



РЕНТГЕНОВСКАЯ МИКРОСКОПИЯ И МИКРОТОМОГРАФИЯ - ИСТОРИЯ И РАЗВИТИЕ

Сенин Р. А.

НИЦ «Курчатовский институт»





First steps

Wilhelm Röntgen
Universität Würzburg
Dec. 1895



Michael Pupin
Columbia University/New York
Feb. 1896



"This is of the hand of a gentleman resident in New York, who, while on a hunting trip in England a few months ago, was so unfortunate as to discharge his gun into his right hand, no less than forty shot lodging in the palm and fingers. The hand has since healed completely; but the shot remain in it, the doctors being unable to remove them, because unable to determine their exact location. The result is that the hand is almost useless, and often painful." - Cleveland Moffett, McClure's Magazine, April 1896





Röntgen first experiments



after W.C. Röntgen

Über eine neue art von Strahlen.

Phys.-Med. Ges., Würzburg, **137**, p. 41, (1895)

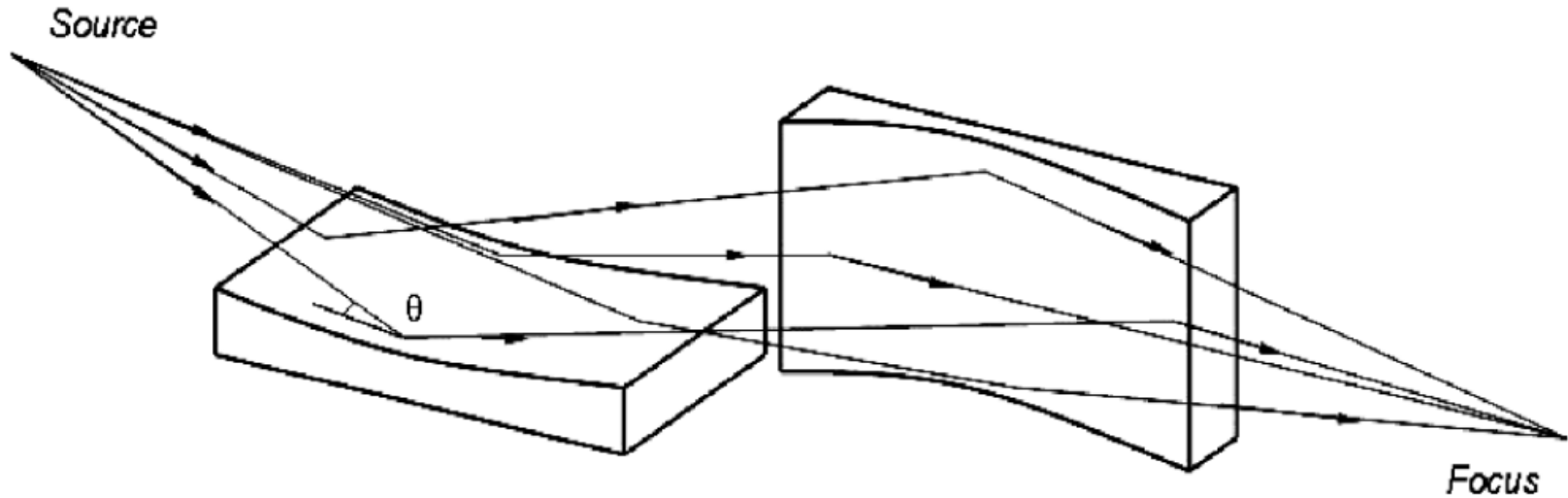
English translation in Nature **53**, p. 274

- **Photographic plates and film are "susceptible to x-rays", providing valuable means of recording the effects..."**
- **"... The refractive index.... cannot be more than 1.05 at most.... X-rays cannot be concentrated by lenses..."**
- **"... Detection of interference phenomena has been tried without success, perhaps only because of their feeble intensity..."**





Kirkpatrick & Baez microscope



Kirkpatrick P and Baez A V 1948
Journal of the Optical Society of America V 38 pp.766–774





X-ray optics milestones

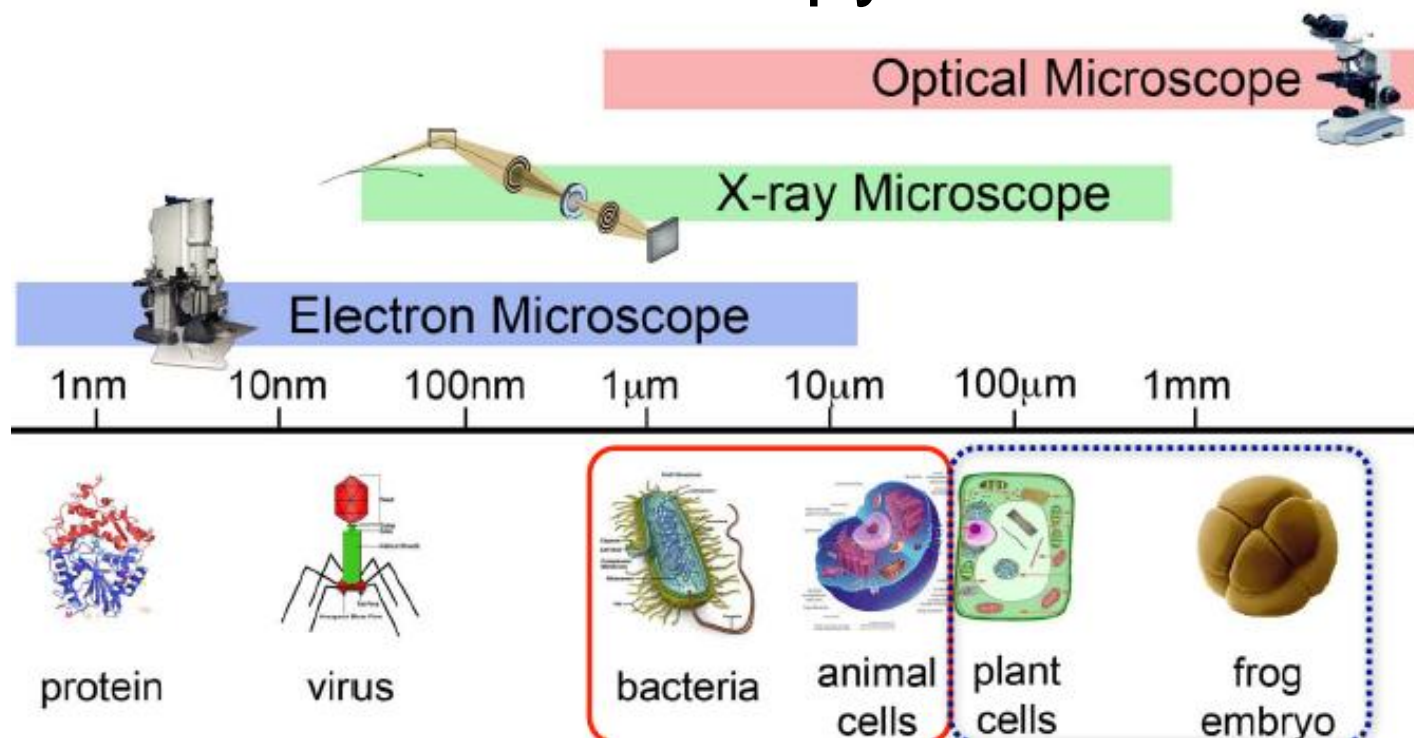
- 1896 X-ray absorption (Röntgen – filters)
- 1912 Bragg diffraction (Laue - monochromators)
- 1913 Radiography (Goby – imaging)
- 1922 Specular reflection (Compton - mirrors)
- 1948 X-ray curved mirrors (Kirkpatrick and Baez)
- 1960 X-ray Fresnel zone plates (Baez)
- 1972 X-ray multilayers (Spiller, Barbee, Underwood)
- 1987 Bragg-Fresnel optics (Aristov)
- 1994 Capillaries (Bilderback)
- 1996 Compound Refractive Lenses (Suehiro, Snigirev)





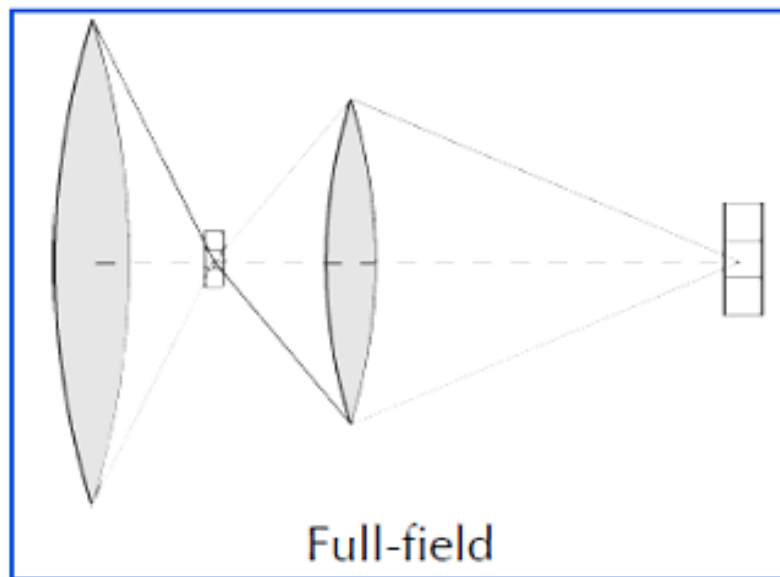
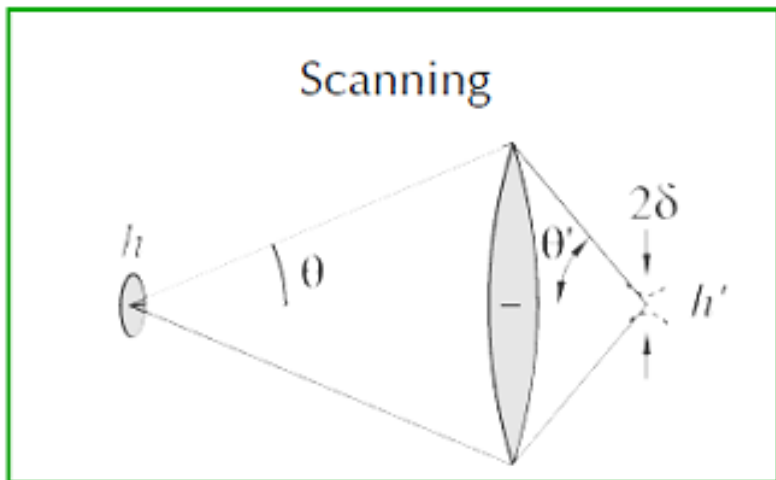
X-ray microscopy:

Bridging the gap between electron and light microscopy





Scanning or full-field?

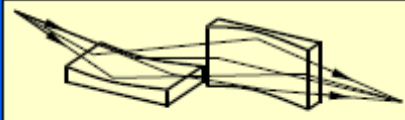

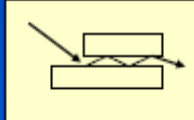
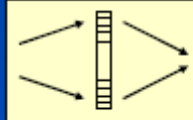
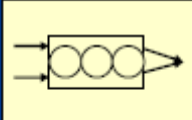


Hard x-rays or soft?





Focusing Optics for Hard X-rays ($E > 6$ keV)

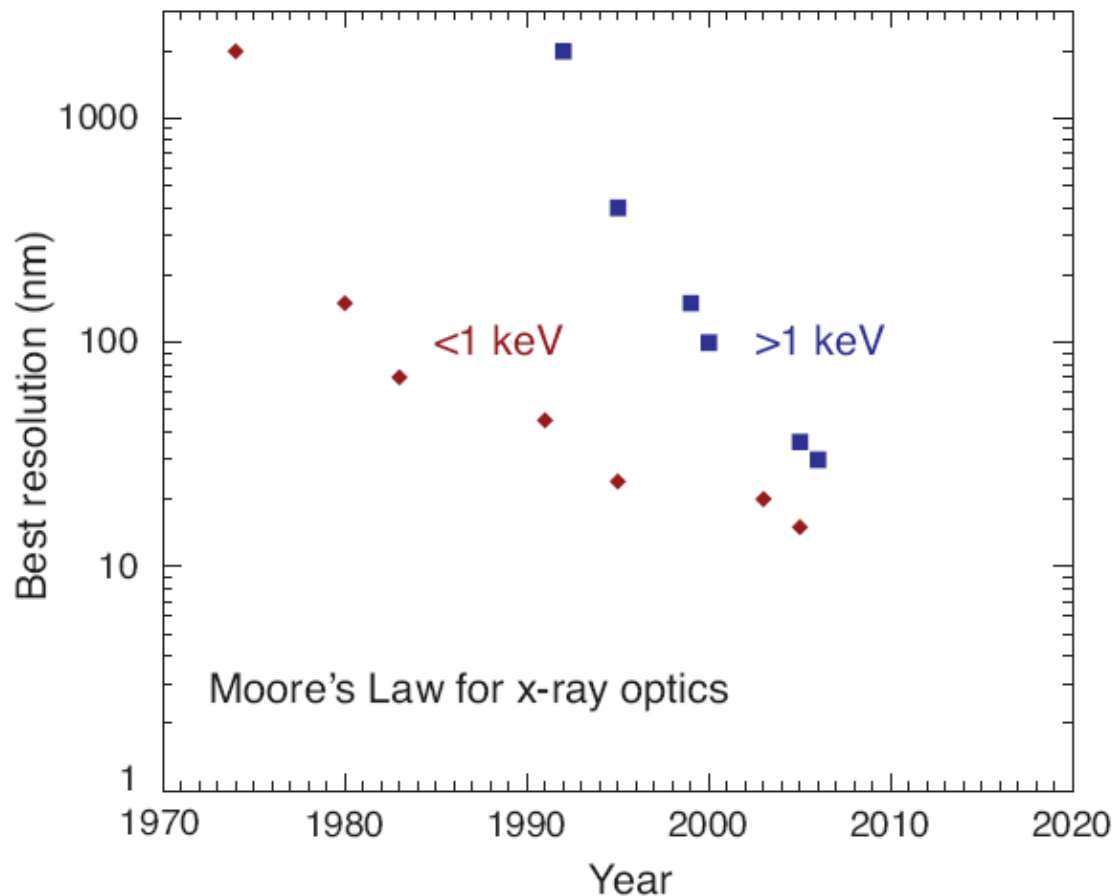
	<i>reflective</i>		<i>diffractive</i>	<i>refractive</i>		
	Kirkpatrick Baez systems		Capillaries	Waveguides	Fresnel Zone plates	Refractive lenses
	mirrors Kirkpatrick Baez, 1948	multilayers Underwood Barbee, 1986	Kreger 1948	Feng et al 1993	Baez 1952	Snigirev et al, 1996
						
Energy	< 30 keV	< 80keV	< 20keV	< 20keV	< 30 keV (80)	<1 MeV
Bandwidth $\Delta E/E$	w. b.	10^{-2}	w.b.	10^{-3}	10^{-3}	10^{-3}
resolution	25 nm @15keV Mimura 2006	41x45nm² @24keV Hignette 2006	50 nm Bilderback 1994	40x25 nm² Salditt 2004	30 nm @20 keV Kang, 2006 17 nm , 2007	50 nm@20keV Schroer, 2004 150nm@50keV Snigirev,2006





Moore's law for x-ray optics

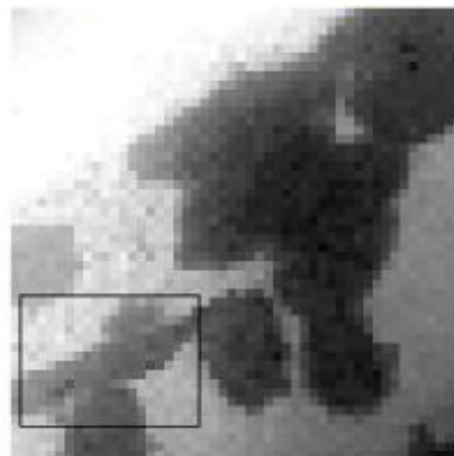
X-ray optics: best resolution





Radiation degradation

- X-rays are ionizing radiation. The dose per high resolution image is about 100,000 times what is required to kill a person
- Makes it hard to view living cells!



Experiment by V. Oehler, J. Fu, S. Williams, and C. Jacobsen, Stony Brook using specimen holder developed by Jerry Pine and John Gilbert, CalTech

10 μm
 $6.0 \cdot 10^2$ Gray, ET=2 min.



5 μm
 $1.2 \cdot 10^5$ Gray, ET=9.5 min.



5 μm
 $2.4 \cdot 10^5$ Gray, ET=17 min.



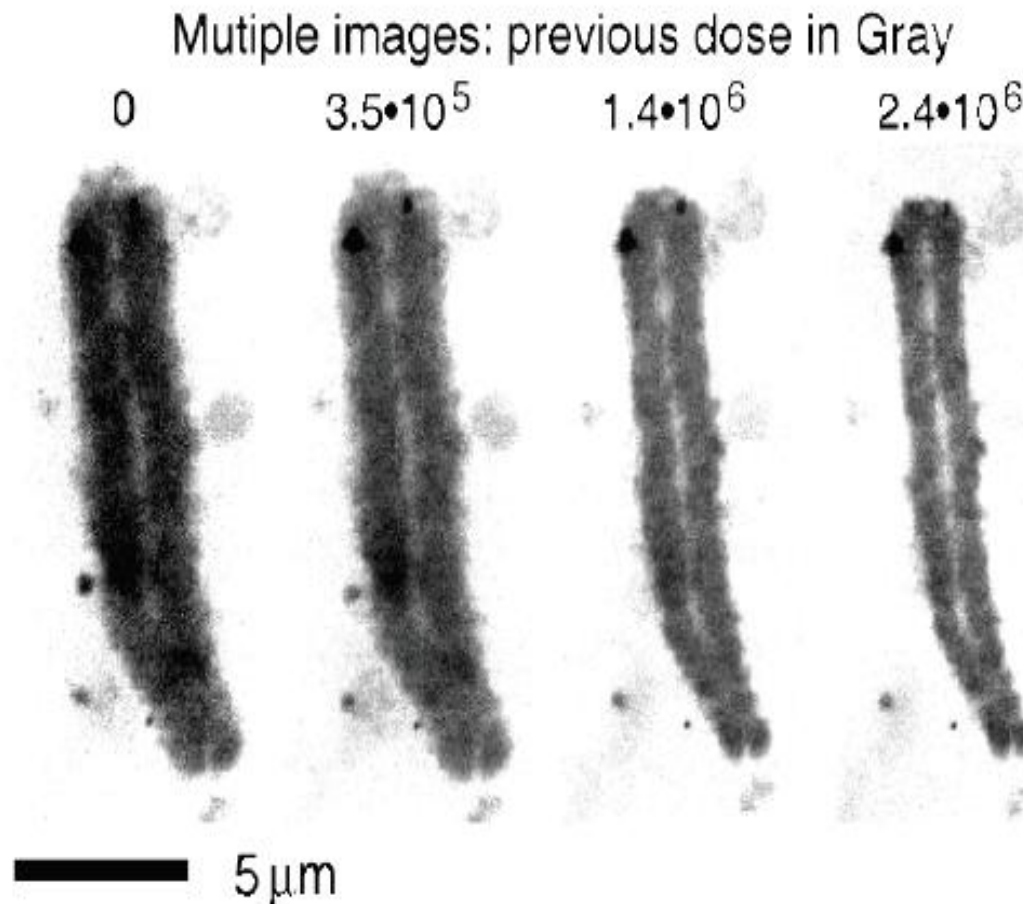
5 μm
 $3.7 \cdot 10^5$ Gray, ET=24.5 min.





Radiation degradation

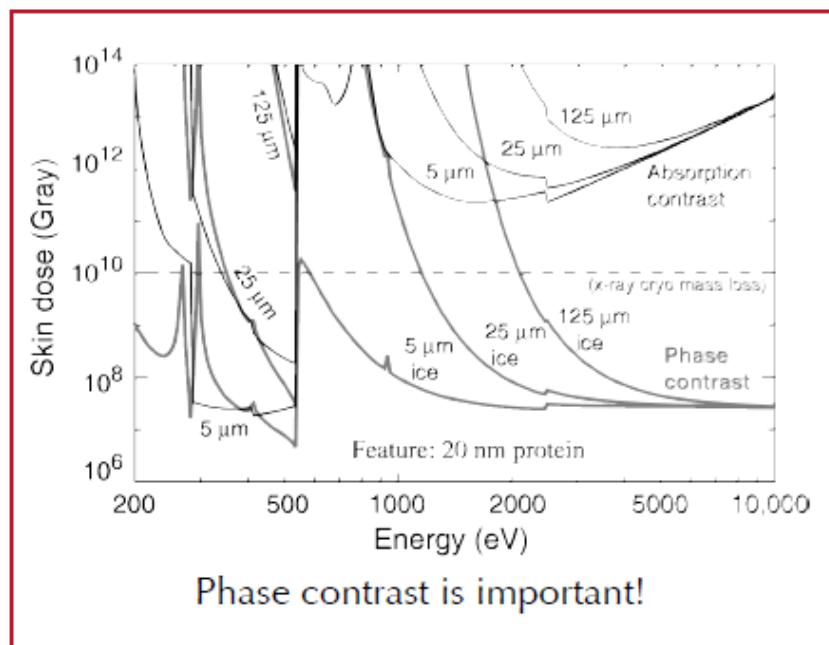
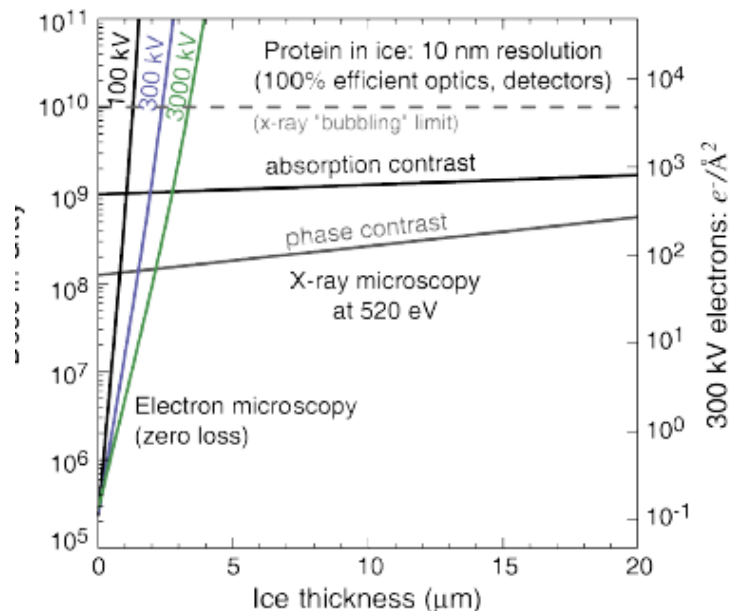
- Chromosomes are among the most sensitive specimens.
- *V. faba* chromosomes fixed in 2% glutaraldehyde. S. Williams et al., J. Microscopy **170**, 155 (1993)
- Repeated imaging of one chromosome shows mass loss, shrinkage





Electrons vs. soft X-rays

X-rays: better for thicker specimens. Sayre *et al.*, *Science* **196**, 1339 (1977); Schmahl & Rudolph in *X-ray Microscopy: Instrumentation and Biological Applications* (Springer, 1987)

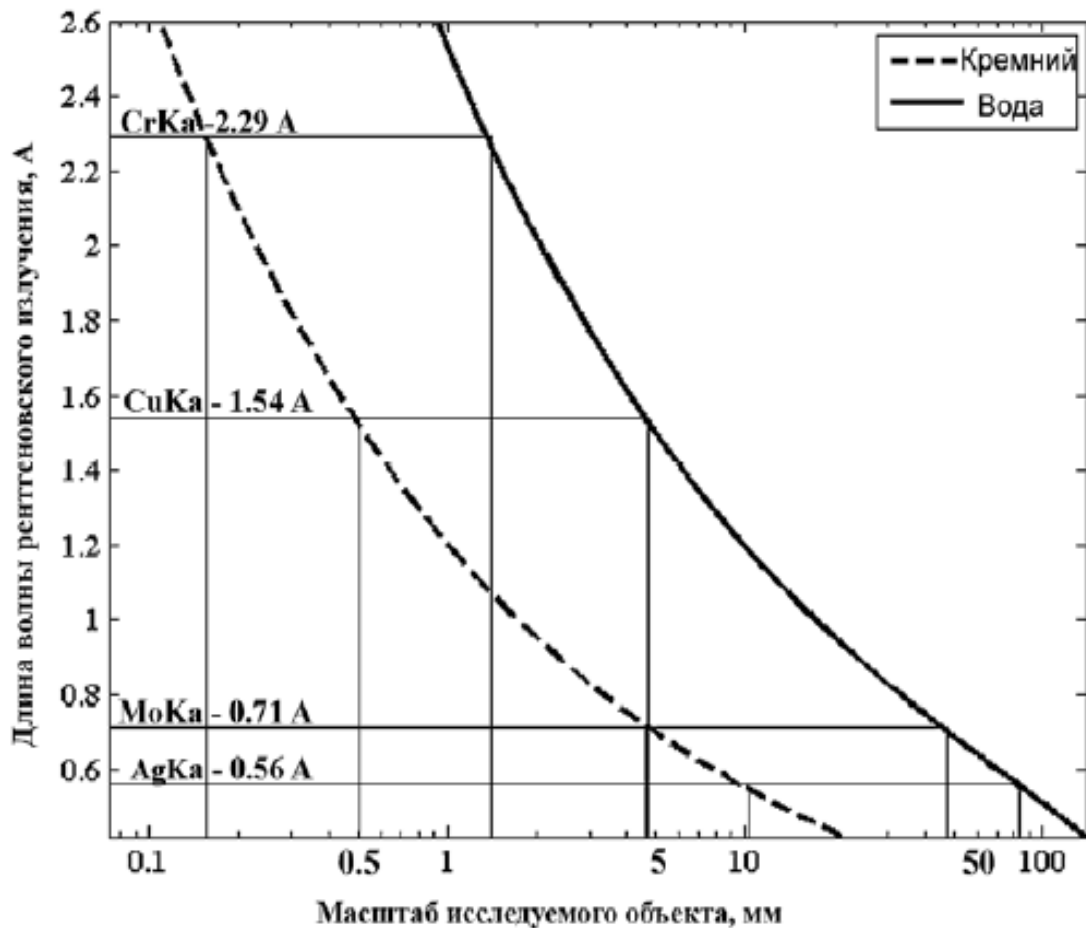


These plots: based on Jacobsen, Medenwaldt, and Williams, in *X-ray Microscopy & Spectromicroscopy* (Springer, 1998)





Hard X-ray: Choosing optimal wavelength

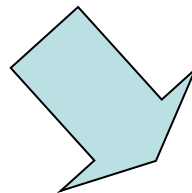


В.Е.Асадчиков, А.В.Бузмаков, Д.А.Золотов и др. Кристаллография, 2010, том 55, № 1, с. 167–176



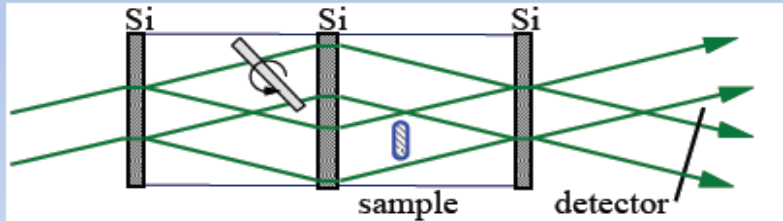


Imaging phase





Phase sensitive imaging

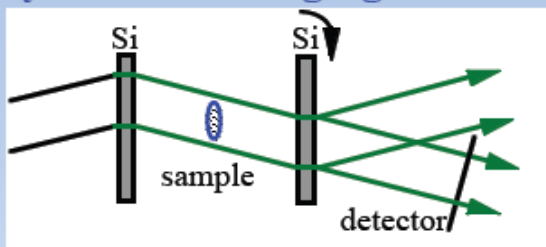


$$\varphi$$

Interference with reference beam

Ando & Hosoya, 1972
Bonse; Momose

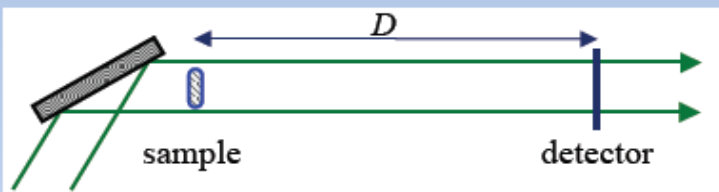
Analyser based imaging



$$\frac{d\varphi}{dx}$$

Analyser as angular filter

Zaumseil, 1980
Belyaevskaia & Ingal,
Chapman



$$\frac{d^2\varphi}{d^2x}$$

Fresnel diffraction

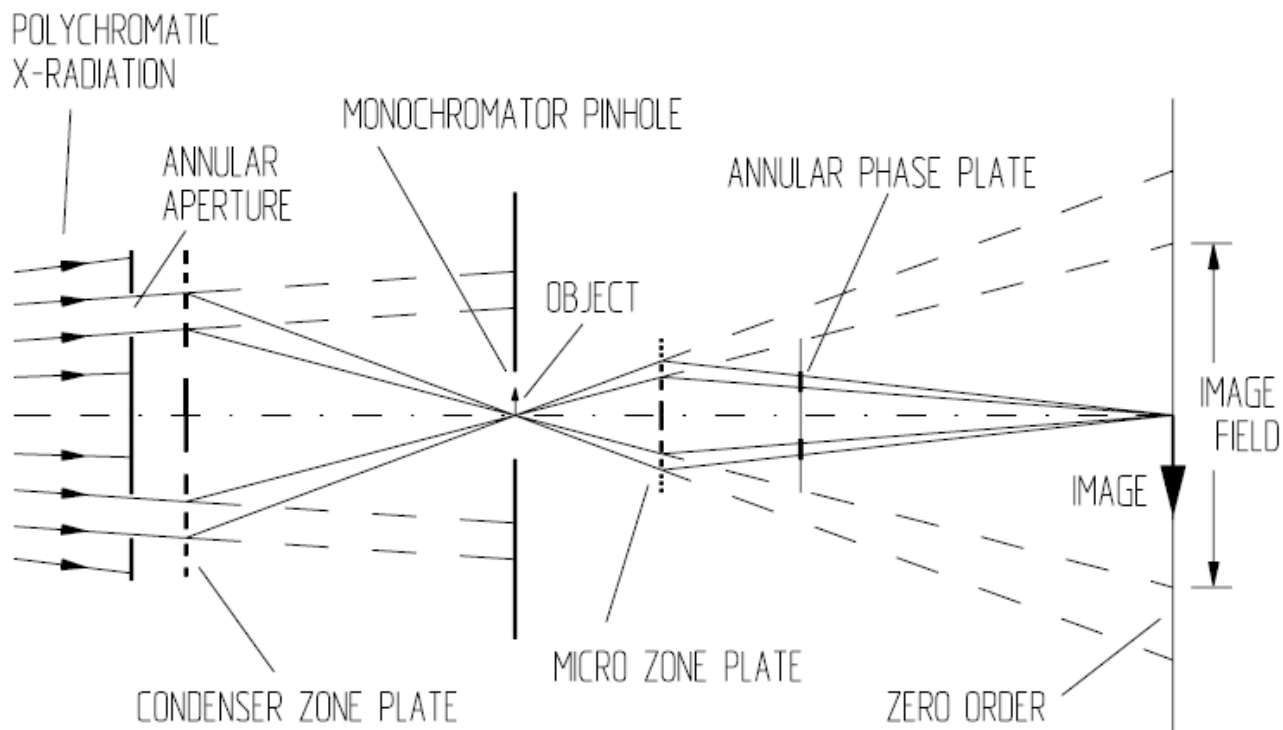
Hartman (1994)
Cloetens; Snigirev; Wilkins

By the courtesy P.Cloetens, HERCULES lectures, ESRF





Zernike phase contrast



From Kirz, Jacobsen, and Howells, Q. Rev. Biophys. 28, 33-130 (1995)

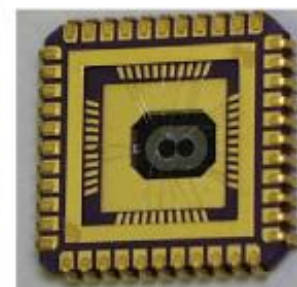
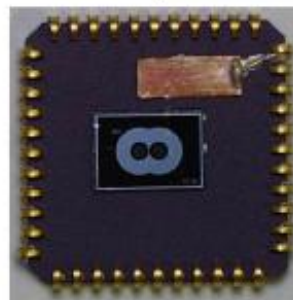
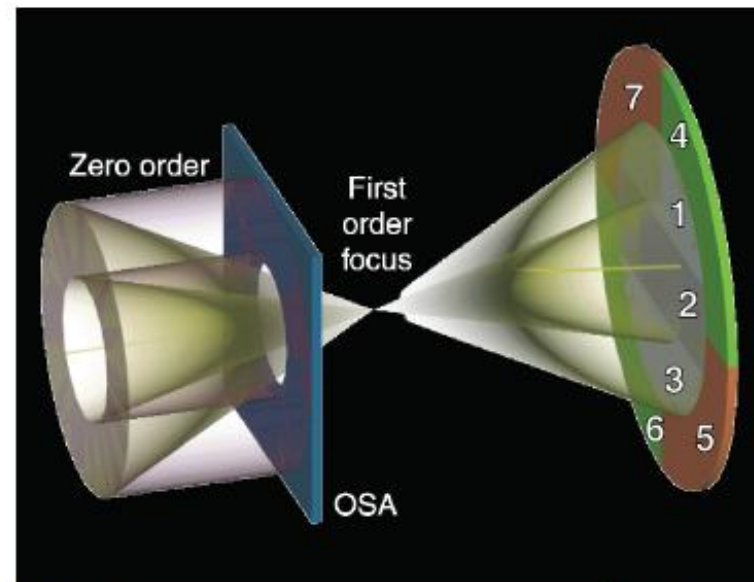




Fast phase contrast detector

M. Feser, B. Hornberger, C. Jacobsen (Stony Brook); P. Rehak, G. de Geronimo (BNL Instrumentation); L. Strüder, P. Holl (MPI München)

- Silicon drift detector
- Simultaneous recording of bright field, dark field, differential contrast at msec pixel times
- No significant upper limit to signal rate. Acceptable dark noise (~ 8 photons/msec equivalent; room temperature)
- High quantum efficiency ($>90\%$)



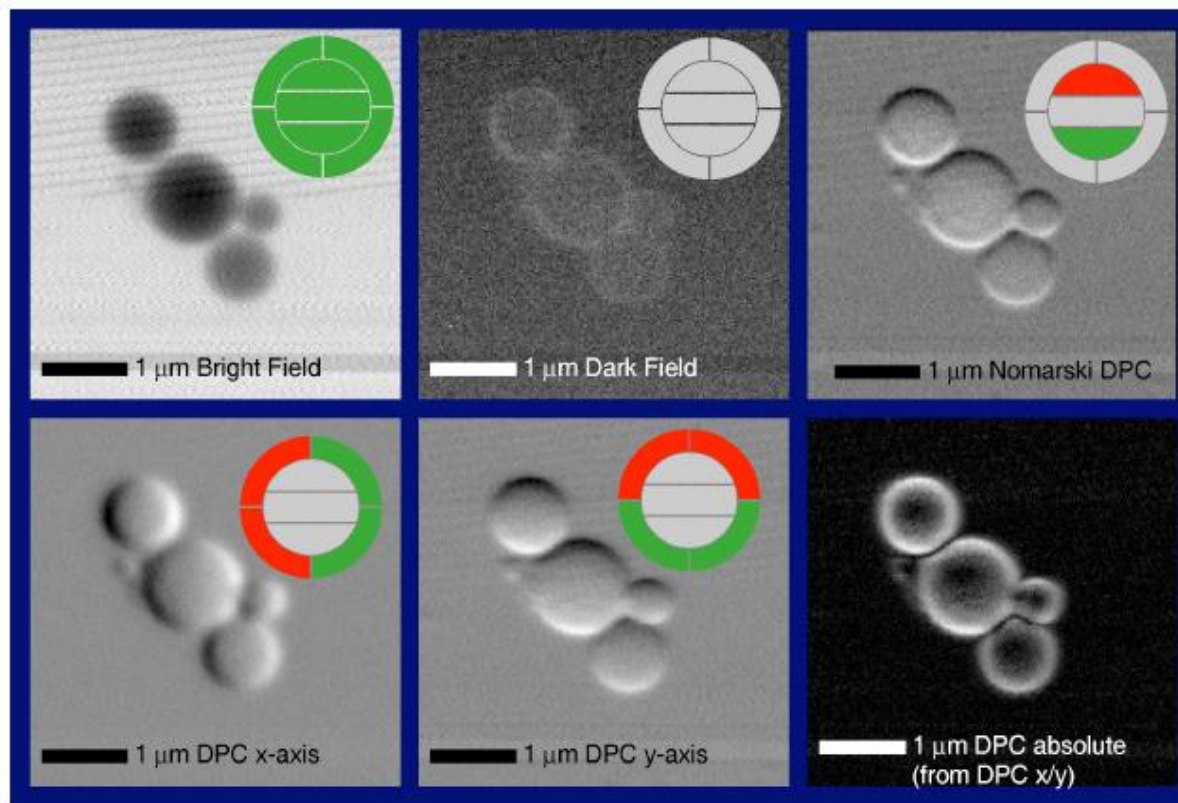
Assembly: 40 mm across
Active area: 600 μm





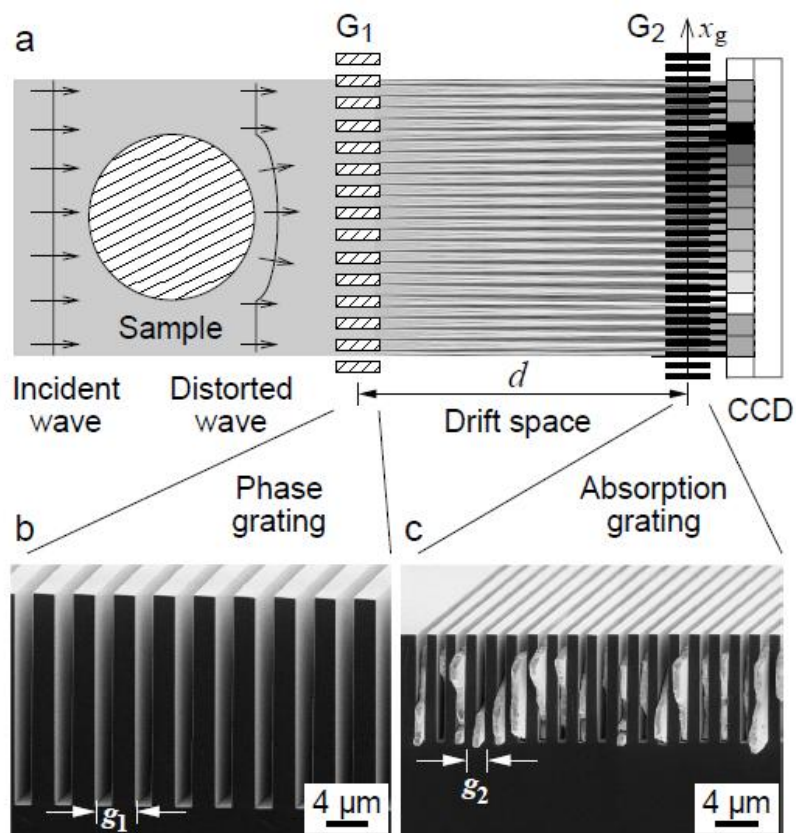
Simultaneous Availability Of Contrast Modes

- Silica spheres 1 μm diameter or less
- Differential phase contrast filters out intensity fluctuations of the source!



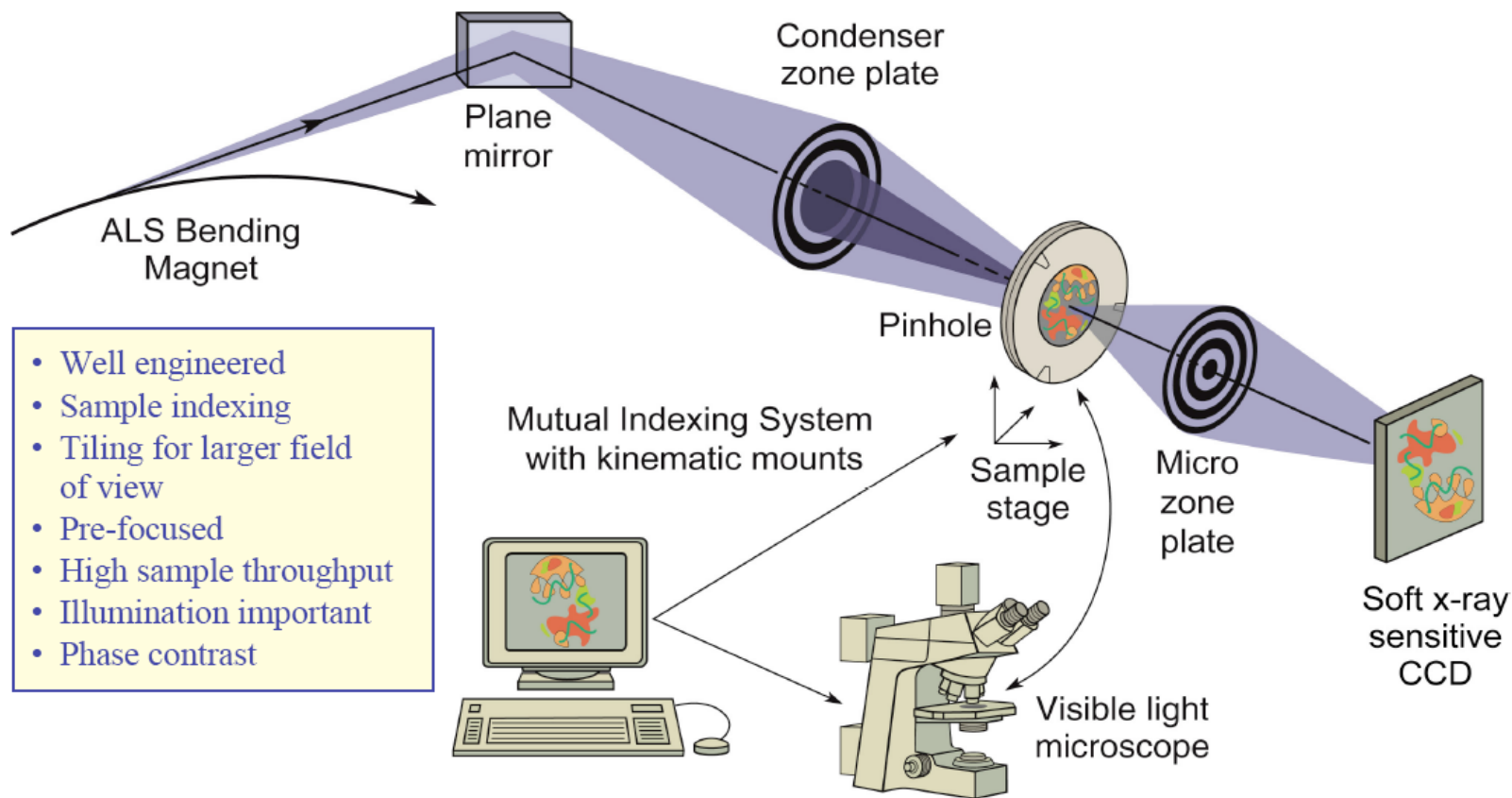


Talbot interferometer



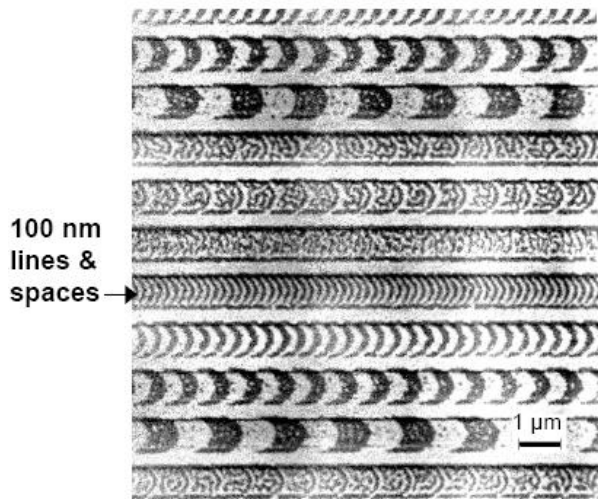


High Resolution Zone-Plate Microscope XM-1 at the ALS





Magnetic Recording Materials

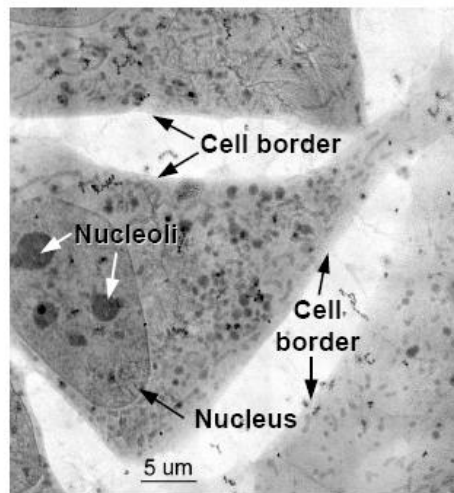


Fe L₃ @ 707.5 eV

FeTbCo Multilayer
with Al Capping Layer

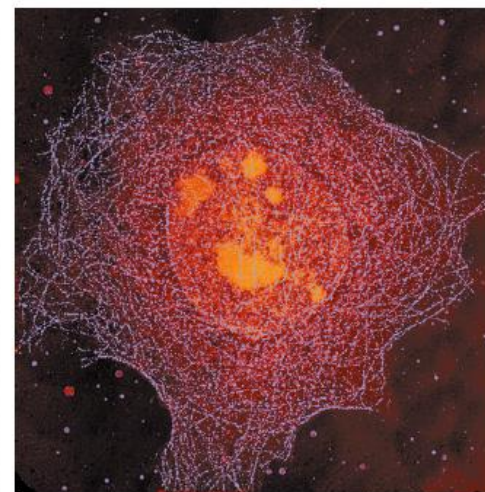
Courtesy of P. Fischer (Max Planck)
and G. Denbeaux (CXRO/LBNL)

Cryo Microscopy for the Life Sciences



Cryo X-Ray Microscopy
of 3T3 Fibroblast Cells

Courtesy of C. Larabell (UCSF)
and W. Meyer-Illse (CXRO/LBNL)

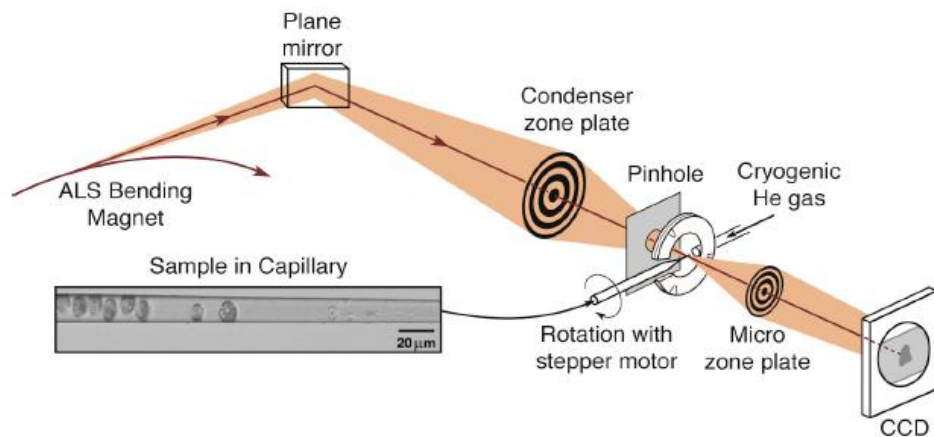


Protein Labeled
Microtubule Network

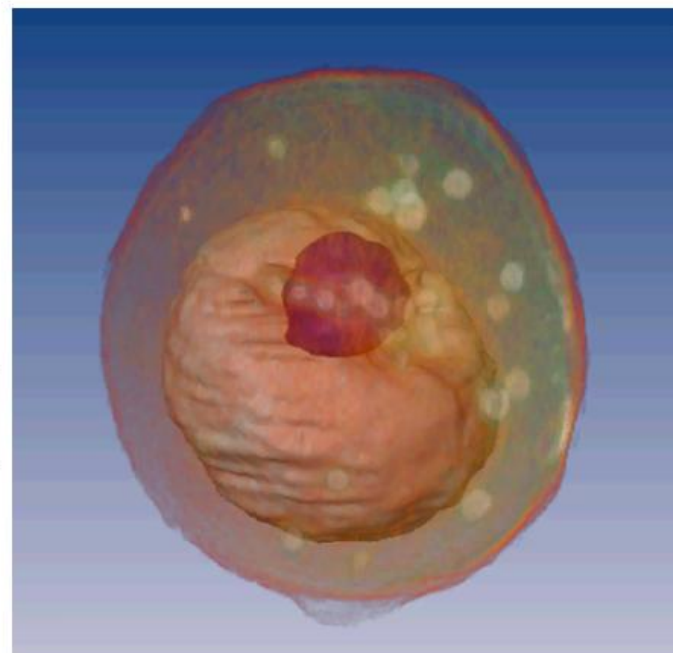




Nanotomography of Cryogenic Fixed Cells



Soft X-Ray Nanotomography of a Yeast Cell



$\lambda = 2.5 \text{ nm}$

C. Larabell and M. LeGros,
Molec. Bio. Cell 15, 957 (2004)

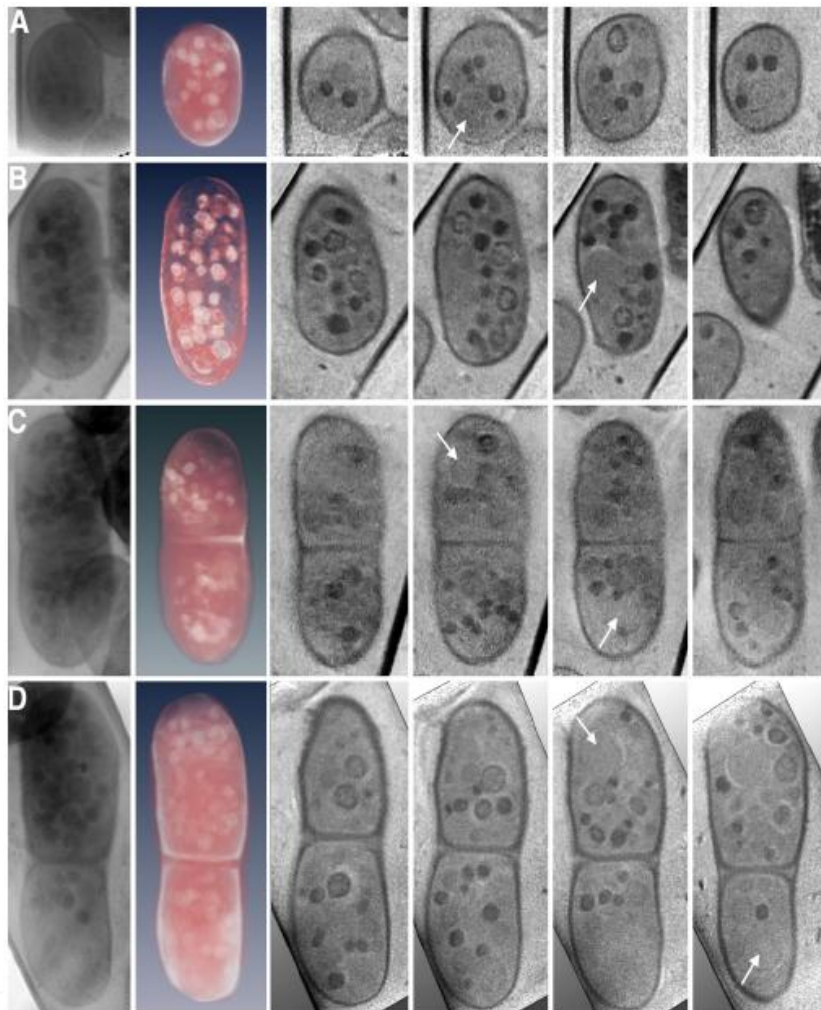


NCXT





Schizosaccharomyces pombe



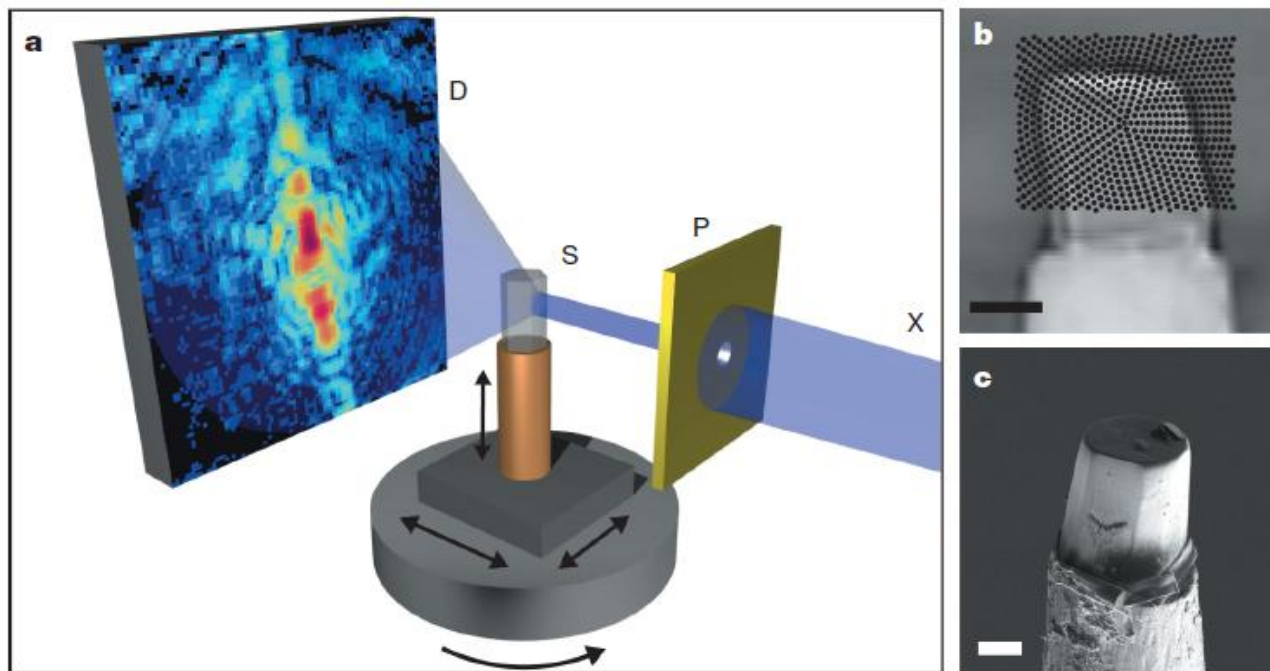
- Newborn cell
- Adult cell
- Dividing cell (early)
- Dividing cell (late)

*W. Gu, L. D. Etkin, M. A. Le Gros,
and C. A. Larabell. (2007)
Differentiation. 75:529-535*





Ptychographic X-ray computed tomography at the nanoscale



Dierolf M, Menzel A, Thibault P, et al. Nature, Vol 467, 2010, pp 436-440.

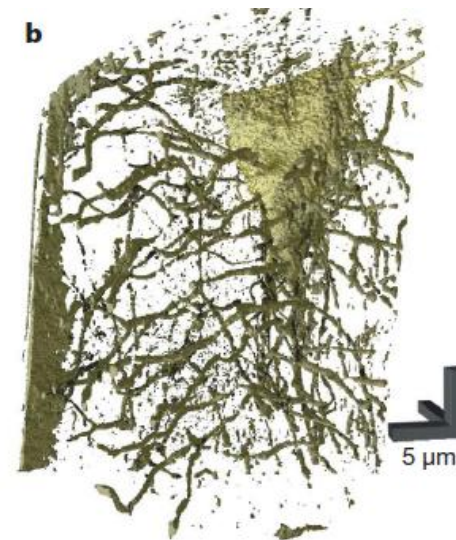
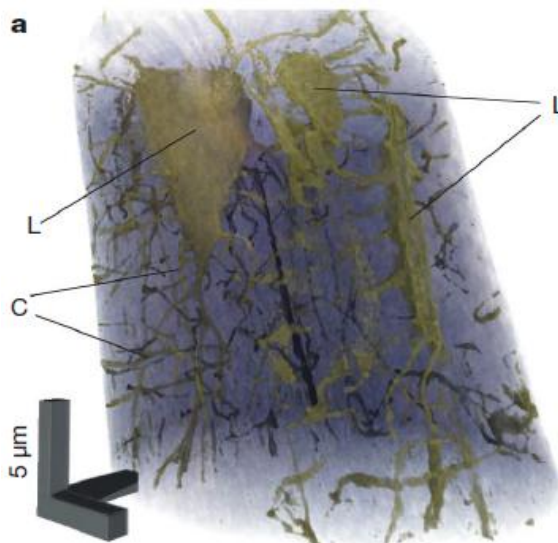
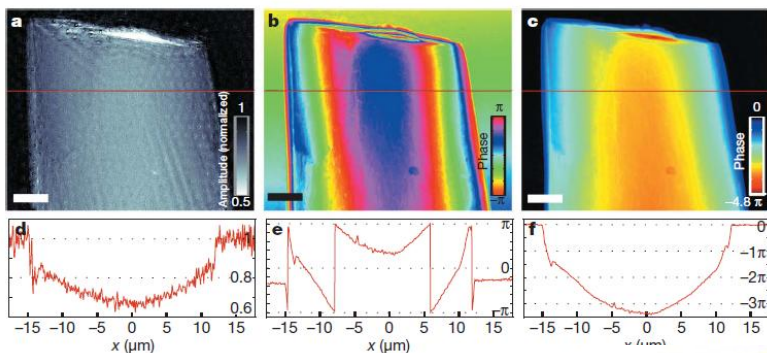




Ptychographic X-ray computed tomography

Spatial resolution - 150 nm

Dose: ~2MGy
($2 \cdot 10^{11}$ Photons total)

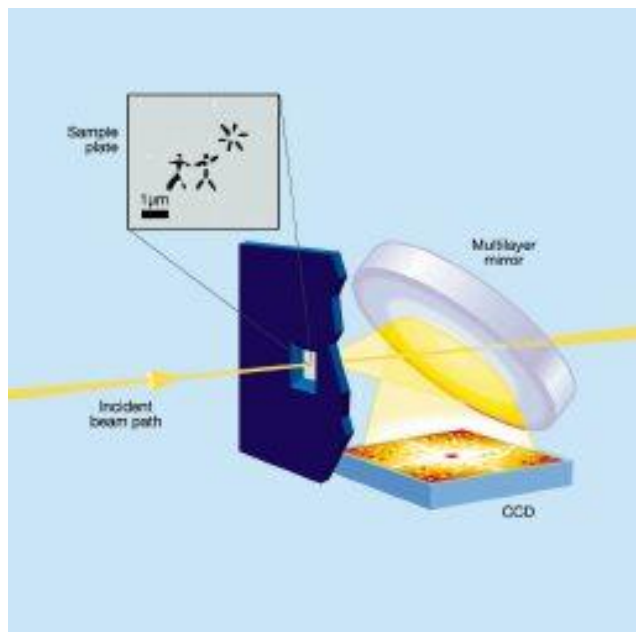


Dierolf M, Menzel A, Thibault P, et al. Nature, Vol 467, 2010, pp 436-440.



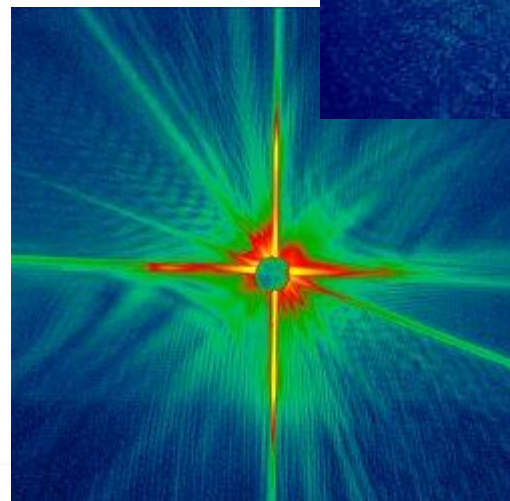
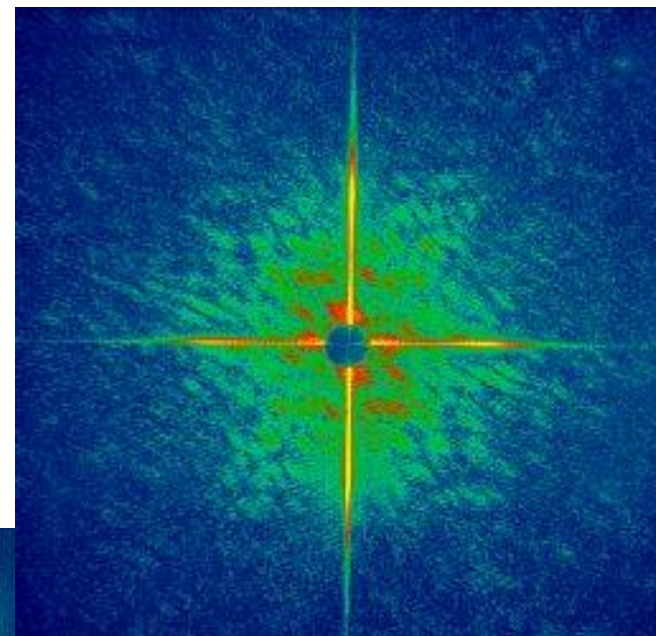


A single shot image



A nanoscale object can be imaged by a single femtosecond FEL pulse before the sample explodes

A coherent diffraction pattern recorded from a single 25 fs pulse

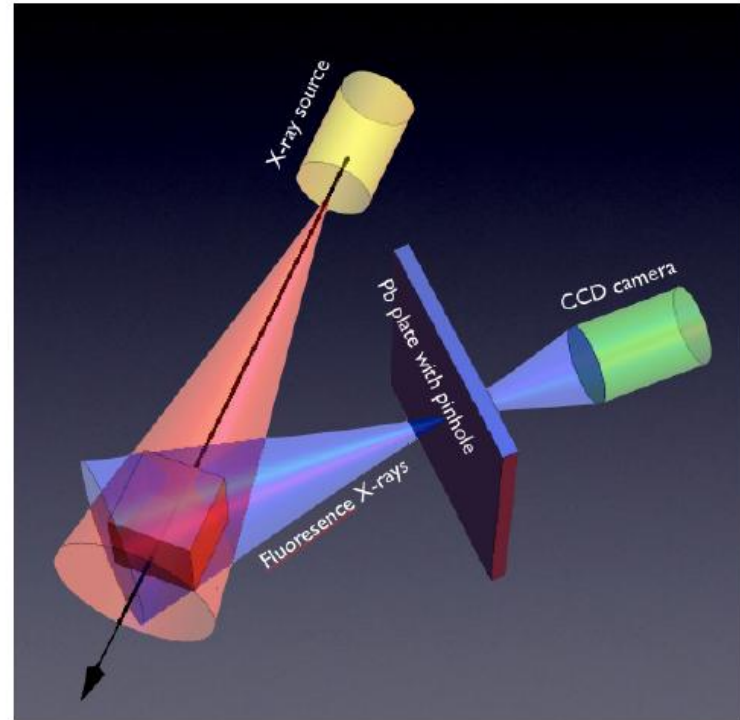
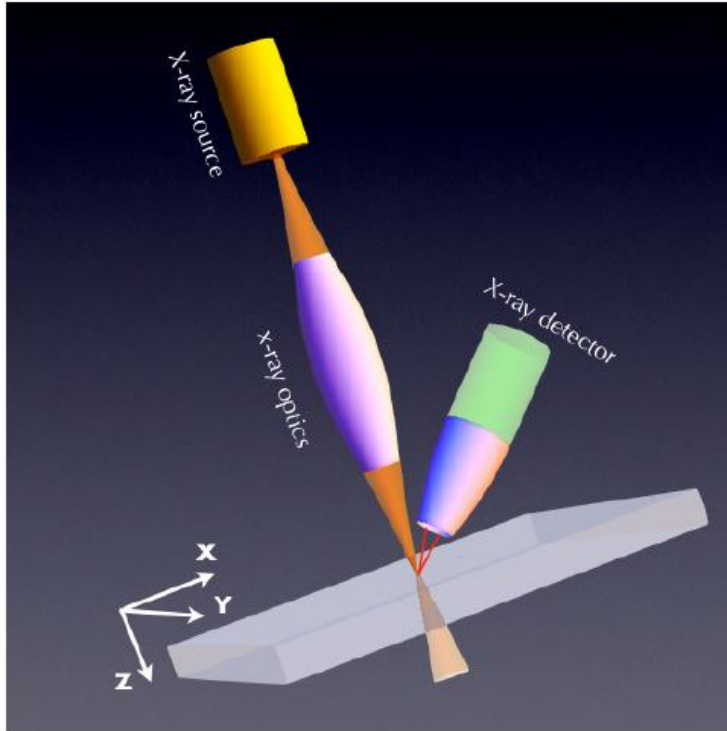


Diffraction pattern from a subsequent pulse showing that the sample was destroyed after recording the image





X-ray fluorescence microtomography



X. Liu, P. Bruyndonckx, and A. Sasov, Proc. SPIE 7804, 78041A-1 (2010).



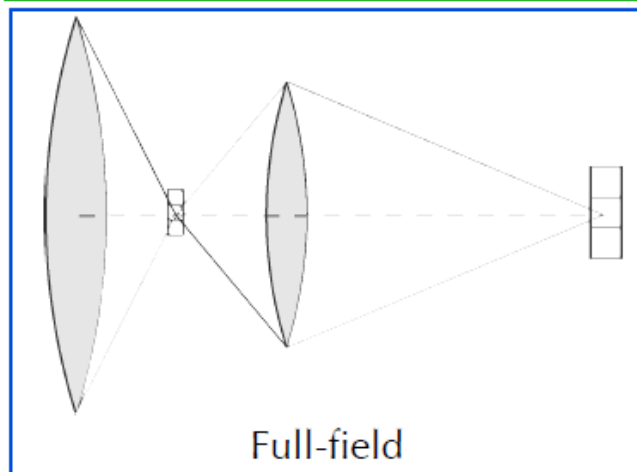
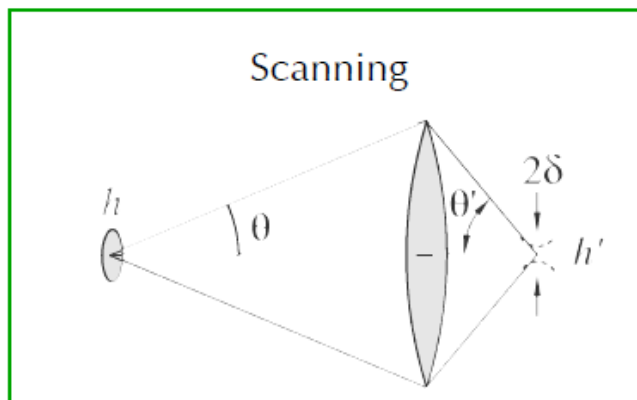


Thank you for your attention!





Scanning or full-field?



Hard x-rays or soft?

