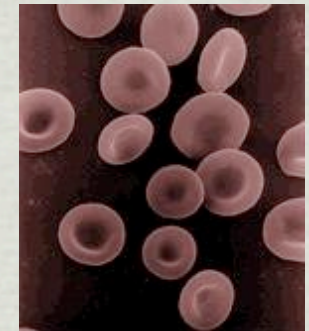
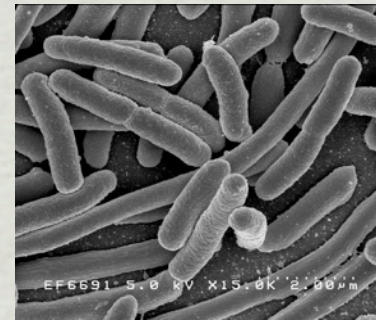
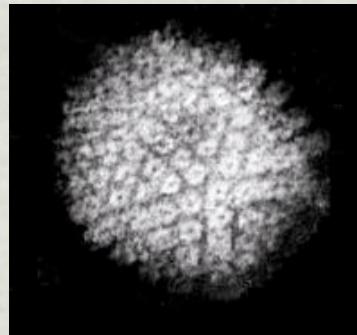
Key Biological Sizes

proteins

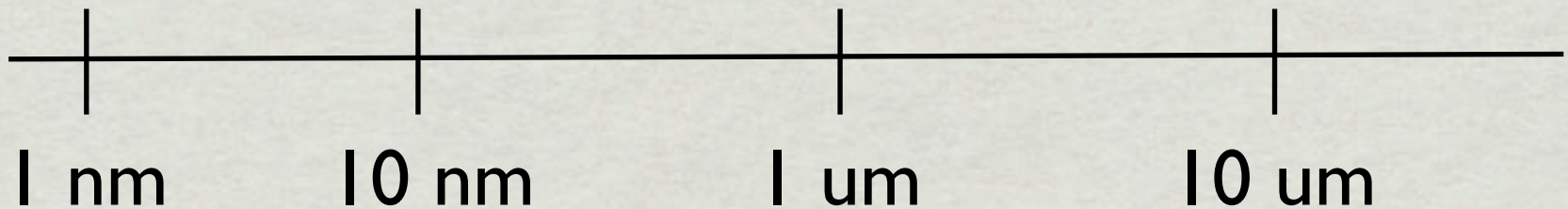
virus

bacteria

eukaryotic
cells



small
solute

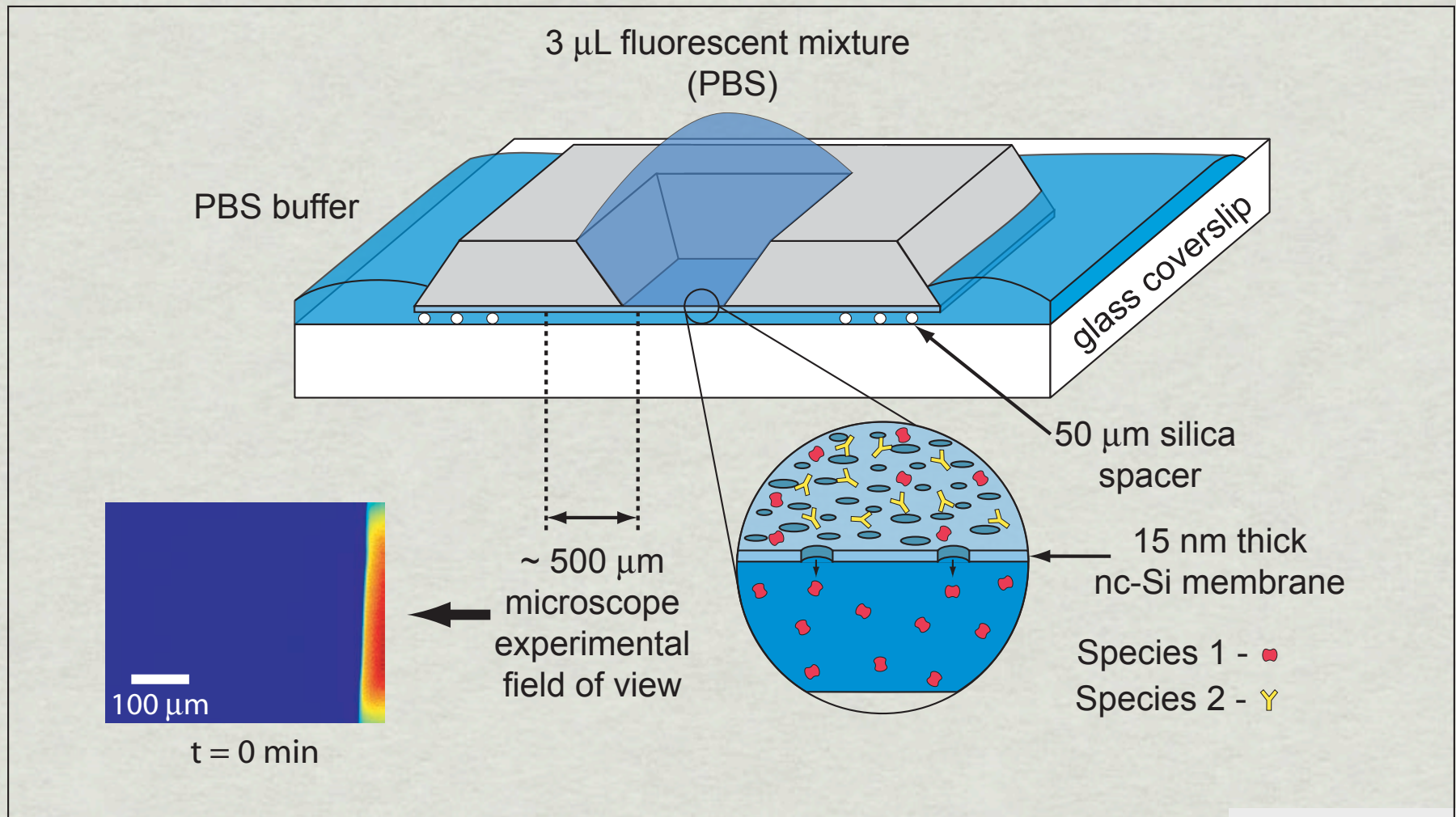


PNC-SI (5-80 NM)

@ 0.1%-15%

 **NRG 2009**

Size Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

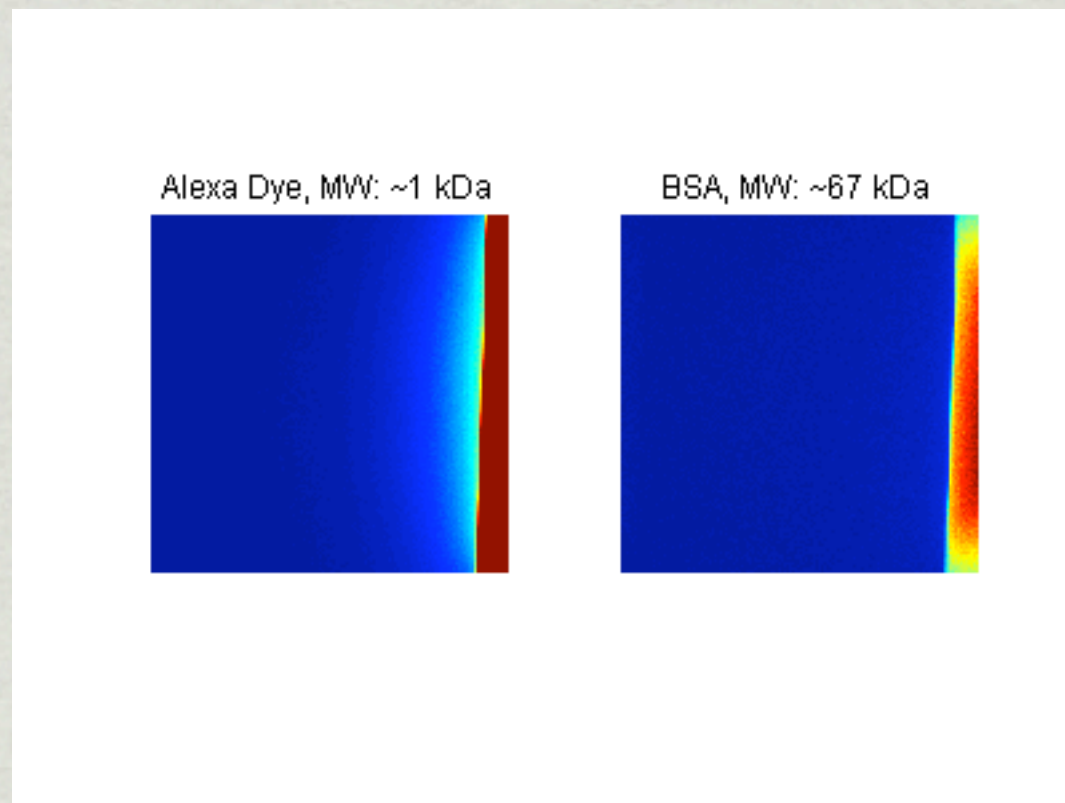
NRG 2009

Size Based Separation

Striemer, C.S, et al. 2007, Nature, 445:749-53

 **NRG 2009**

Size Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

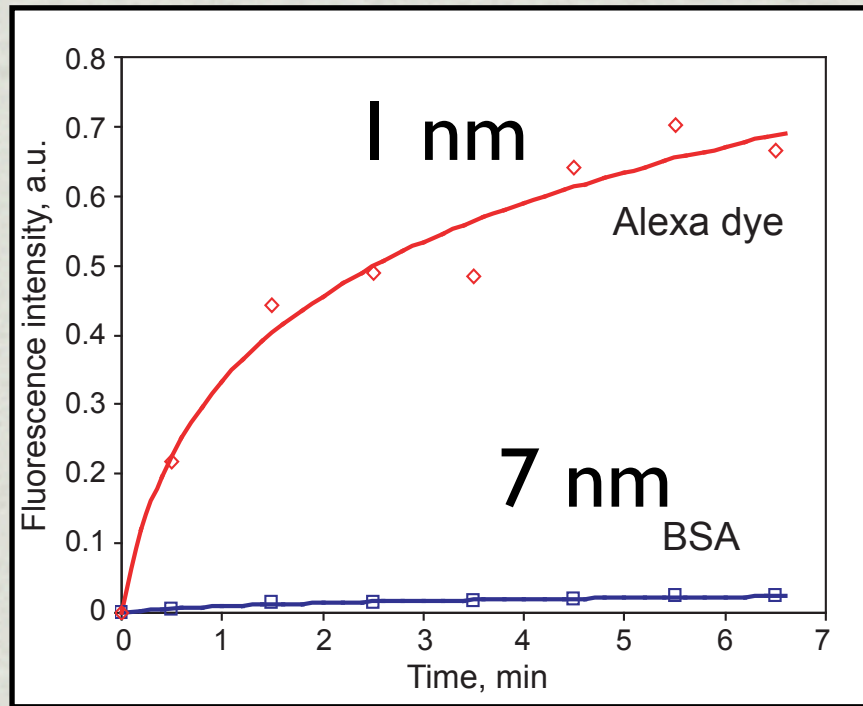
 NRG 2009

Size Based Separation

Striemer, C.S, et al. 2007, Nature, 445:749-53

 **NRG 2009**

Size Based Separation

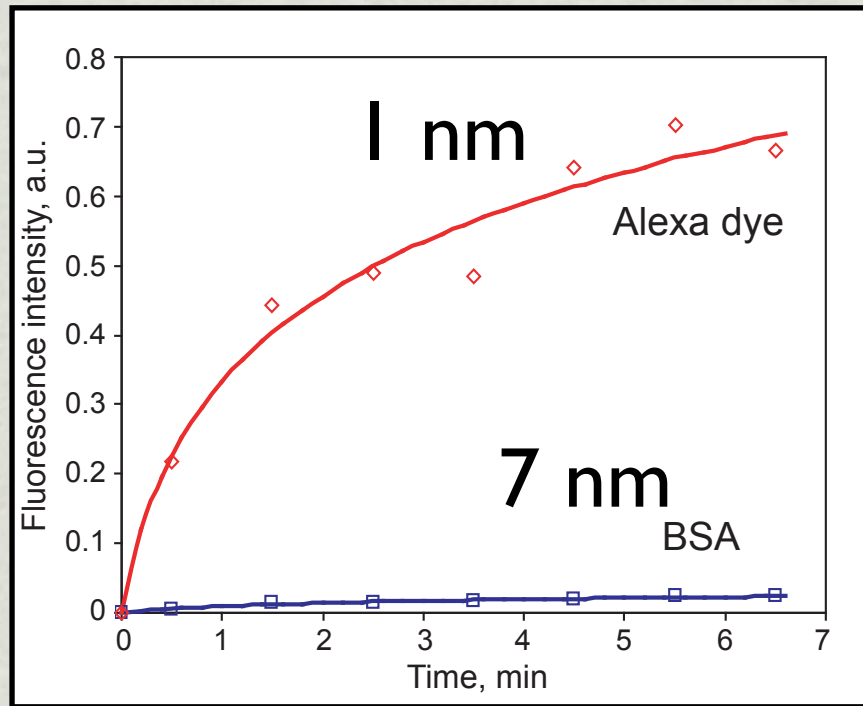


MEMBRANE A

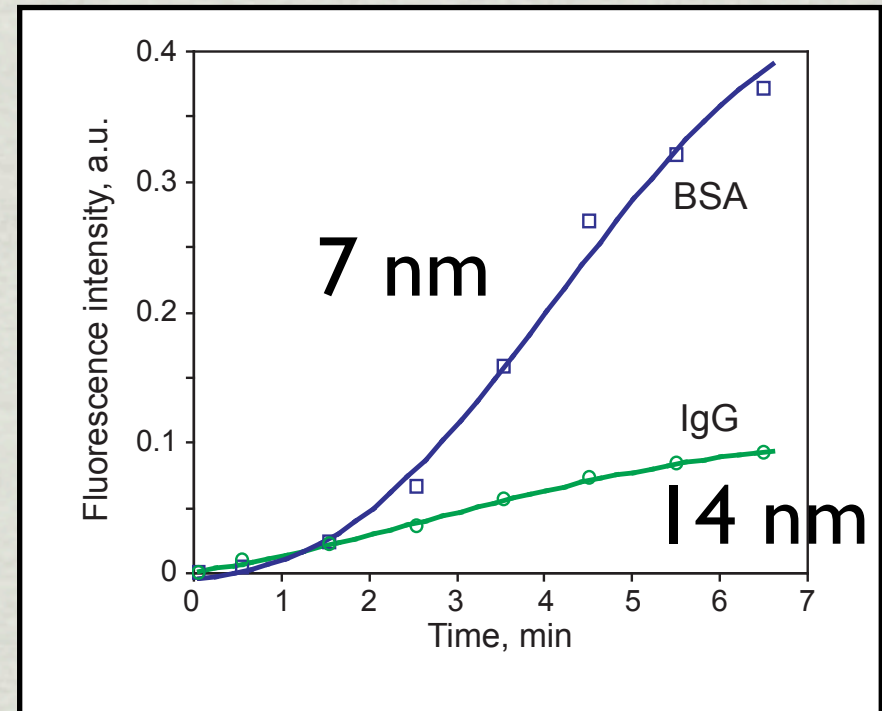
Striemer, C.S, et al. 2007, Nature, 445:749-53

 **NRG 2009**

Size Based Separation



MEMBRANE A

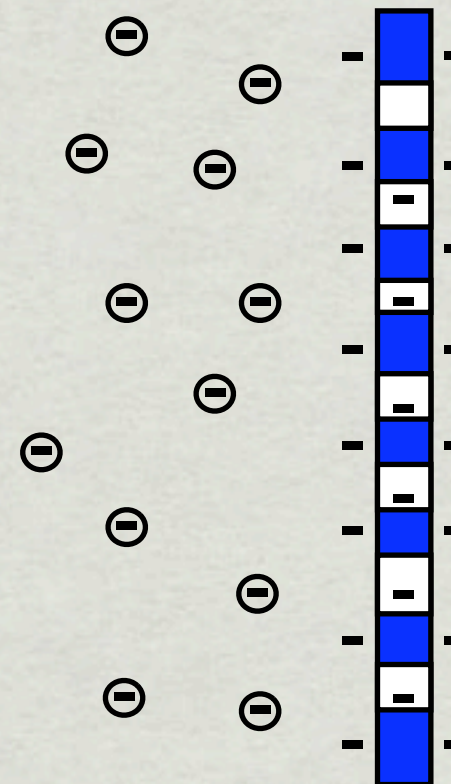
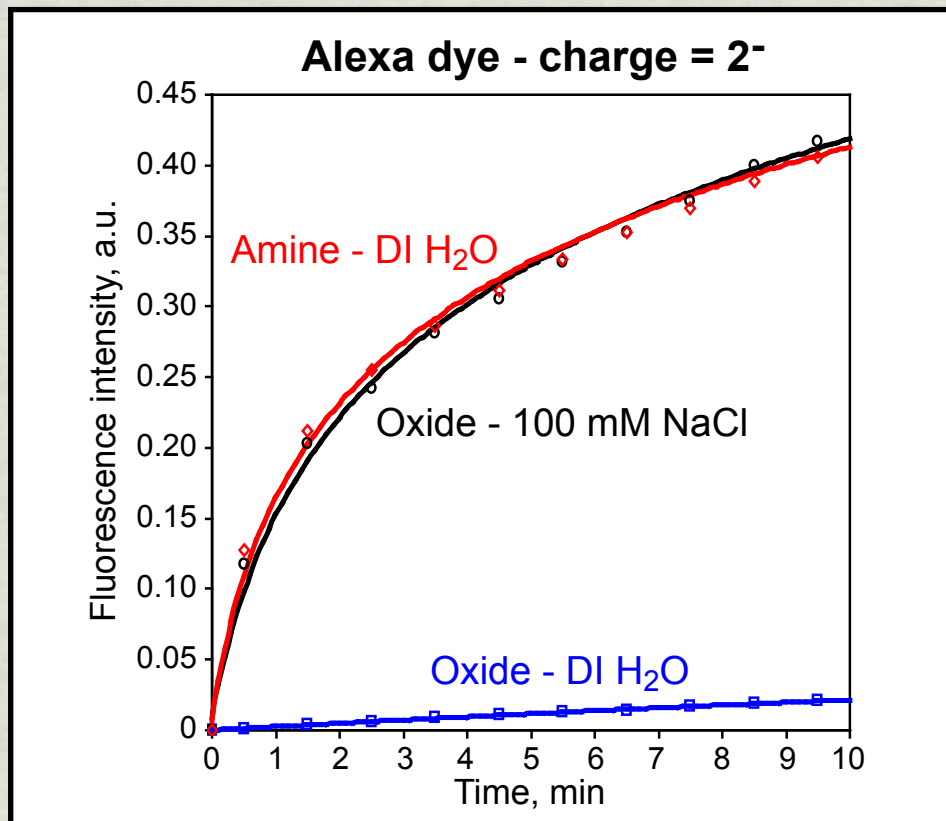


MEMBRANE B

Striemer, C.S, et al. 2007, Nature, 445:749-53

NRG 2009

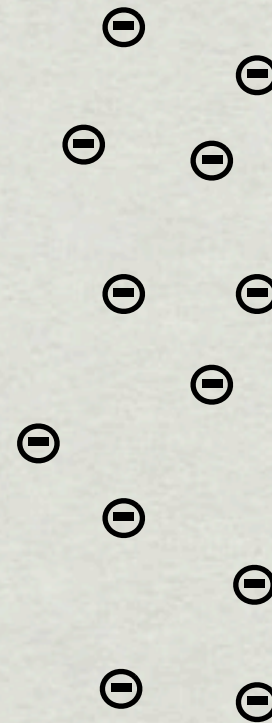
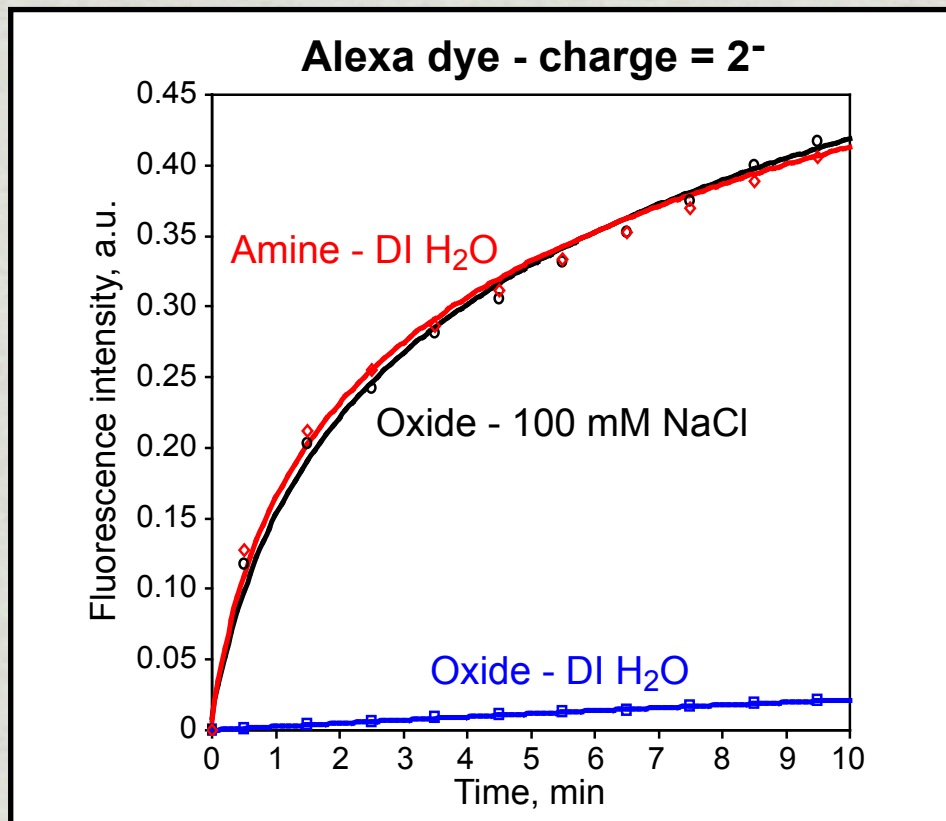
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

NRG 2009

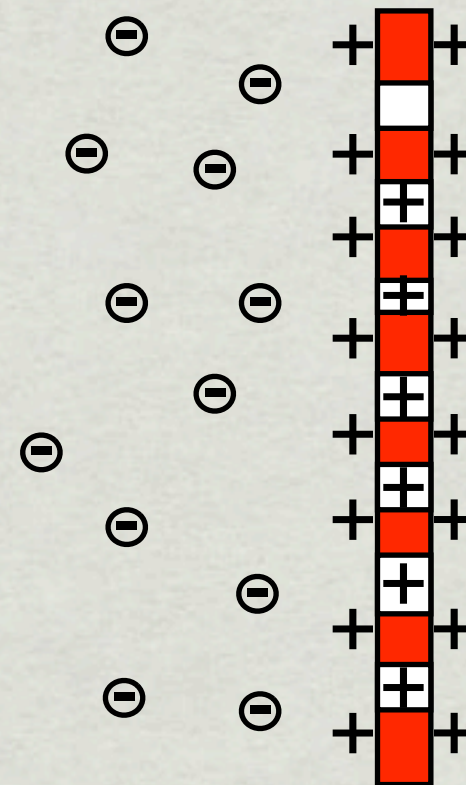
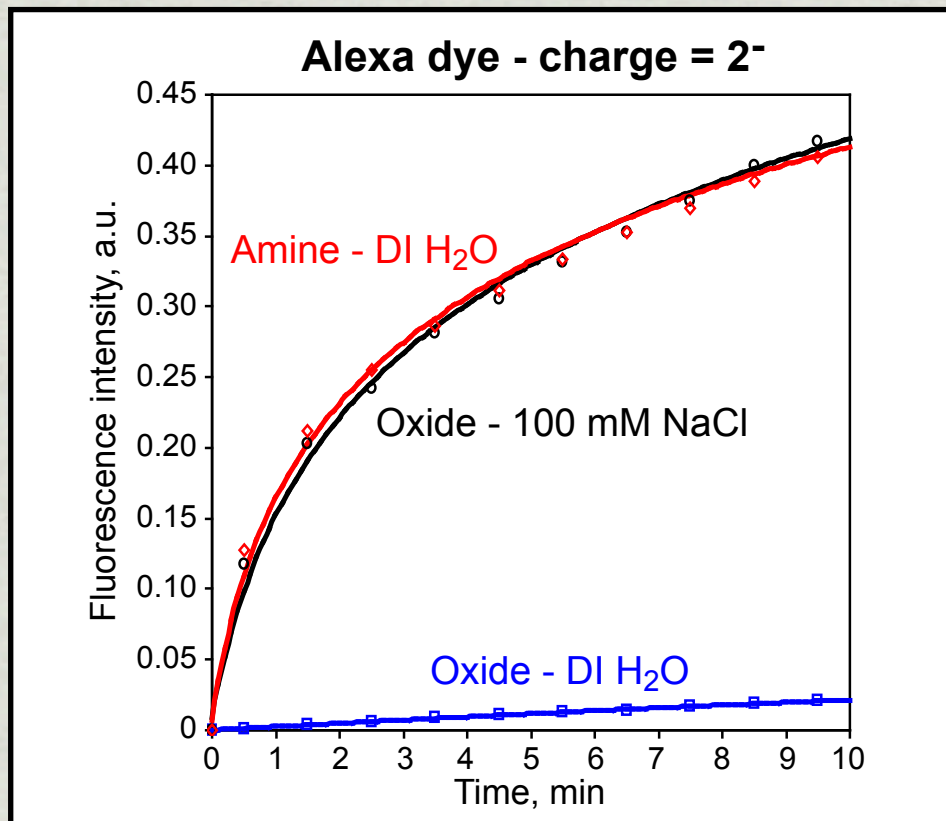
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

 NRG 2009

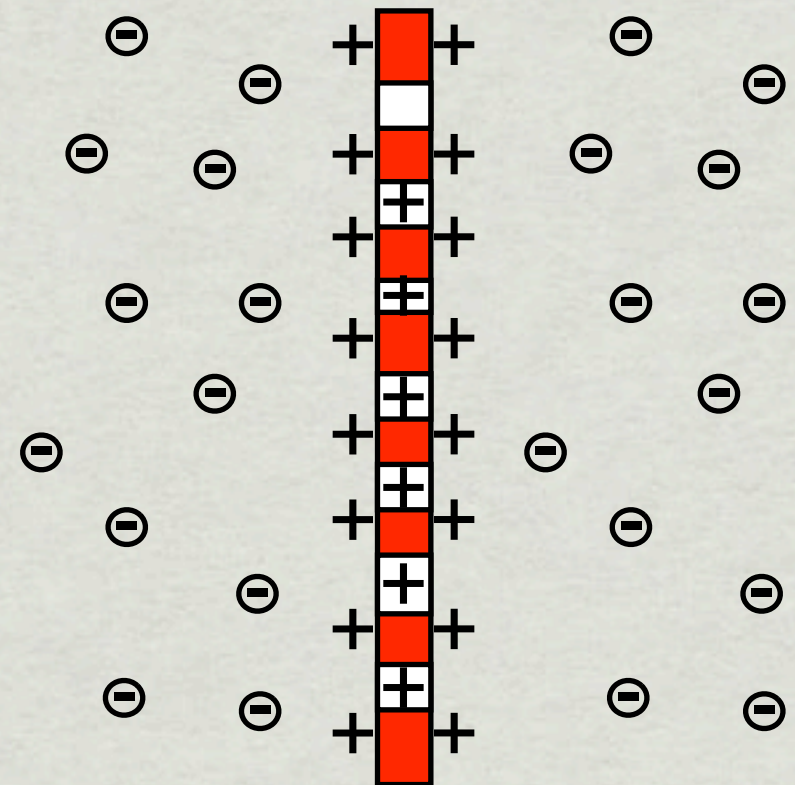
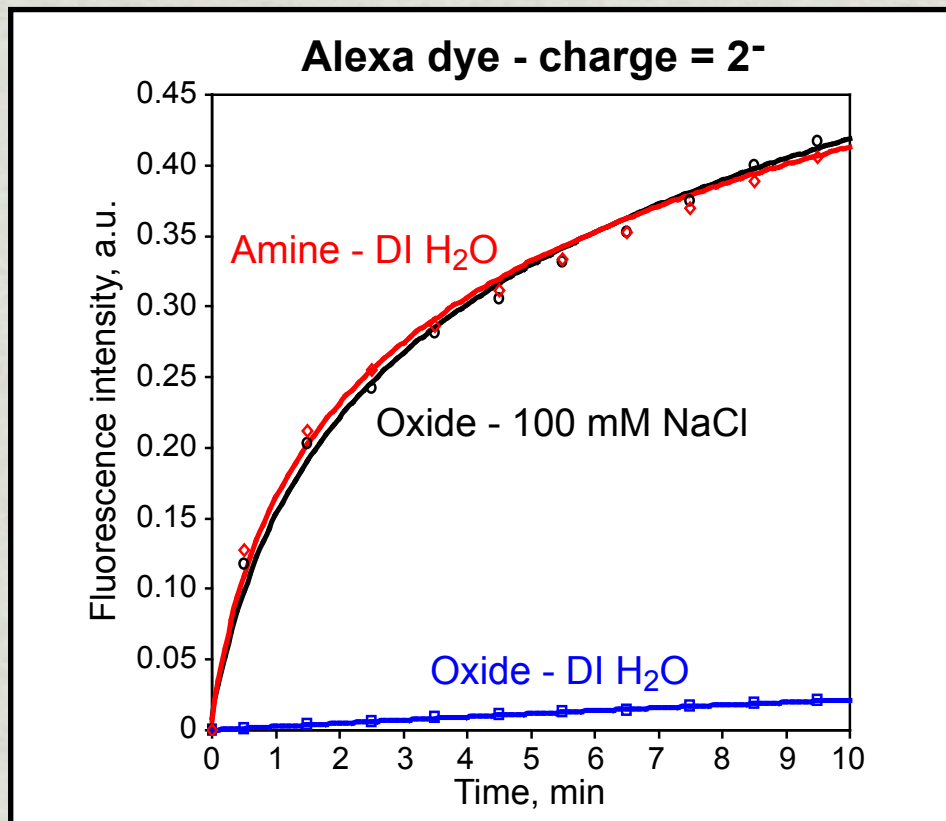
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

NRG 2009

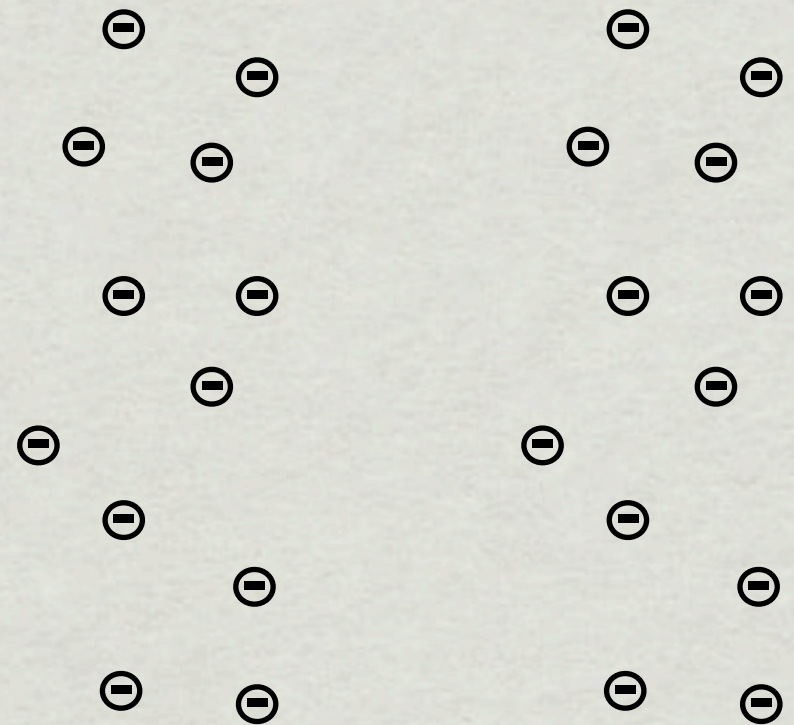
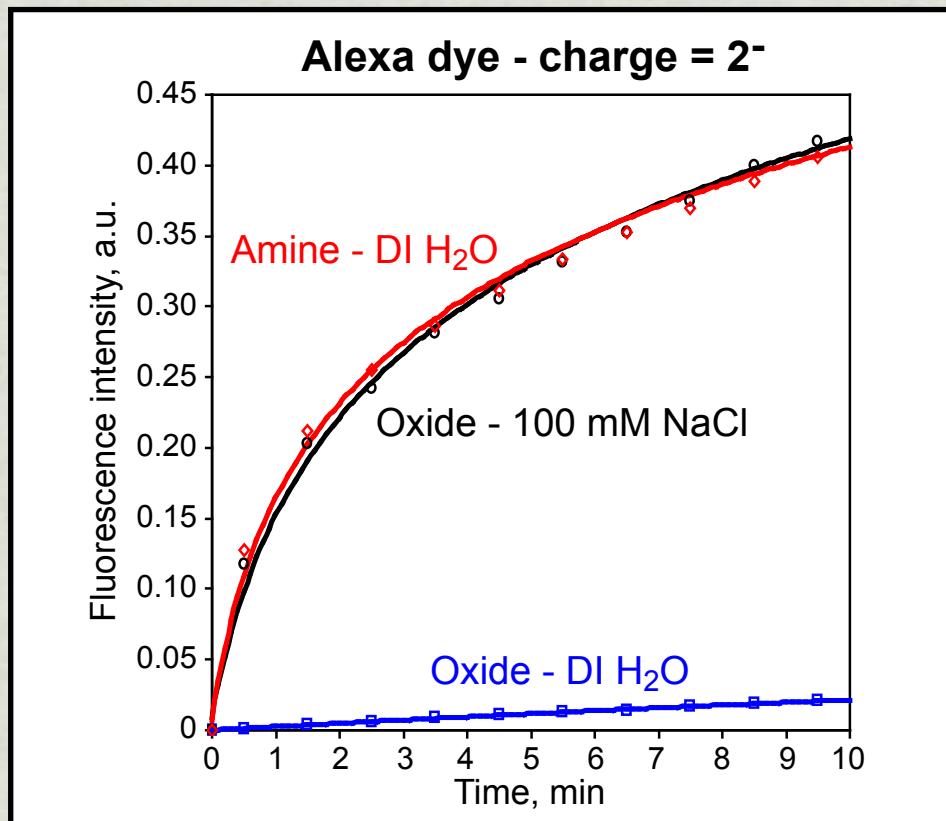
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

NRG 2009

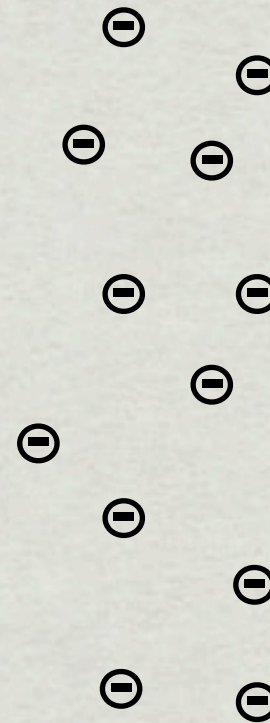
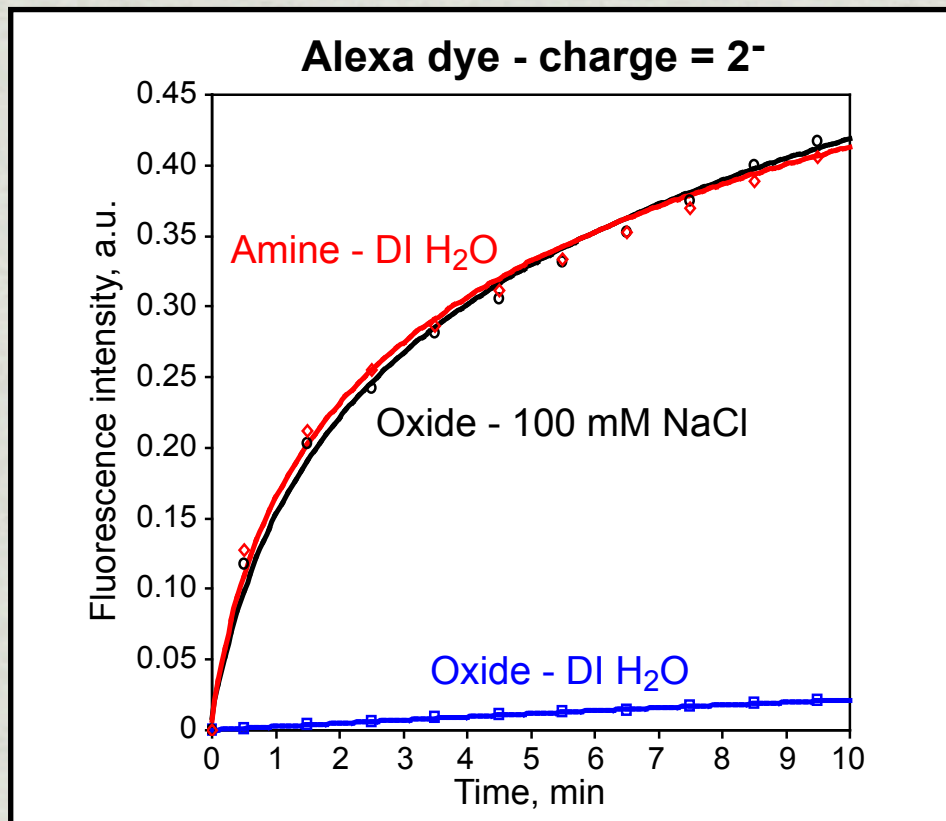
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

NRG 2009

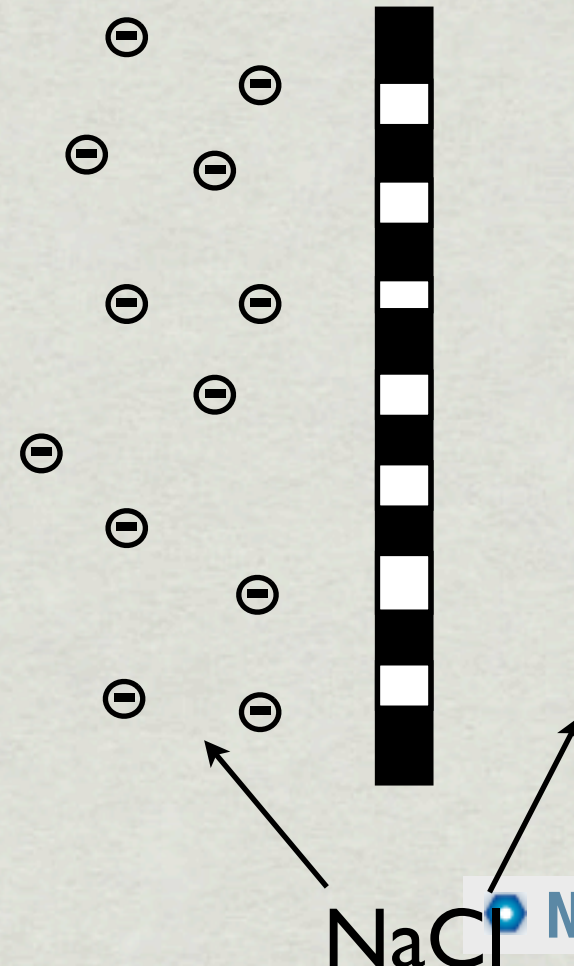
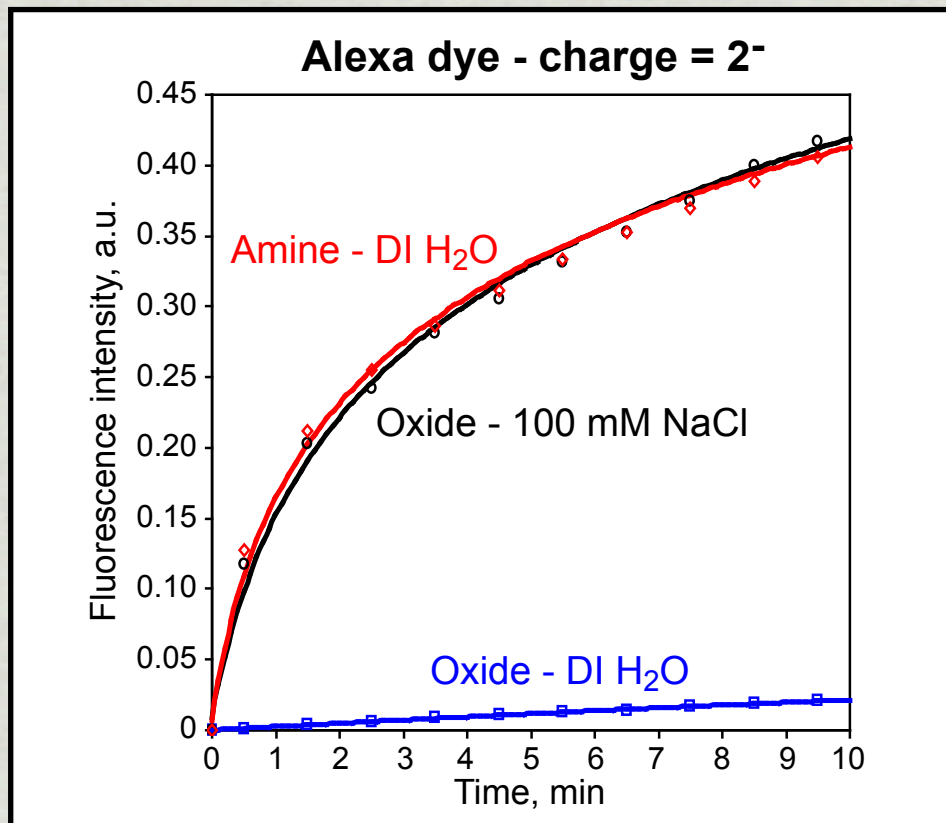
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

 NRG 2009

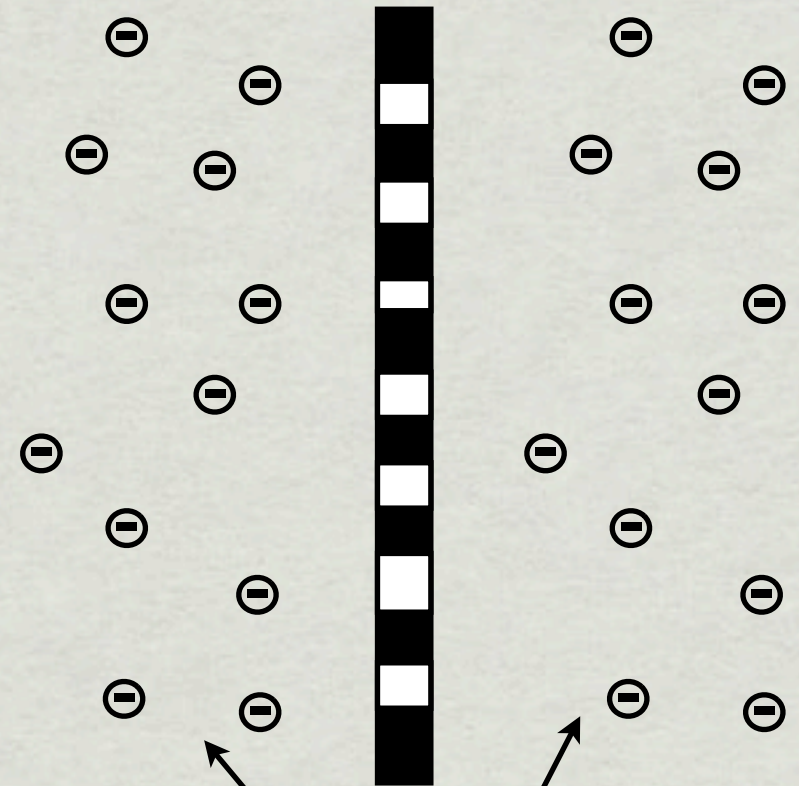
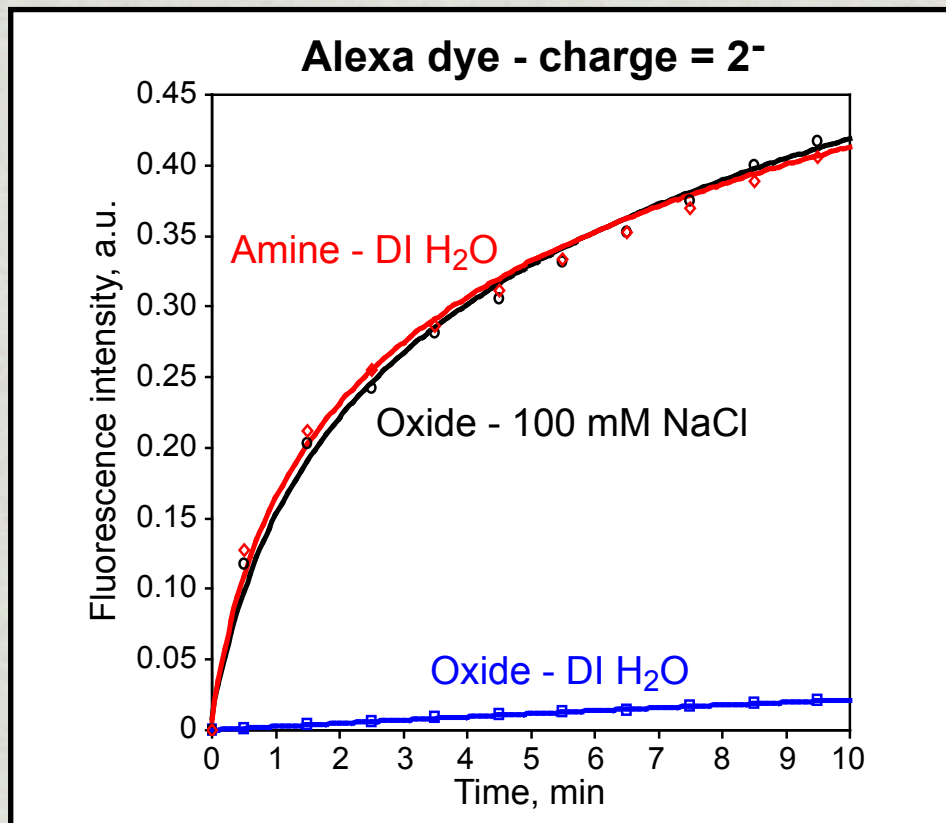
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

NaCl NRG 2009

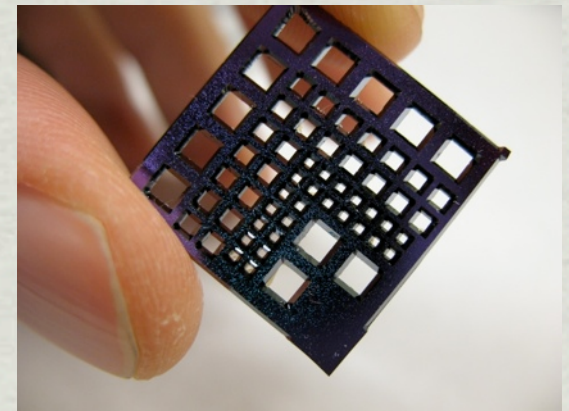
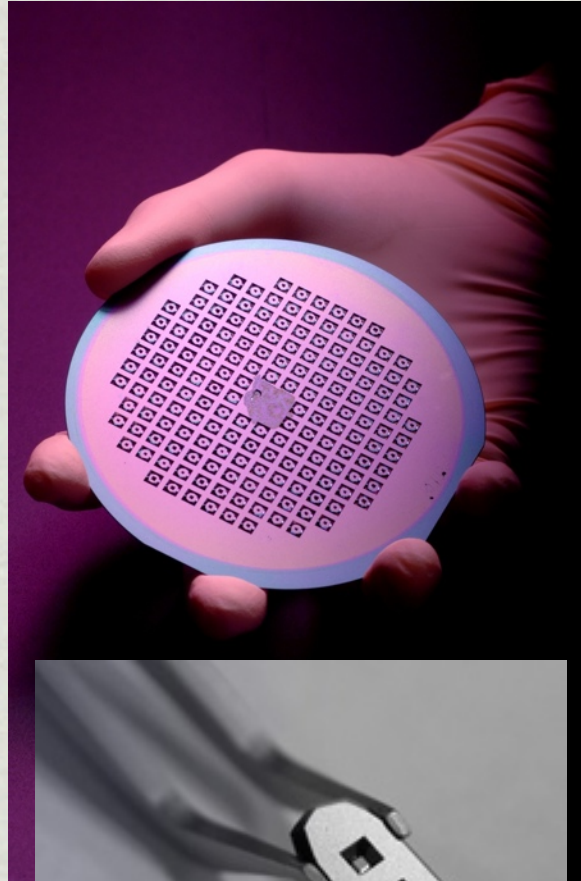
Charge Based Separation



Striemer, C.S, et al. 2007, Nature, 445:749-53

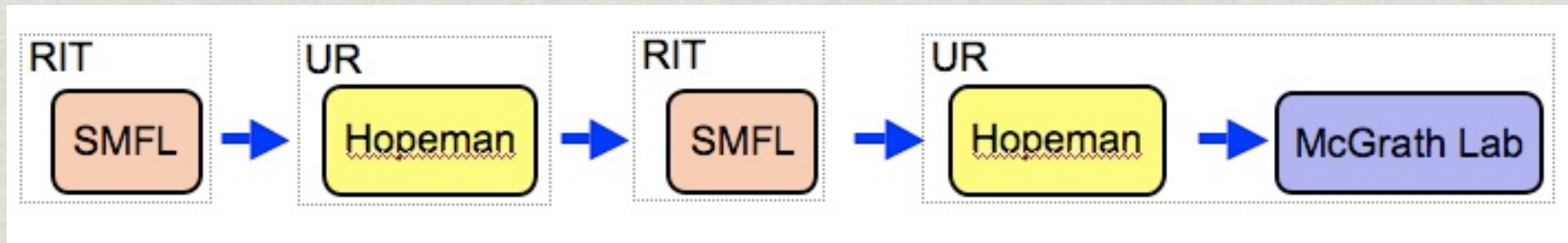
NaCl NRG 2009

Scalable, flexible, economical fabrication



Production Flow

PAST



PATTERNING

DEPOSITION

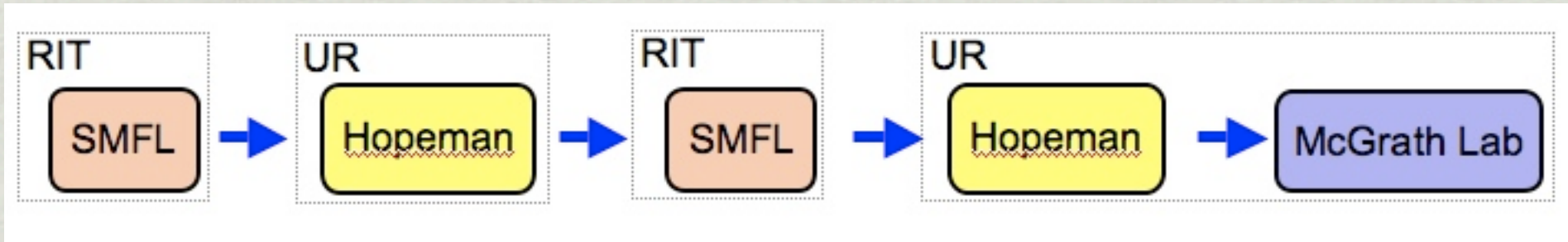
**THERMAL
ANNEALING**

ETCHING

**TESTING &
APPLICATIONS**

Production Flow

PAST



PATTERNING

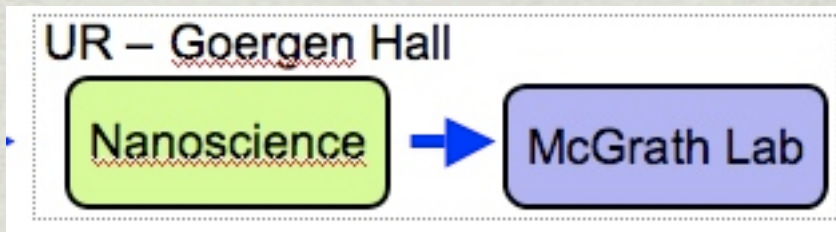
DEPOSITION

THERMAL
ANNEALING

ETCHING

TESTING &
APPLICATIONS

NEAR FUTURE

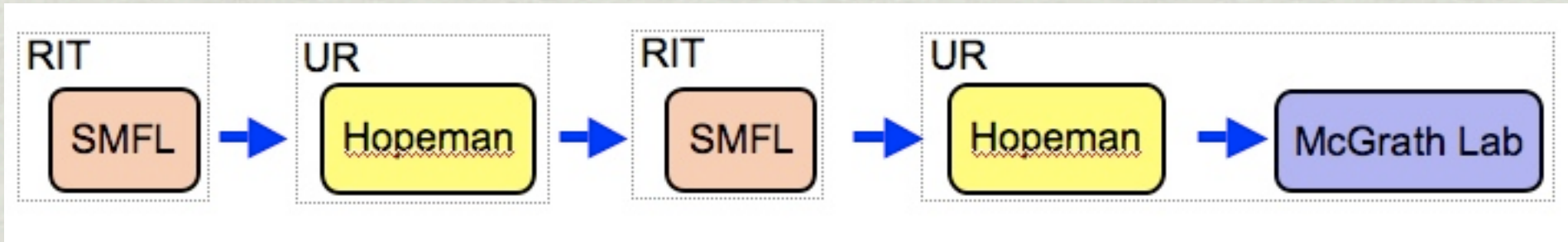


ALL FABRICATION

TESTING &
APPLICATIONS

Production Flow

PAST



PATTERNING

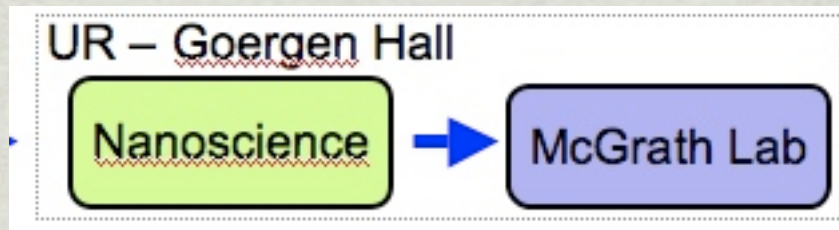
DEPOSITION

THERMAL
ANNEALING

ETCHING

TESTING &
APPLICATIONS

NEAR FUTURE



ALL FABRICATION

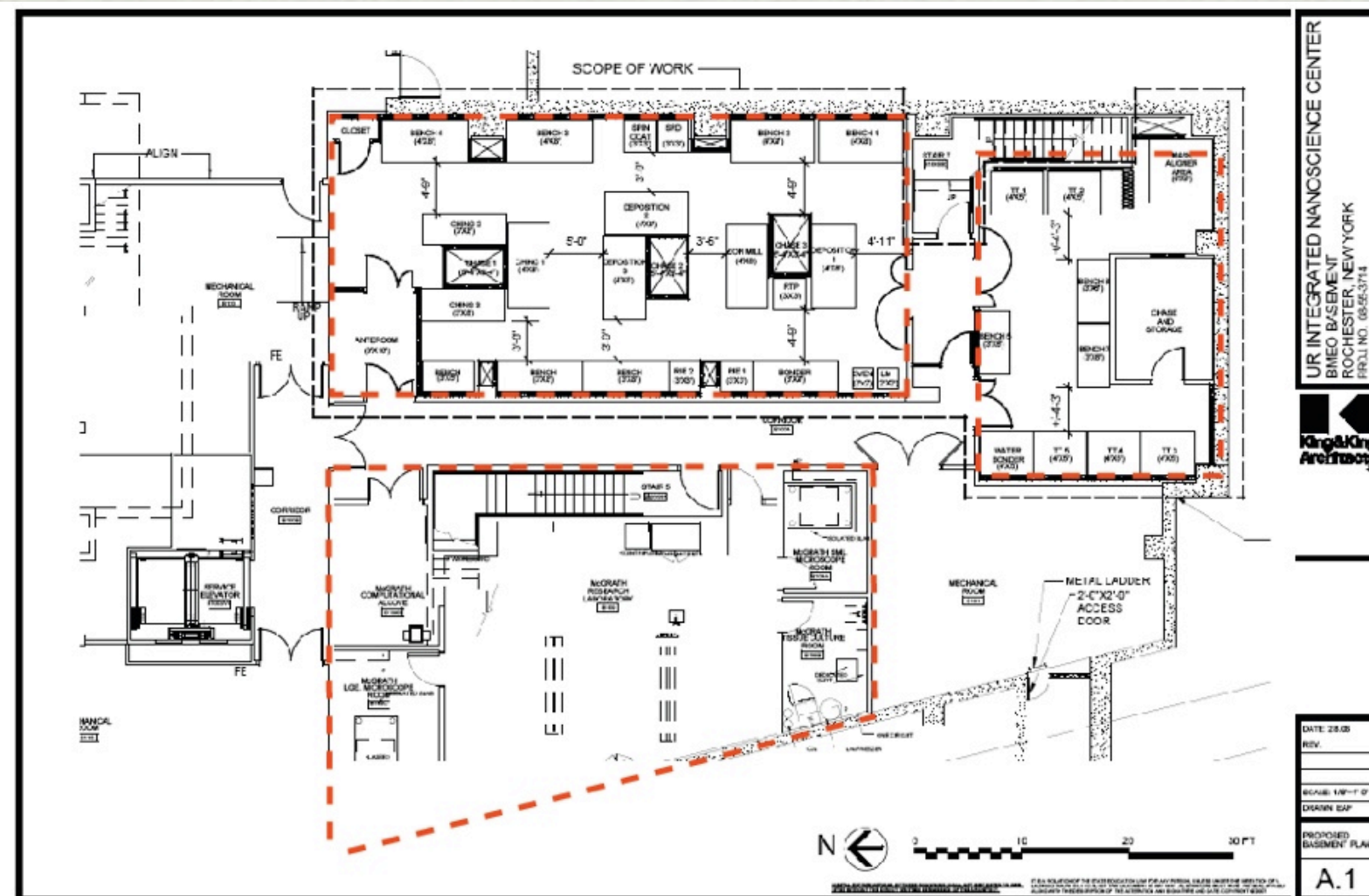
TESTING &
APPLICATIONS

**MADE POSSIBLE BY
PURCHASE OF NEW SYSTEMS**

- SURFACE PREP
- ANNEALING
- DEPOSITION

 NRG 2009

UR's Nanoscience Center



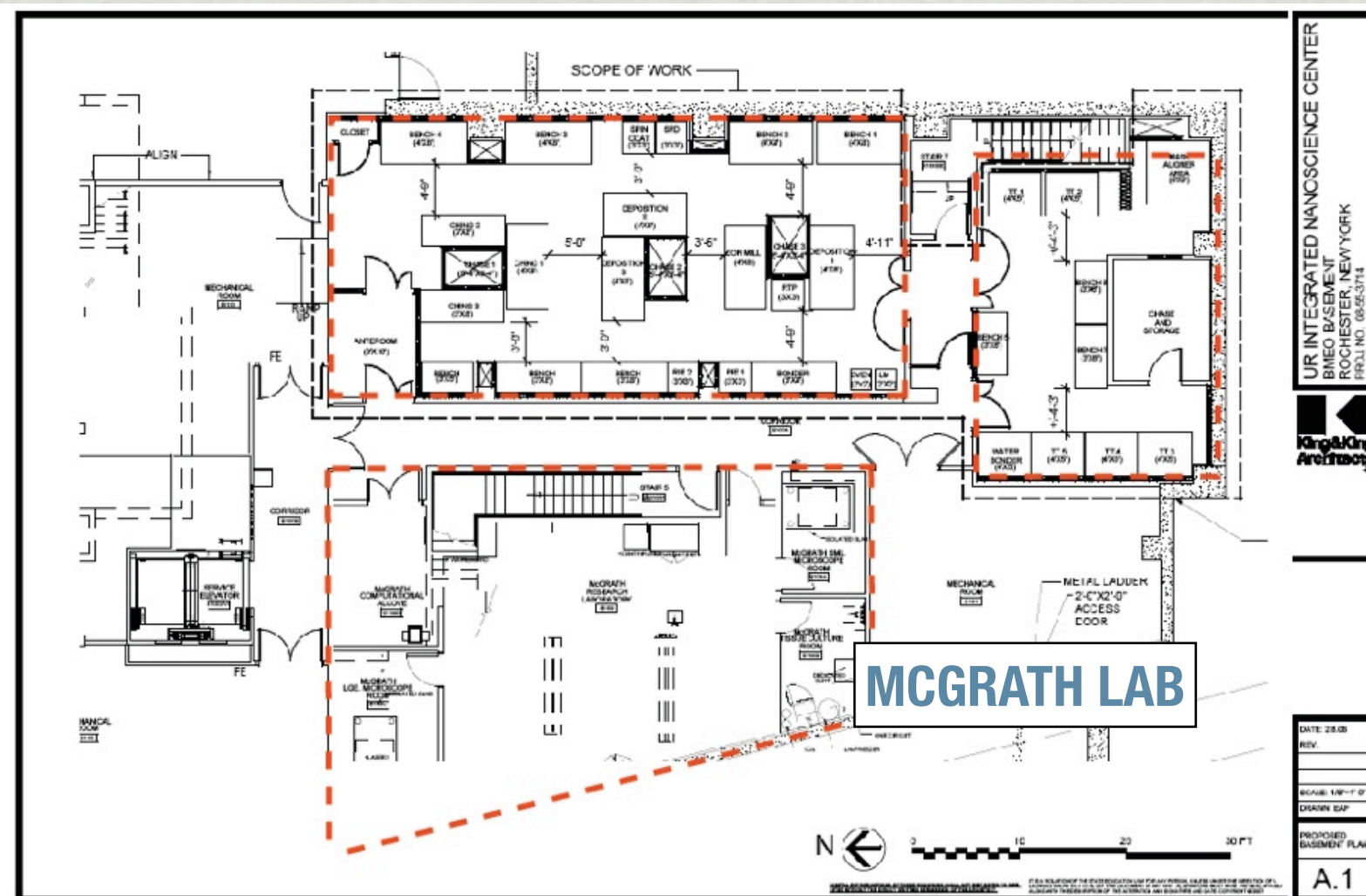
UR INTEGRATED NANOSCIENCE CENTER
 BME0 BASEMENT
 ROCHESTER, NEW YORK
 PROJ. NO. 06-55-3714



DATE: 28.03
REV:
SCALE: 1/8" = 1'-0"
DIAGN. EAP
PROPOSED BASEMENT PLAN
A.1



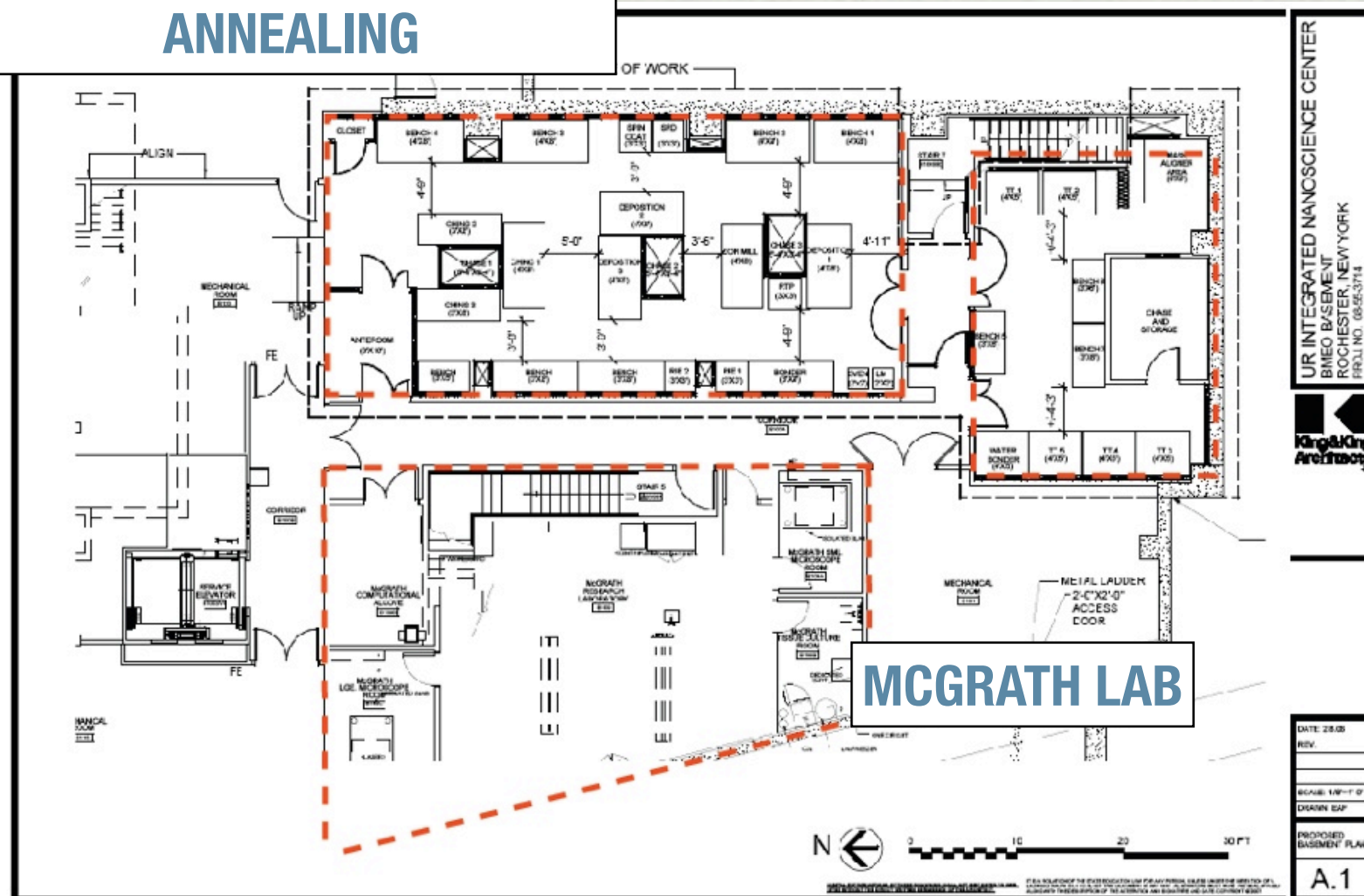
UR's Nanoscience Center



NRG 2009

UR's Nanoscience Center

**SURFACE PREP, DEPOSITION
AND
ANNEALING**



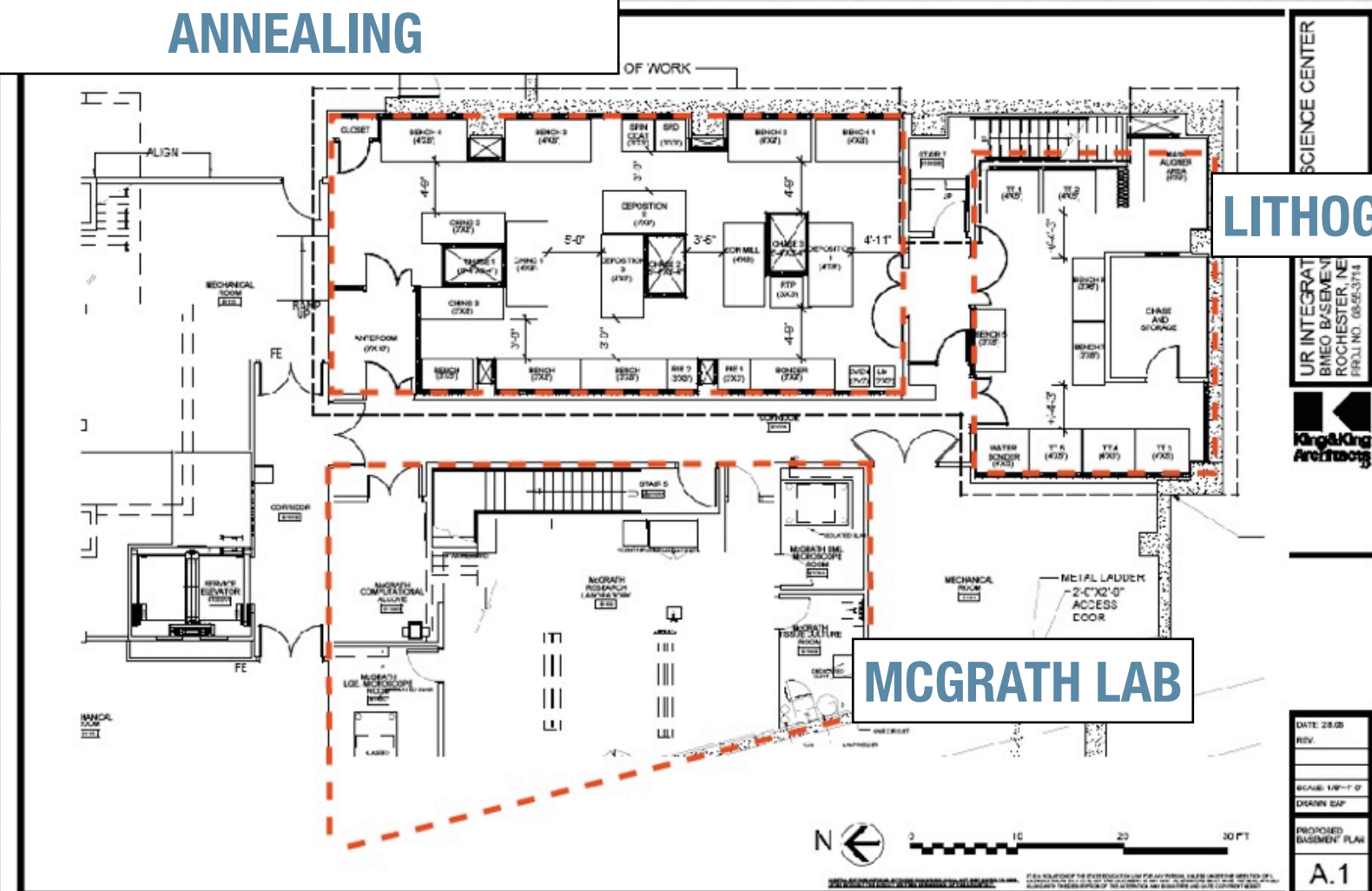
NRG 2009

UR's Nanoscience Center

**SURFACE PREP, DEPOSITION
AND
ANNEALING**

LITHOGRAPHY

MCGRATH LAB



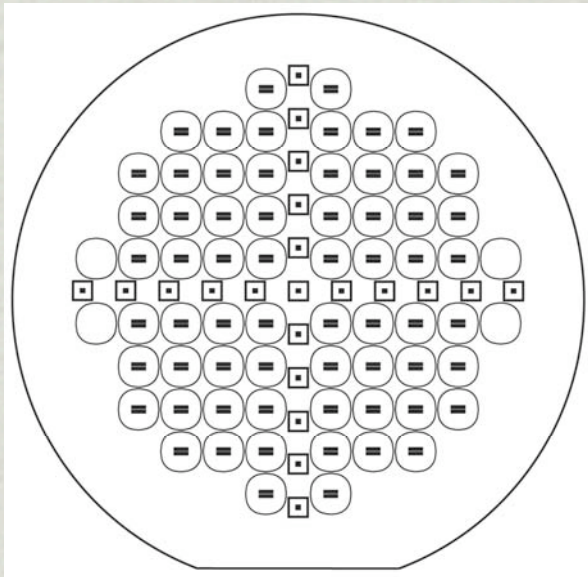
SCIENCE CENTER
UR INTEGRATED
BMEO BASEMENT
ROCHESTER, NY
PROJ. NO. 06-55-3714



DATE: 28.03
REV.
SCALE: 1/8" = 1'-0"
DIAGN: EAP
PROPOSED BASEMENT PLAN
A.1

NRG 2009

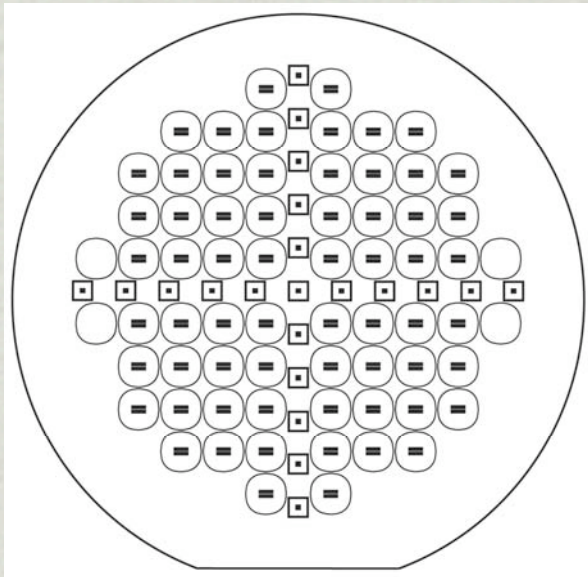
Manufacturing Possibilities and Challenges



4" wafer

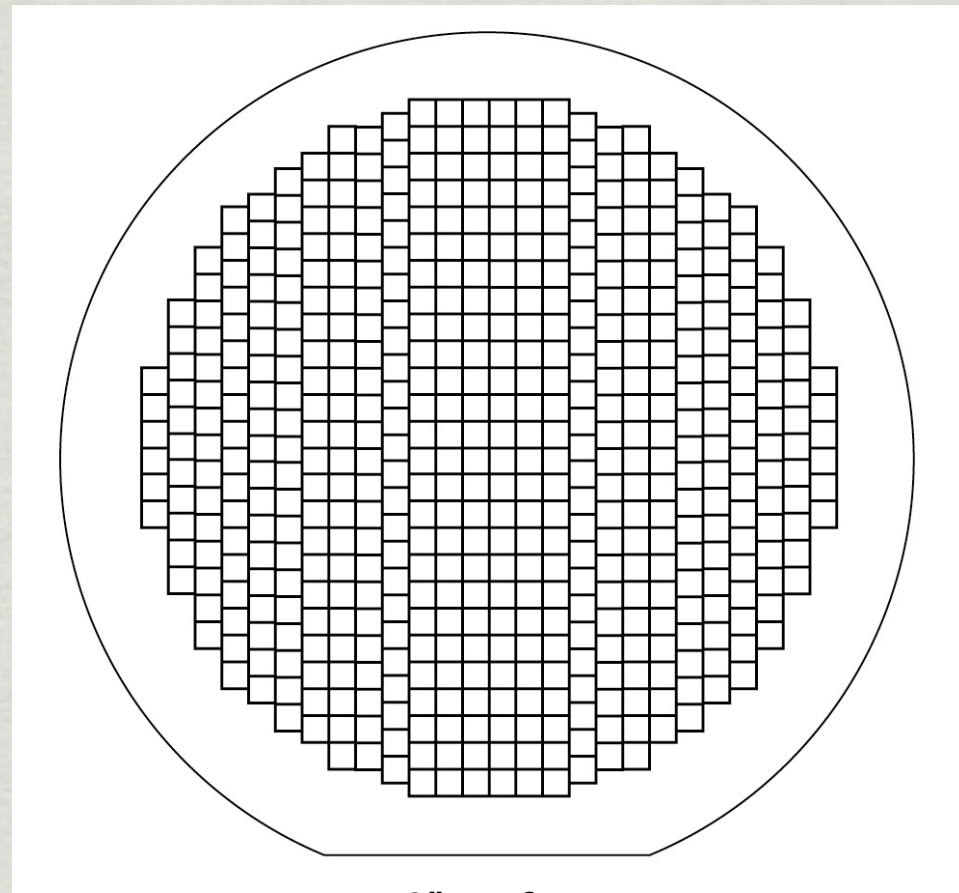
**CURRENTLY: 68
INSERTS PER WAFER**

Manufacturing Possibilities and Challenges



4" wafer

**CURRENTLY: 68
INSERTS PER WAFER**

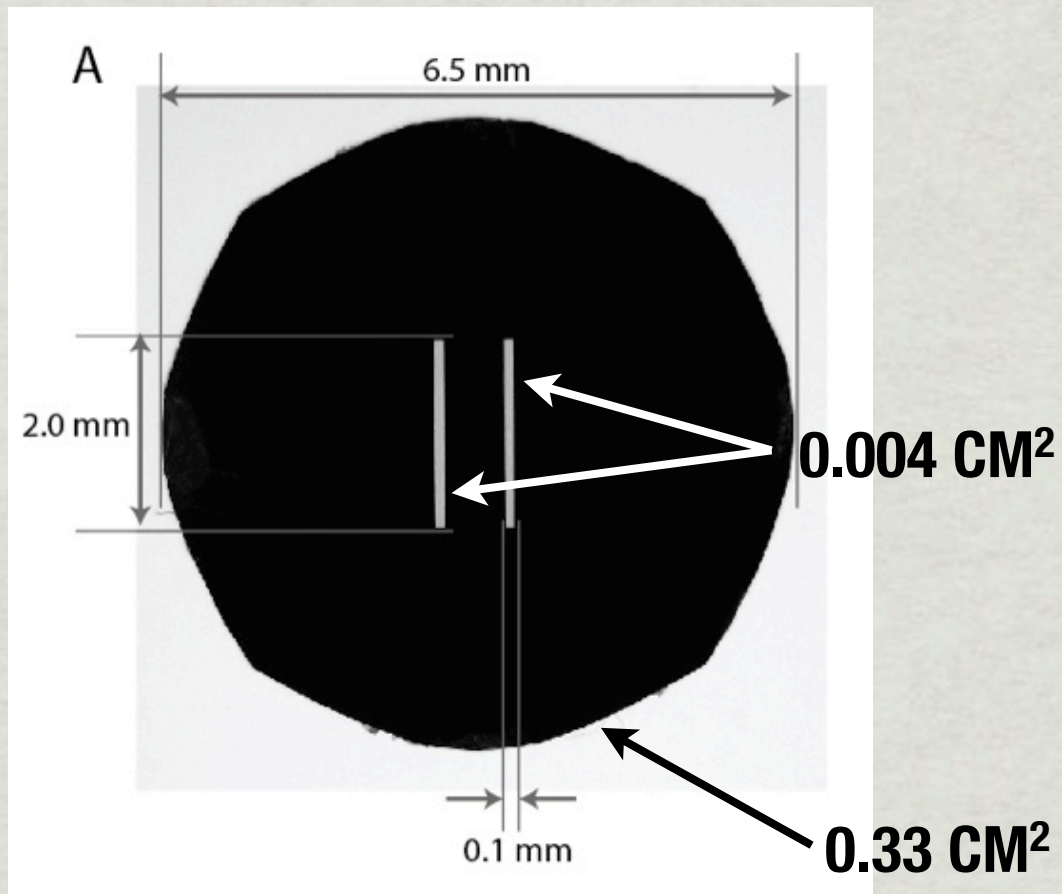


6" wafer

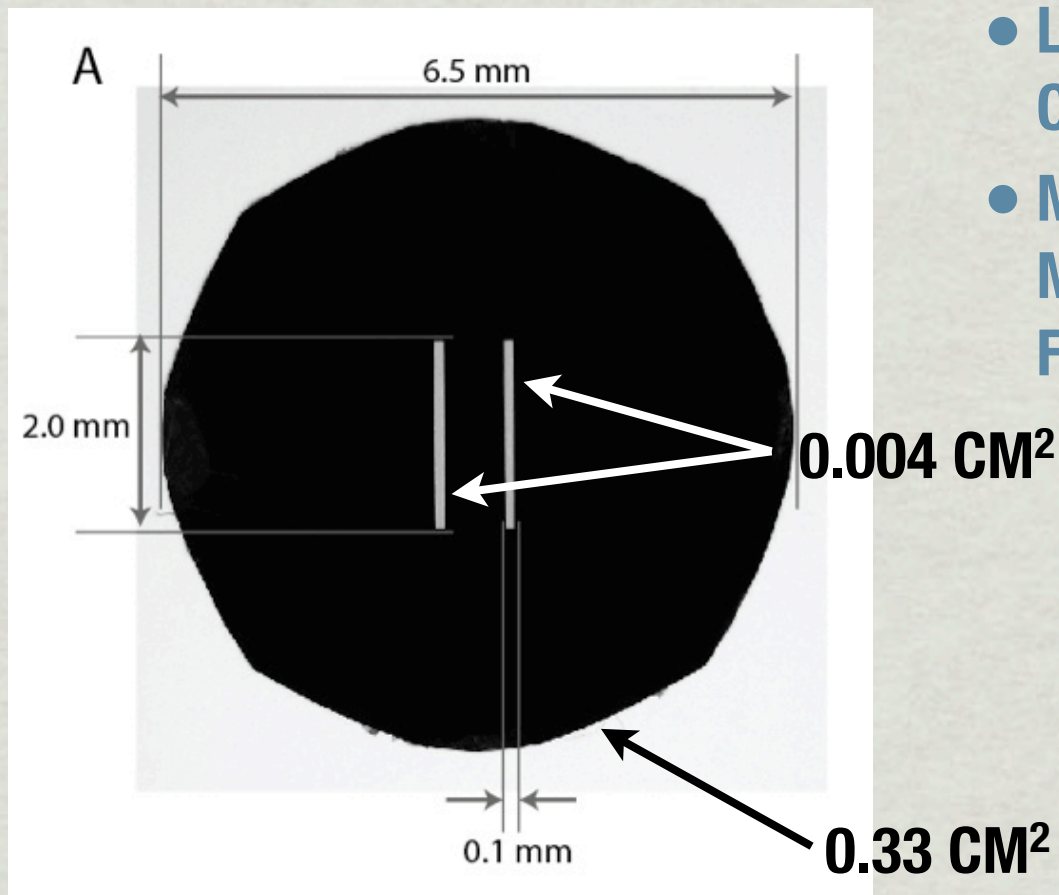
> 500 INSERTS PER WAFER

 **NRG 2009**

Manufacturing Possibilities and Challenges

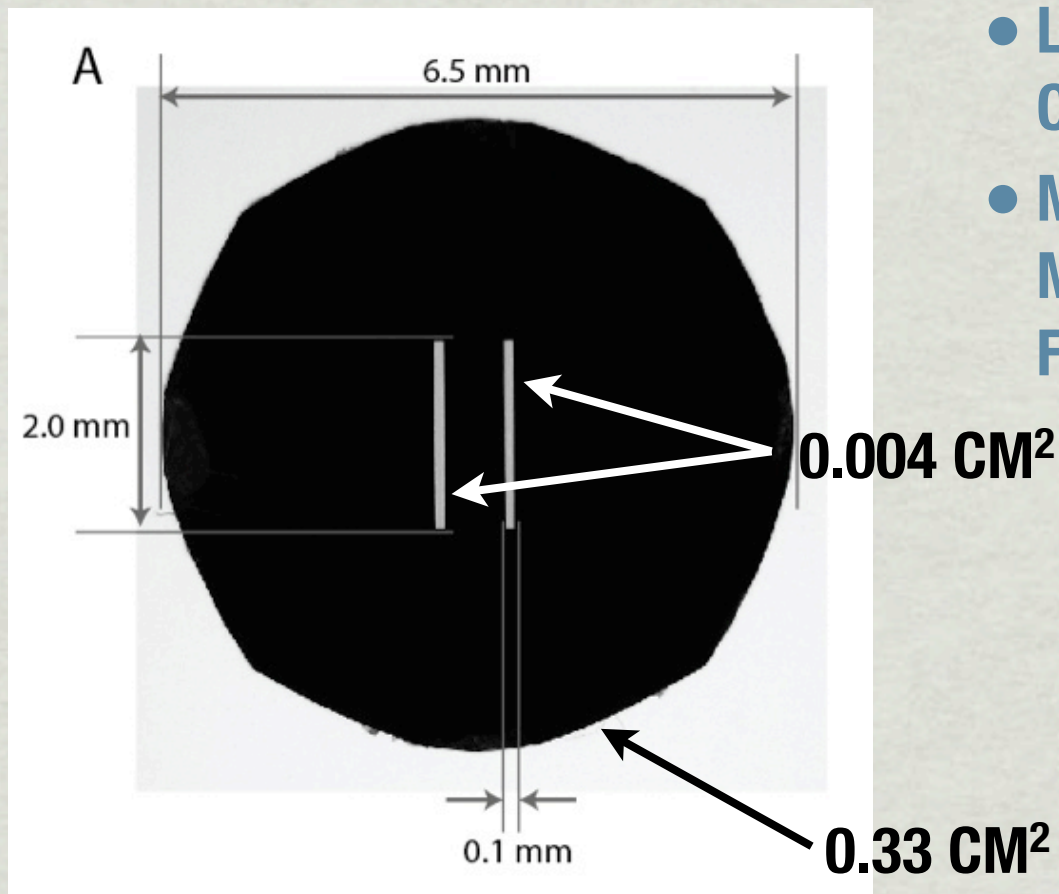


Manufacturing Possibilities and Challenges

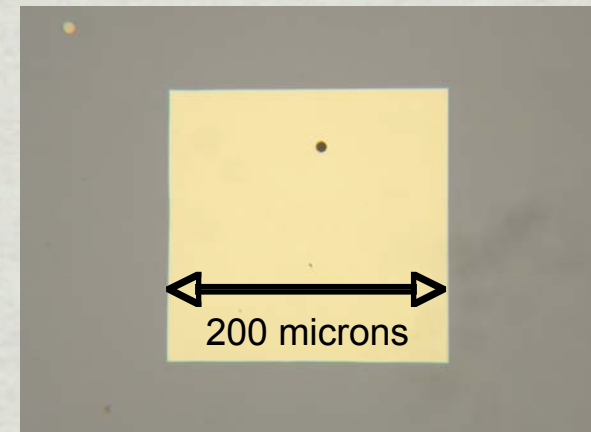


- LESS THAN 1% OF AREA IS CURRENTLY ACTIVE
- MADE NECESSARY BY ETCHING, MECHANICS, AND DEFECT FREQUENCY

Manufacturing Possibilities and Challenges

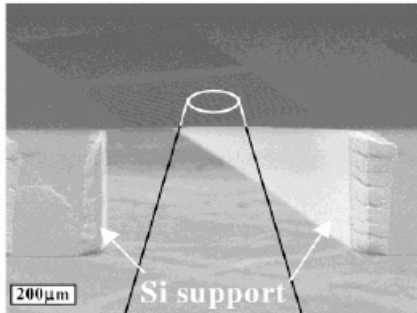


- LESS THAN 1% OF AREA IS CURRENTLY ACTIVE
- MADE NECESSARY BY ETCHING, MECHANICS, AND DEFECT FREQUENCY

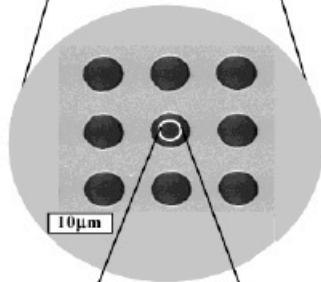


Ultrathin SiN membranes

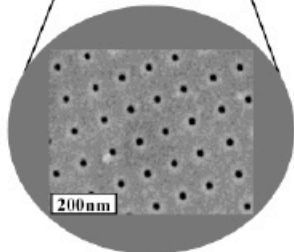
a) Si frame holding a microsieve



b) SiN microsieve holding a nanosieve



c) SiN nanosieve

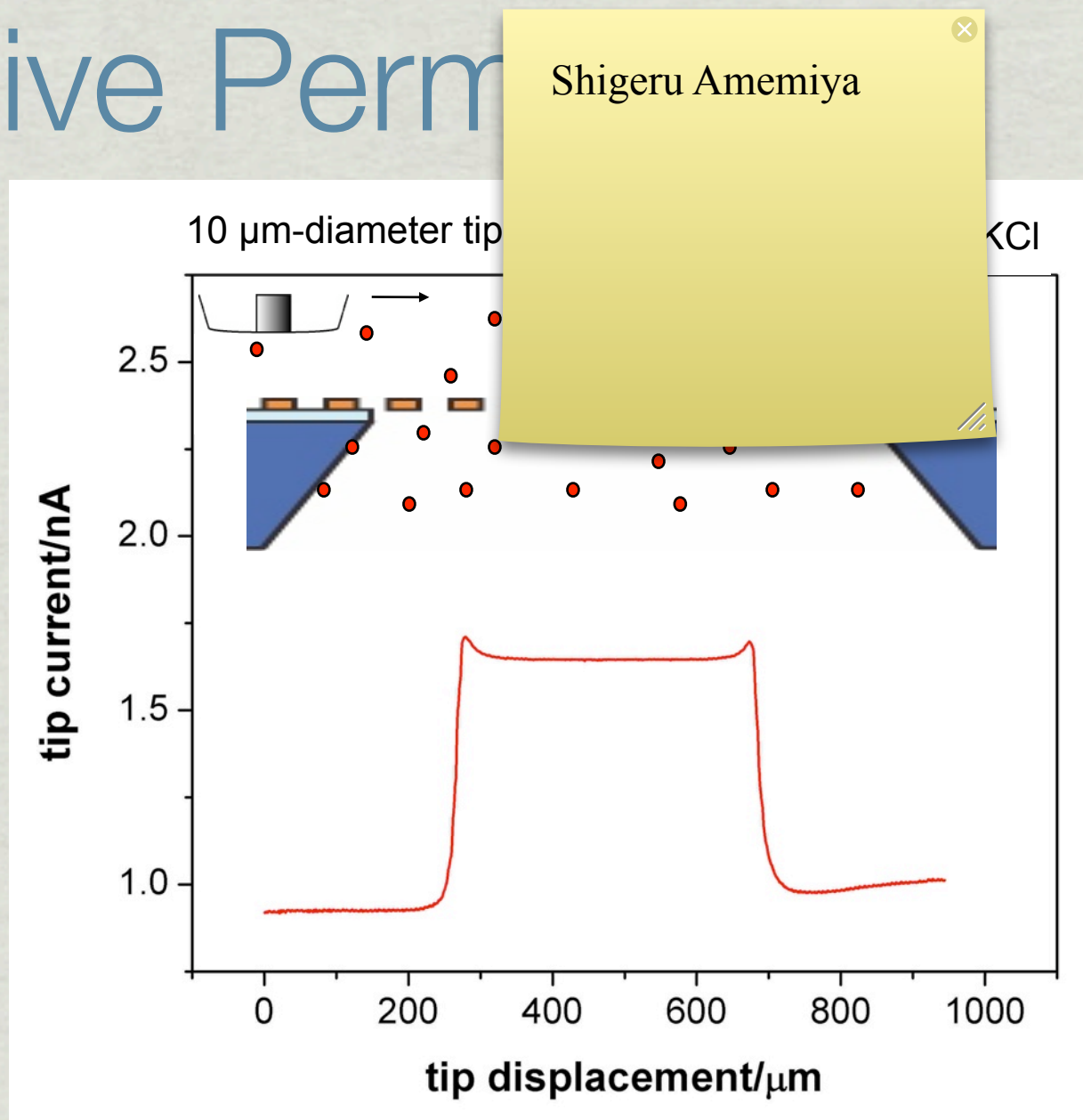


Tong, et al. (2004). Silicon Nitride Nanosieve Membrane. Nano Letters 4, 283-287

- * SiN - 10 nm thick but elaborate and impractical
- * No demonstrated separations
- * Appreciated all the potential and issues (air flow, water permeability, and mechanics)

High Diffusive Perm

- * Intrinsic permeability of 5.2×10^{-2} cm/s measured using Scanning Electrochemical Microscopy.
- * Experimental results & pore histograms are consistent with theory that neglects pore resistance
- * Between 2-3 orders higher than small molecule diffusion permeability of reconstituted cellulose or PES.



Kim et al. 2008, JACS 130:4230-4231

NRG 2009

High Diffusive Perm

Shigeru Amemiya

- * Intrinsic permeability of 5.2×10^{-2} cm/s measured using Scanning Electrochemical Microscopy.
- * Experimental results & pore histograms are consistent with theory that neglects pore resistance
- * Between 2-3 orders higher than small molecule diffusion permeability of reconstituted cellulose or PES.

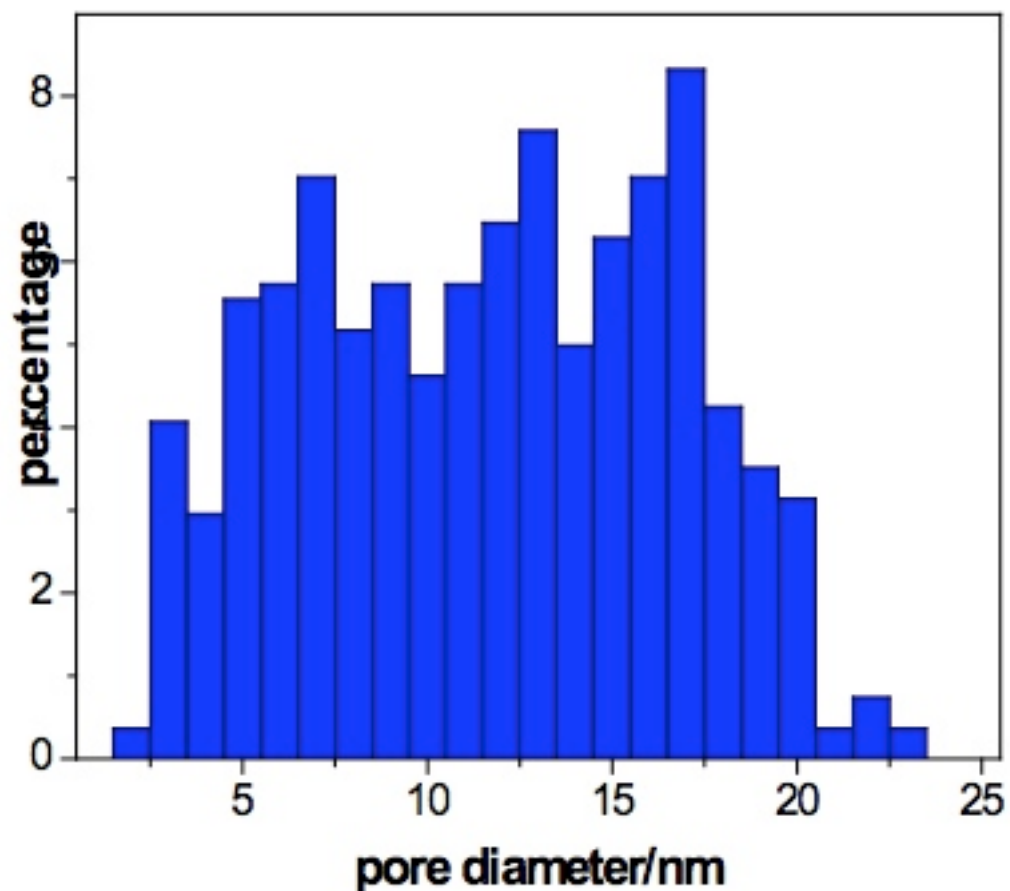
Kim et al. 2008, JACS 130:4230-4231

 NRG 2009

High Diffusive Perm

Shigeru Amemiya

- * Intrinsic permeability of 5.2×10^{-2} cm/s measured using Scanning Electrochemical Microscopy.
- * Experimental results & pore histograms are consistent with theory that neglects pore resistance
- * Between 2-3 orders higher than small molecule diffusion permeability of reconstituted cellulose or PES.



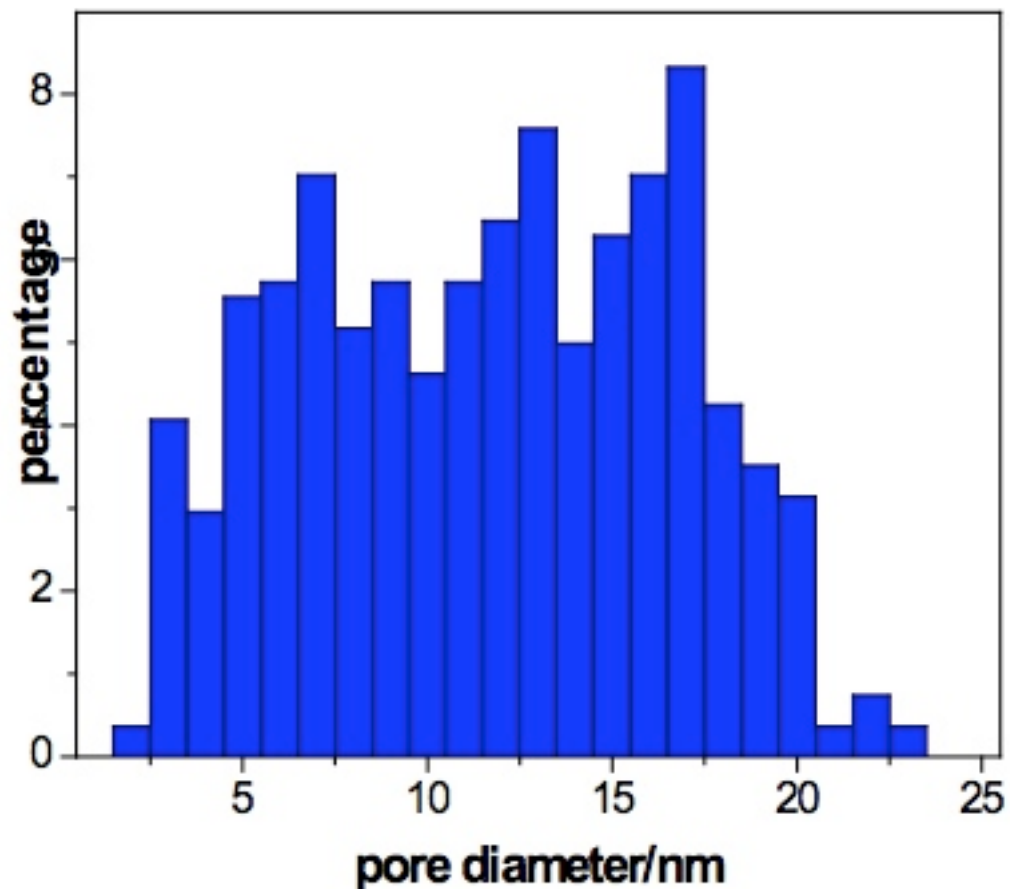
Kim et al. 2008, JACS 130:4230-4231

NRG 2009

High Diffusive Perm

Shigeru Amemiya

- * Intrinsic permeability of 5.2×10^{-2} cm/s measured using Scanning Electrochemical Microscopy.
- * Experimental results & pore histograms are consistent with theory that neglects pore resistance
- * Between 2-3 orders higher than small molecule diffusion permeability of reconstituted cellulose or PES.

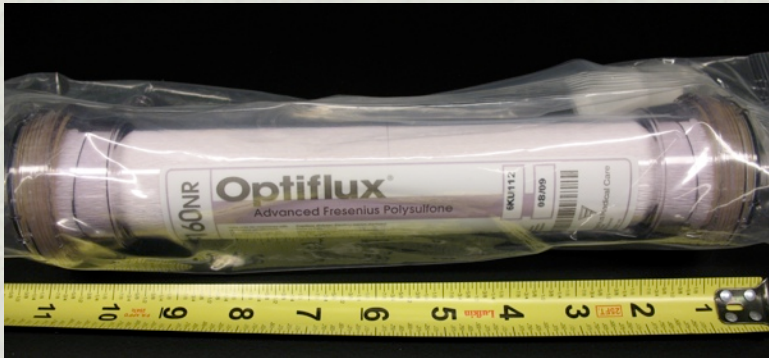


$$k = 2DNr$$

Kim et al. 2008, JACS 130:4230-4231

NRG 2009

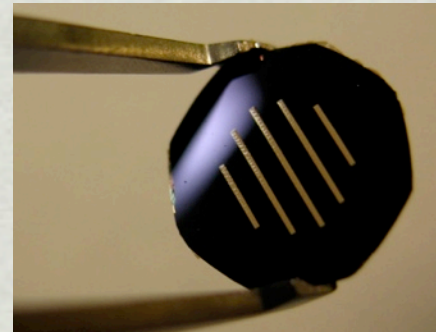
Higher Permeability = Smaller ...



- ✱ *For current devices: active area is 1.8 m² and transmembrane water flow is ~10 ml/min @ 3 psi.*
- ✱ *1000x improved permeability would require only 18 cm². So 10 dime-sized membranes with mostly (~80%) active area could support both transmembrane flow and match dialysis in the same period of time.*

● NRG 2008

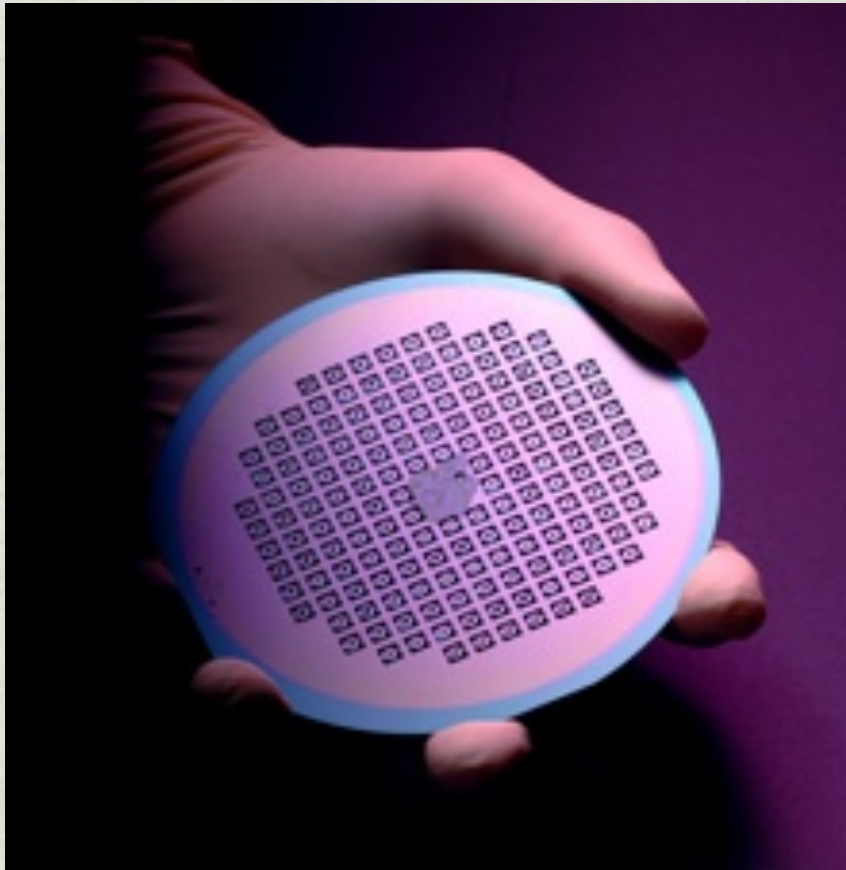
Higher Permeability = Smaller ...



- * *For current devices: active area is 1.8 m² and transmembrane water flow is ~10 ml/min @ 3 psi.*
- * *1000x improved permeability would require only 18 cm². So 10 dime-sized membranes with mostly (~80%) active area could support both transmembrane flow and match dialysis in the same period of time.*

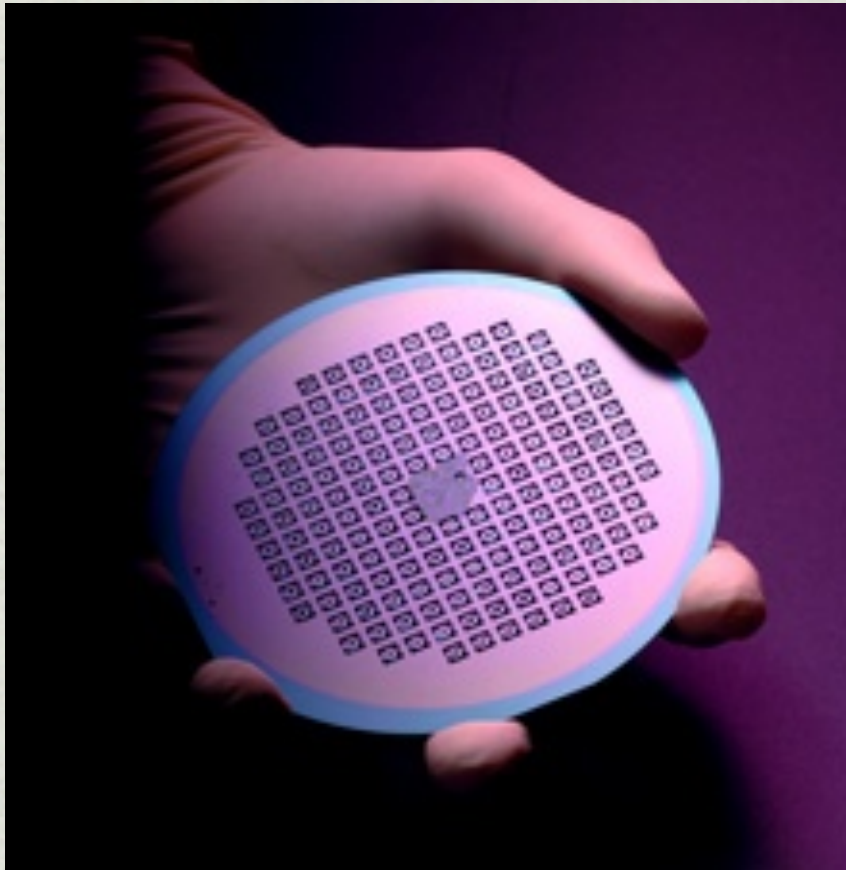
● NRG 2008

... or faster?



- ✱ *Alternatively a 6" wafer of mostly active membrane could achieve the same dialysis in 1/10th of the time.*

... or faster?



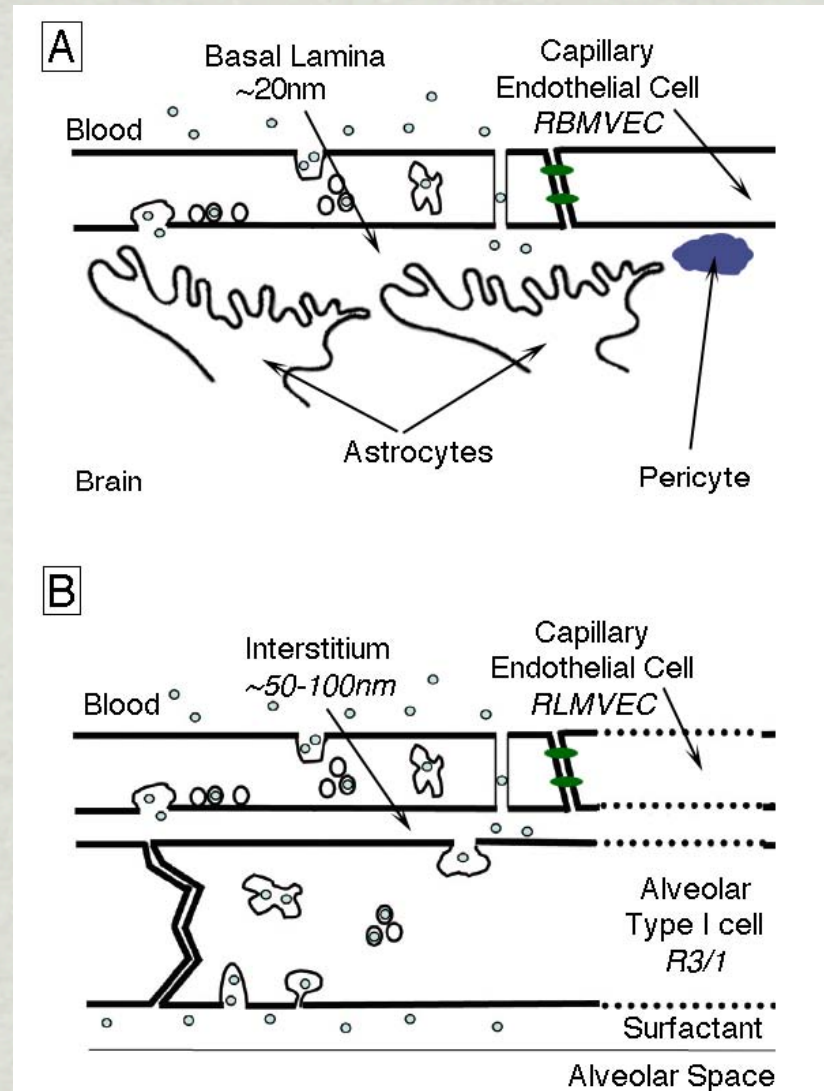
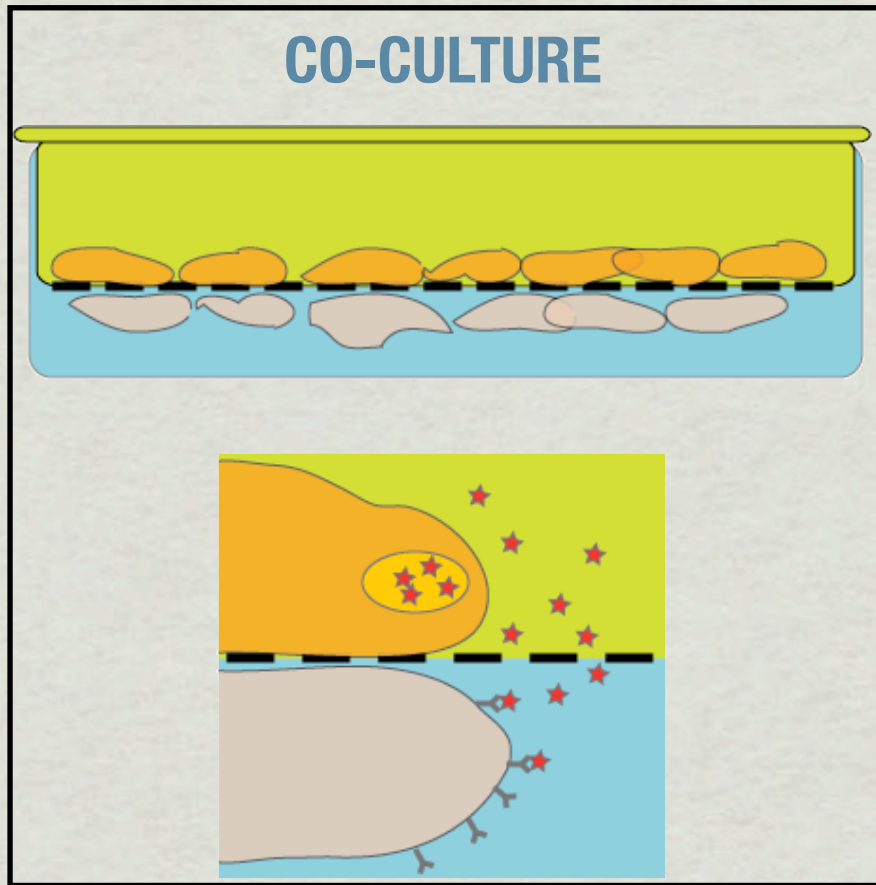
- ✱ *Alternatively a 6" wafer of mostly active membrane could achieve the same dialysis in 1/10th of the time.*

**THE MATERIAL PROVIDES THE
POTENTIAL**

**REALIZING THIS POTENTIAL IS AN
ENGINEERING CHALLENGE**

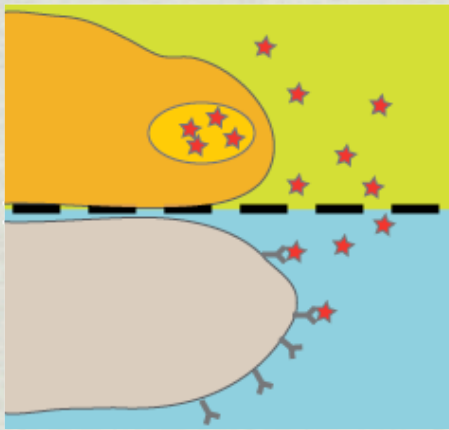
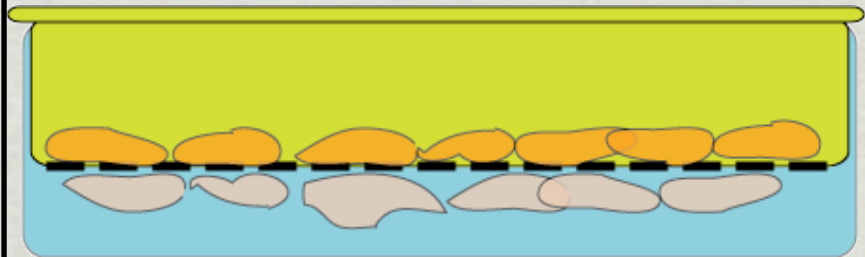
 **NRG 2008**

Cellular Transwell Devices

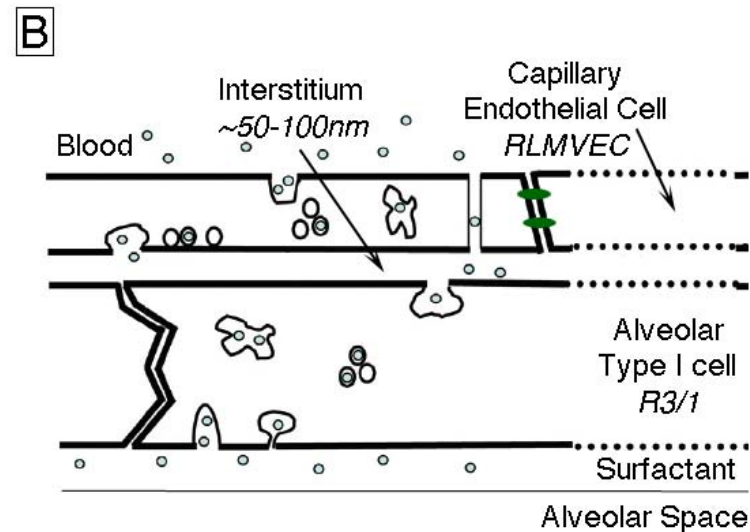
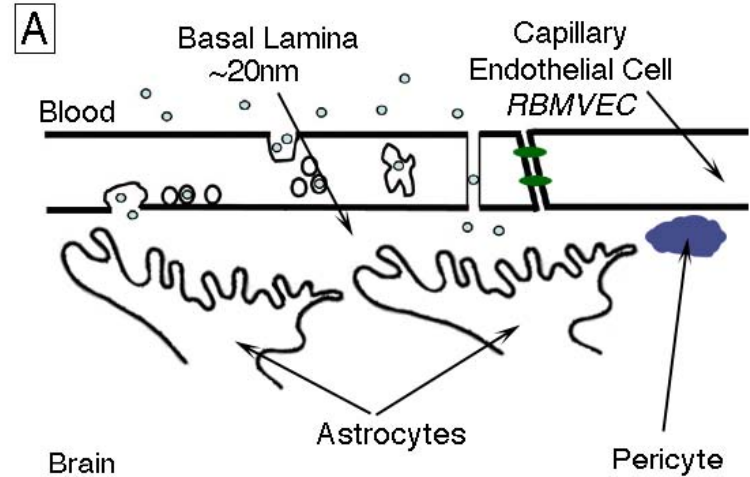
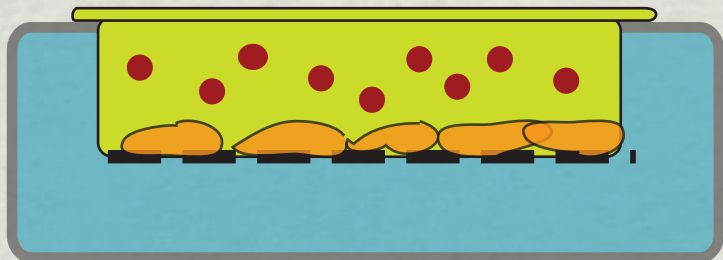


Cellular Transwell Devices

CO-CULTURE



DRUG PERMEATION



Microfluidics

counterflow dialysis



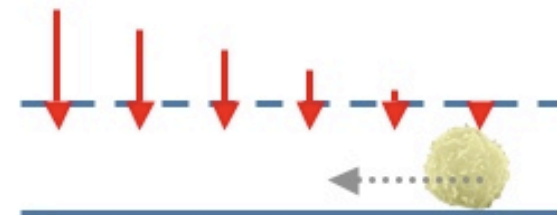
cell/cell Interactions



microbioreactor



chemotaxis



Conclusions

- * Pnc-Si is a new ultrathin nanoporous membrane material. Small membranes can be manufactured on a large scale and incorporated into practical separation devices.
- * Primary application is to small scale separation of biologicals
- * High air and liquid permeabilities w/ demonstrated ability to fractionate proteins, nanoparticles, etc.
- * Viable as a cell-culture substrate. Cell behavior on membranes is normal.
- * Microfluidics, arrayed membranes for screening applications, electrokinetics, and more ...