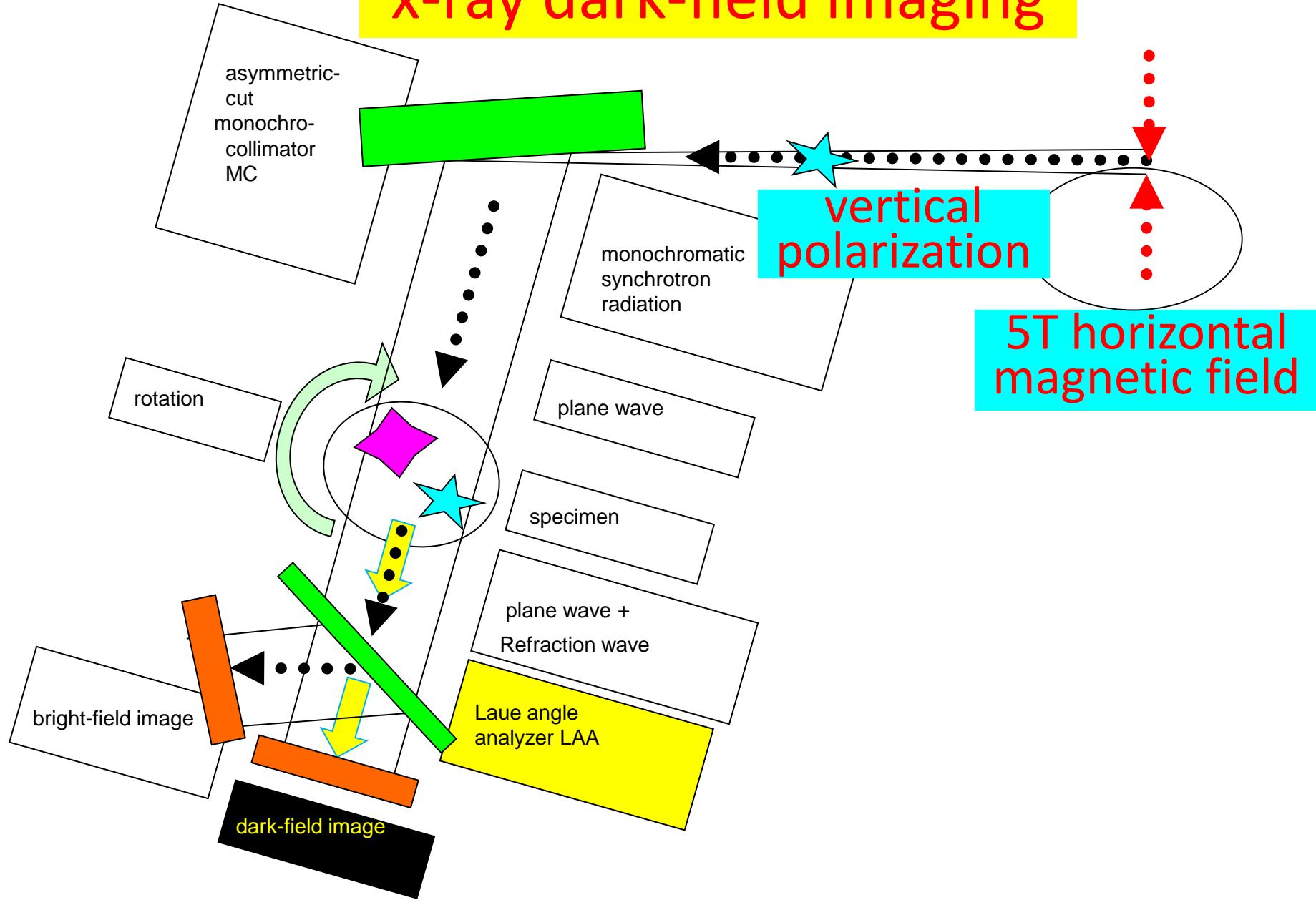


# X-ray Dark-Field Imaging : Recent Developments towards Clinical and Pathological Applications

M. Ando



# x-ray dark-field imaging



# Collaborators

KOREA: KIM Jong-Ki: CUD, Kim Hongtae: CUD, Lim Jae-Hong: PLS,

CHINA: JIANG Xiaoming : BSRF, LI Gang: BSRF,

USA: GUPTA Rajiv: MGH,

AUSTRALIA: MAKSIMENKO Anton : AS

JAPAN: ICHIHARA Shu: NMC, JIN Ge: KIT, MORI Kensaku: NAGOYA U., SUNAGUCHI Naoki: GUNMA U., SUZUKI Yosifumi: KIT, YUASA Tetsuya: YAMAGATA U. , OHURA Norihiko: KYORIN U.

# SOMETHING SPECIAL

- crystal based refraction contrast using a Laue angle analyzer LAA
- vertical polarization that makes setting up x-ray optics so easy and comfortable

# Flow of talk

- (1) what we have been developing ~  
X-ray dark-field imaging (XDFI) optics
- (2) what applications are ongoing
- (3) to which direction we want to go
  - (3-1) clinical
  - (3-2) pathological

# XDFI condition

- $I_{FD}(w, H) = (w^2 + \cos^2\pi H(1 + w^2)^2/\Lambda)) / (1 + w^2)$  (1)
- $I_D(w, H) = (w^2 + \sin^2\pi H(1 + w^2)^2/\Lambda)) / (1 + w^2)$  (2)
- available from X-ray dynamical diffraction theory
- w: angular parameter H: thickness of LAA
- $\Lambda = \lambda \cos\Theta_B / |P| |\chi_G|$  : Pendellosung fringe period
- $\Theta_B$ =Bragg angle P:polarization  $\chi_G = (\lambda^2 r_e \pi V) F_G$  : susceptibility  $r_e$  :classical electron diameter V:unit cell volume  $F_G$ :crystal structure factor

- $I_{FD} = \cos^2(\pi H/\Lambda)|_{w=0} = 0$

- $H = (p+1/2)\Lambda : thickness\ of\ LAA$

- $p: integer$

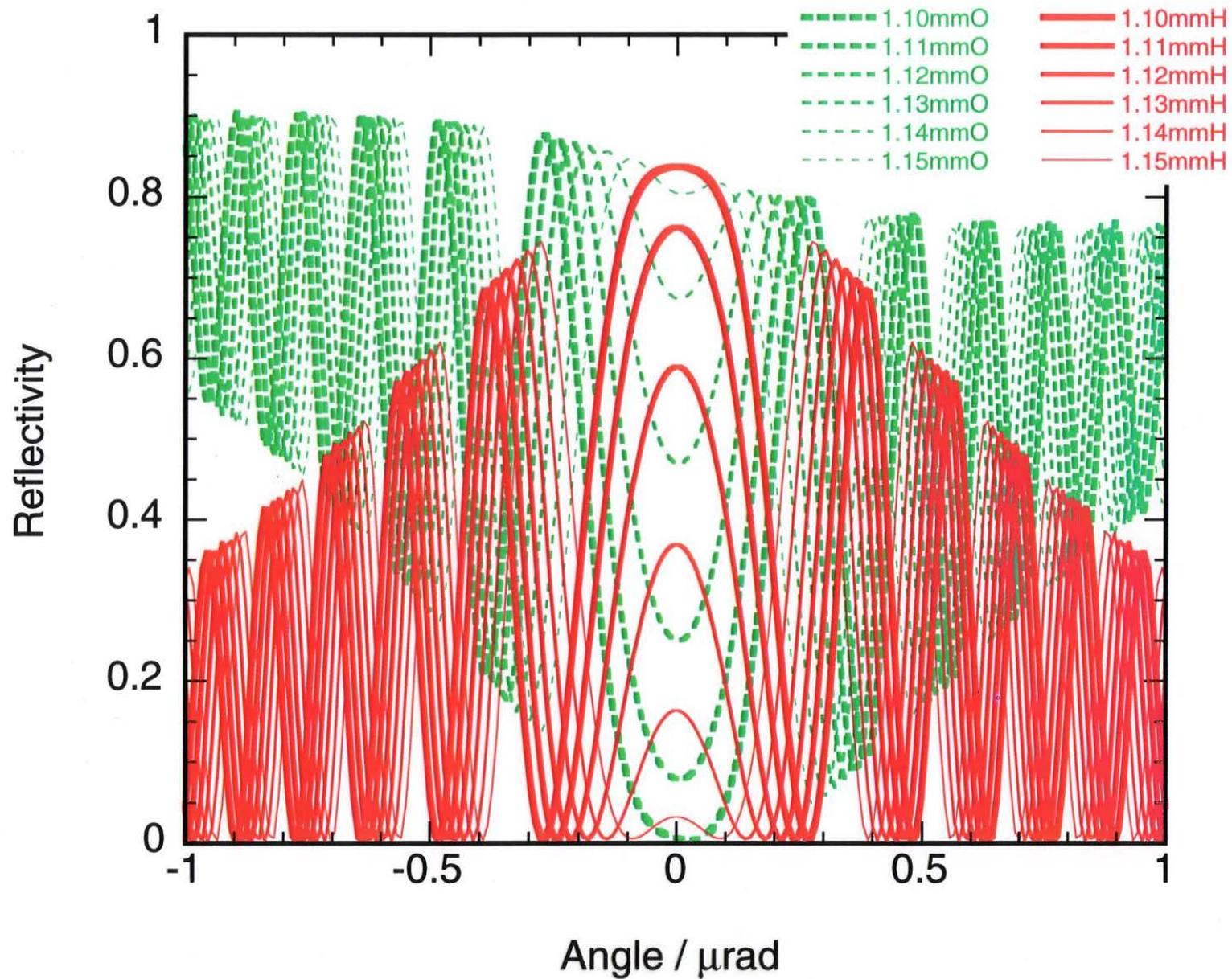
- $\Lambda : Pendellosung\ period\ \sim 100\mu m$

- $I_{FD}(0, H) \Big|_{w=0} = \cos^2 \pi H / \Lambda$  (4)
- $I_D(0, H) \Big|_{w=0} = \sin^2 \pi H / \Lambda$  (5)
- $I_{FD}(0, H) \Big|_{w=0} + I_D(0, H) \Big|_{w=0} = 1$  (6)

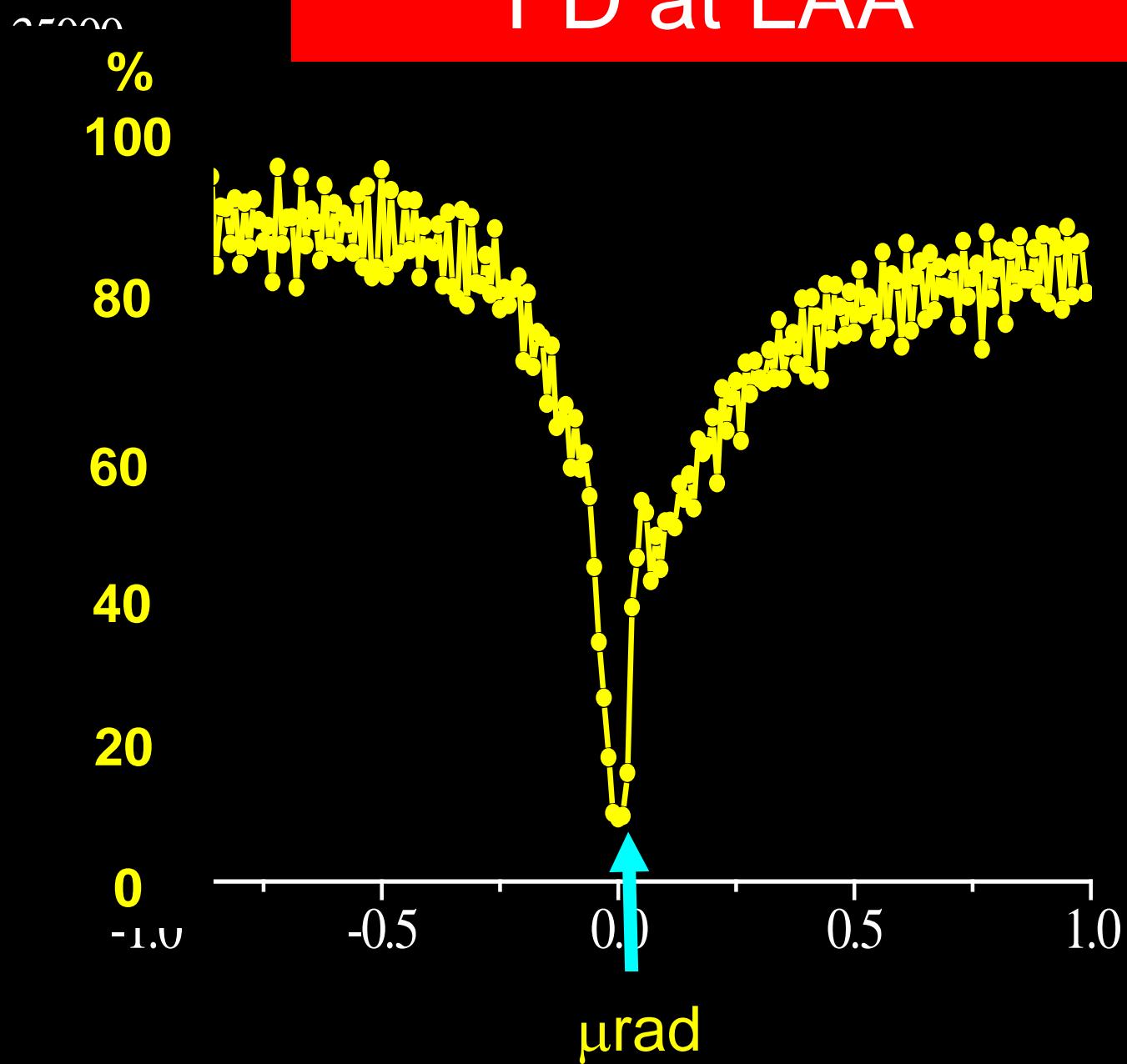
# FD and D profile on Si 440 Laue case at

FD:  
convex  
downward

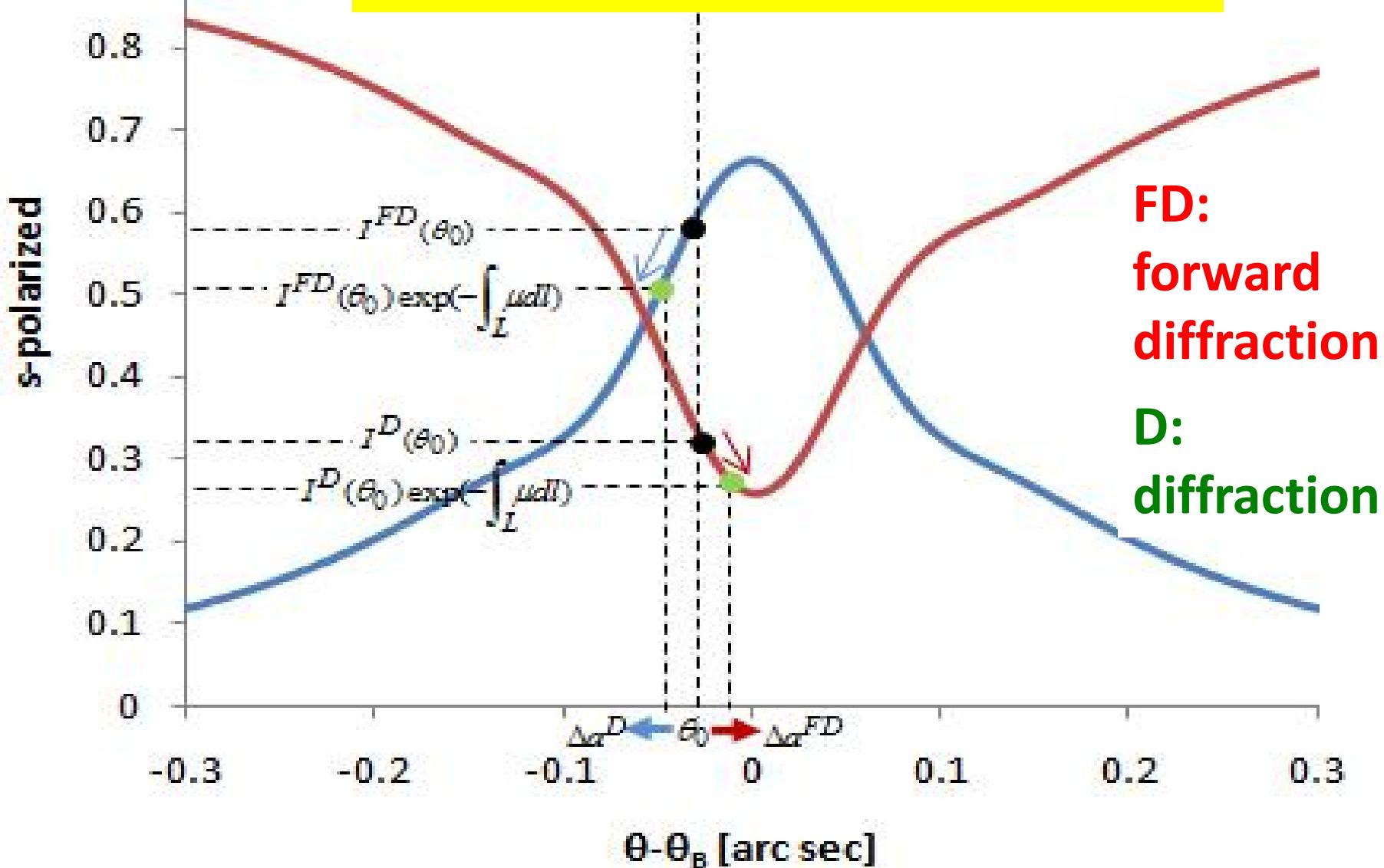
D:  
convex  
upward



# FD at LAA

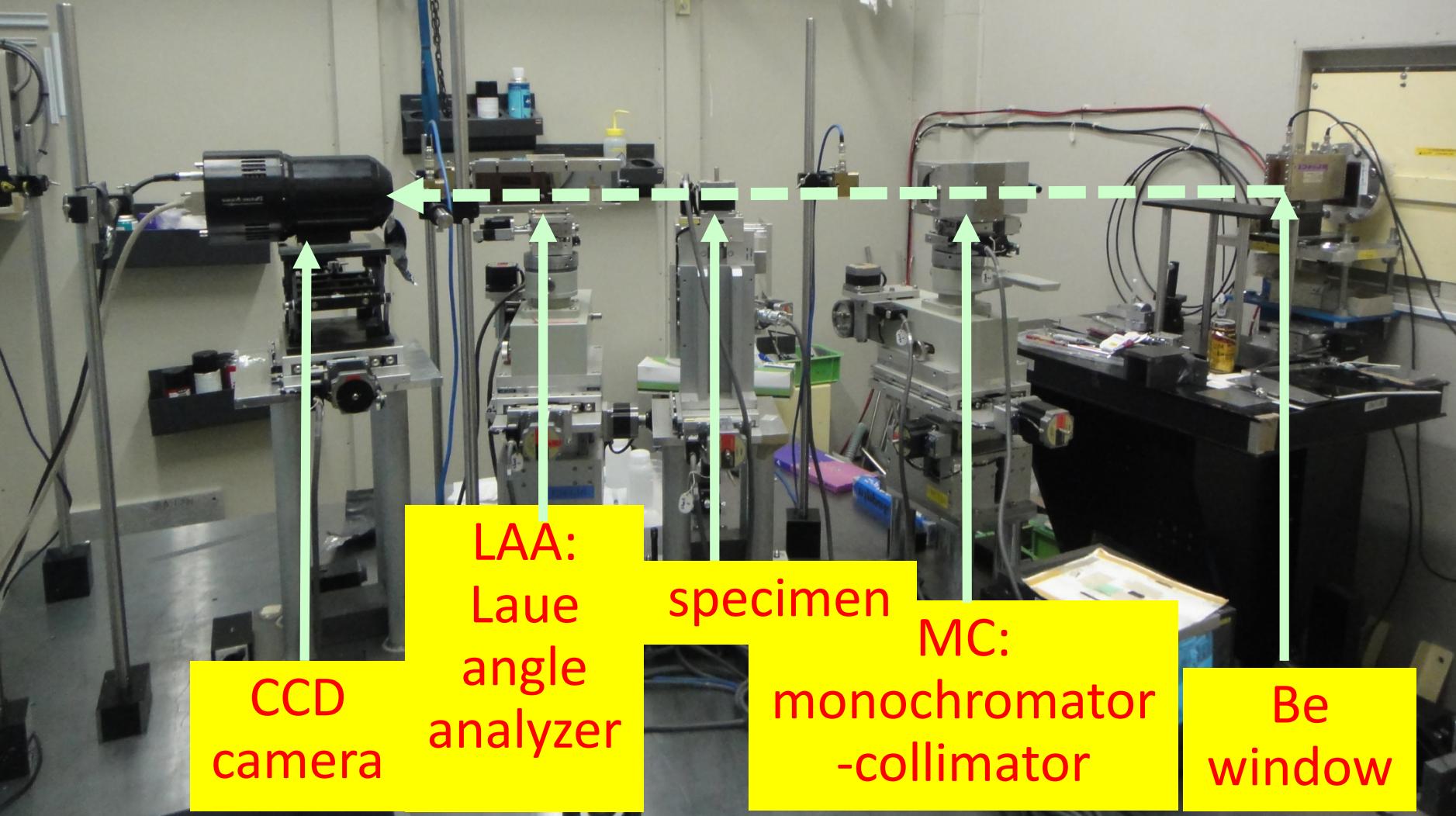


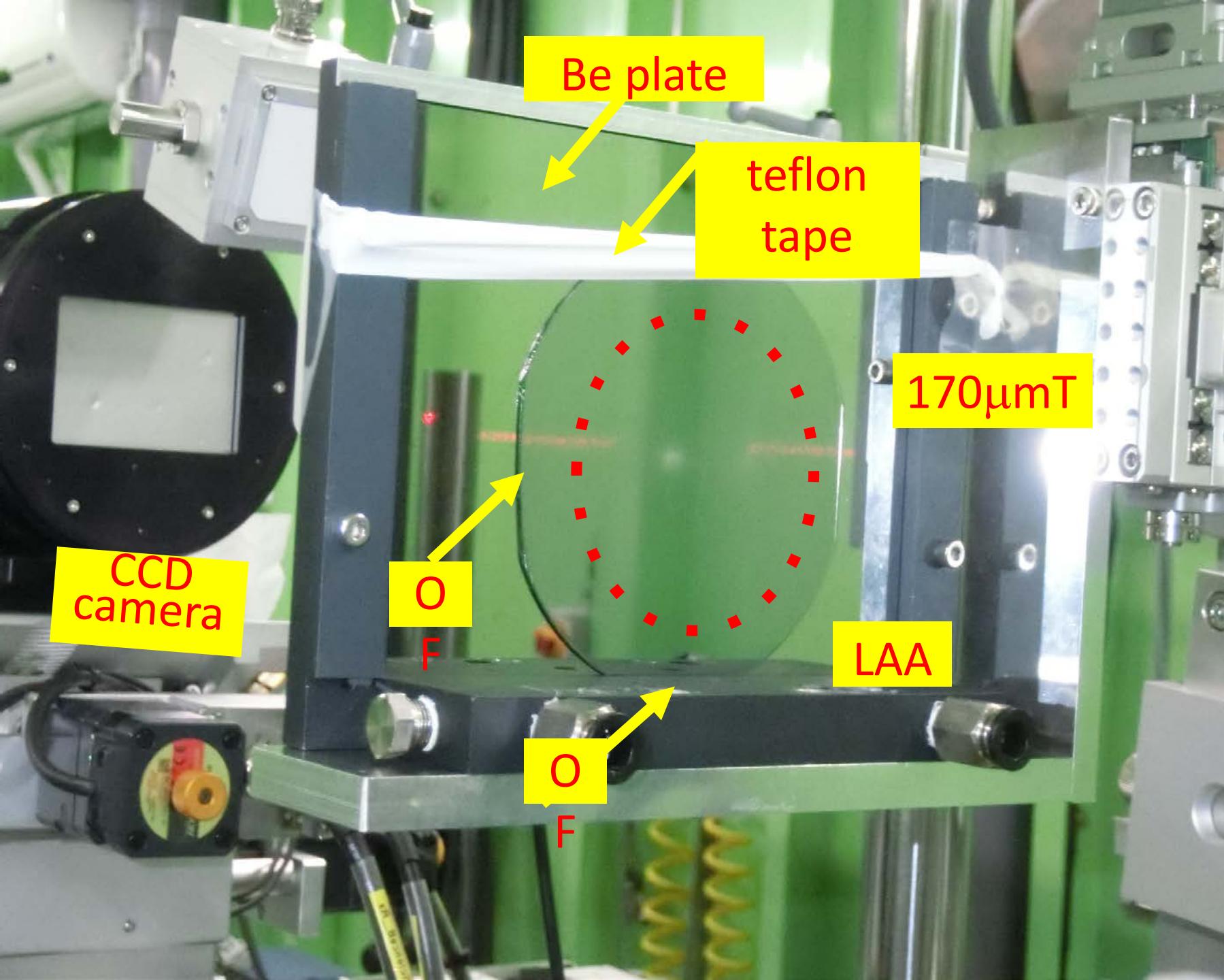
# How to obtain refraction component





BL14 Photon Factory  
5T superconducting vertical wiggler  
 $E_C = 20\text{keV}$ ,  $E=35\text{keV}$





FZ crystal:

**SUMCO**

grinding:

**DISCO**

acid

(HF+HNO<sub>3</sub>)

etching:

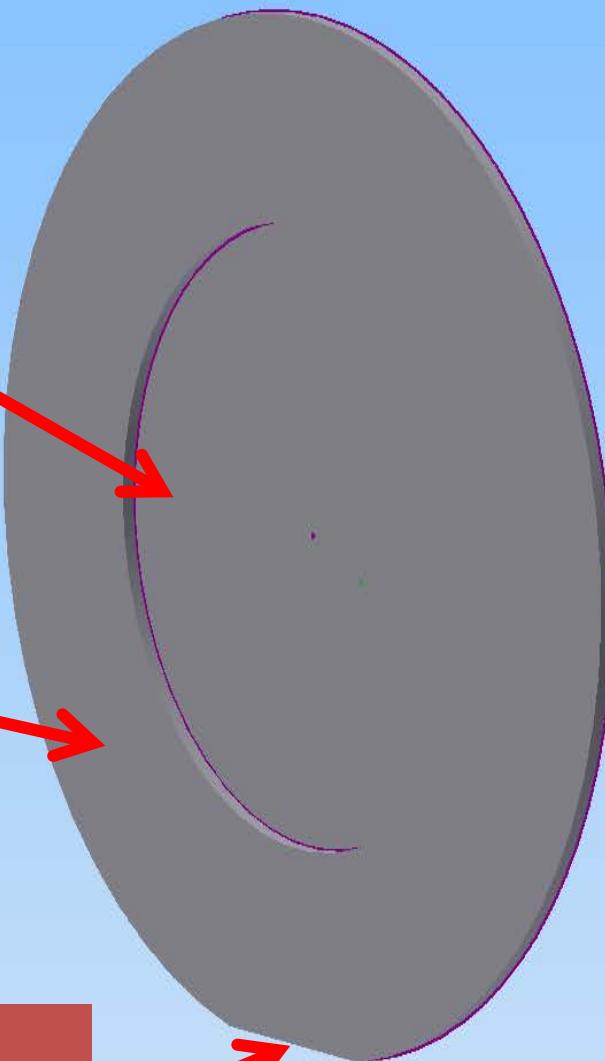
**Mimasu**

170 $\mu\text{m}$  thick

3mm  
thick

<110>

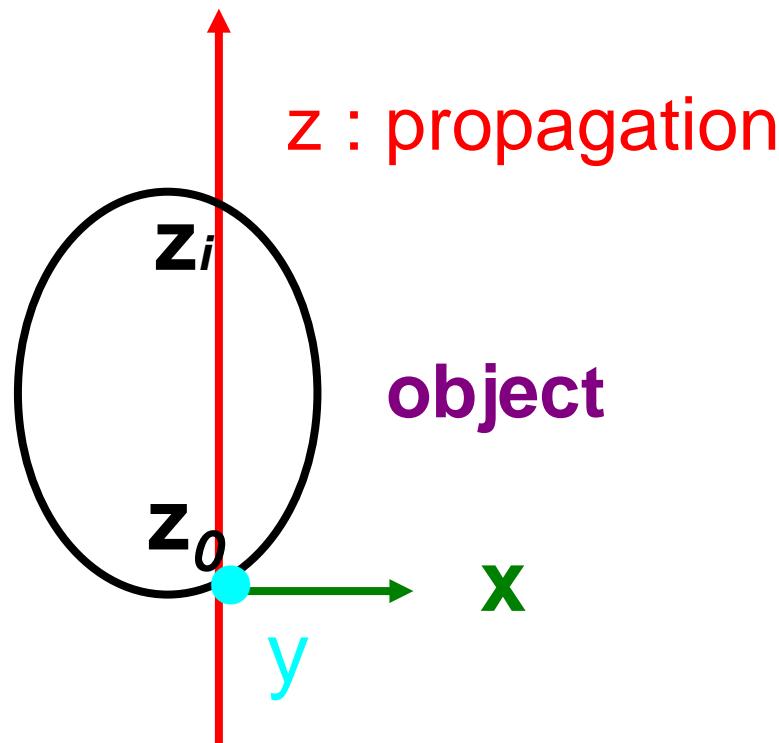
OF :orientation flat



# refraction angle: $\alpha$

$$\alpha = r_e h^2 c^2 / E^2 2\pi \int_{z_0}^{z_i} \partial n(r) / \partial x \, dz$$

$z_0 \quad n = 1 - \delta$

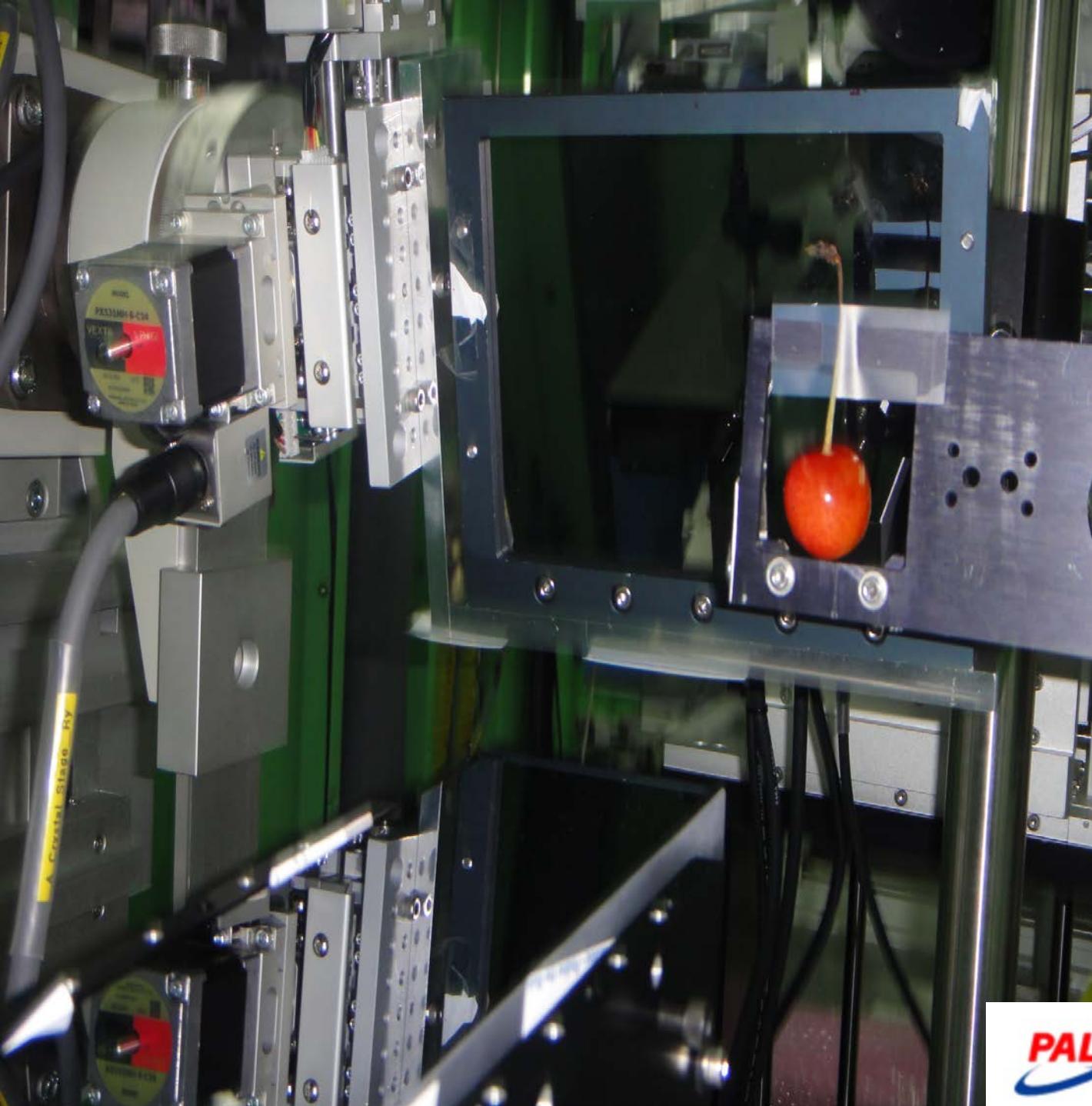


15 keV, 35 keV x-ray

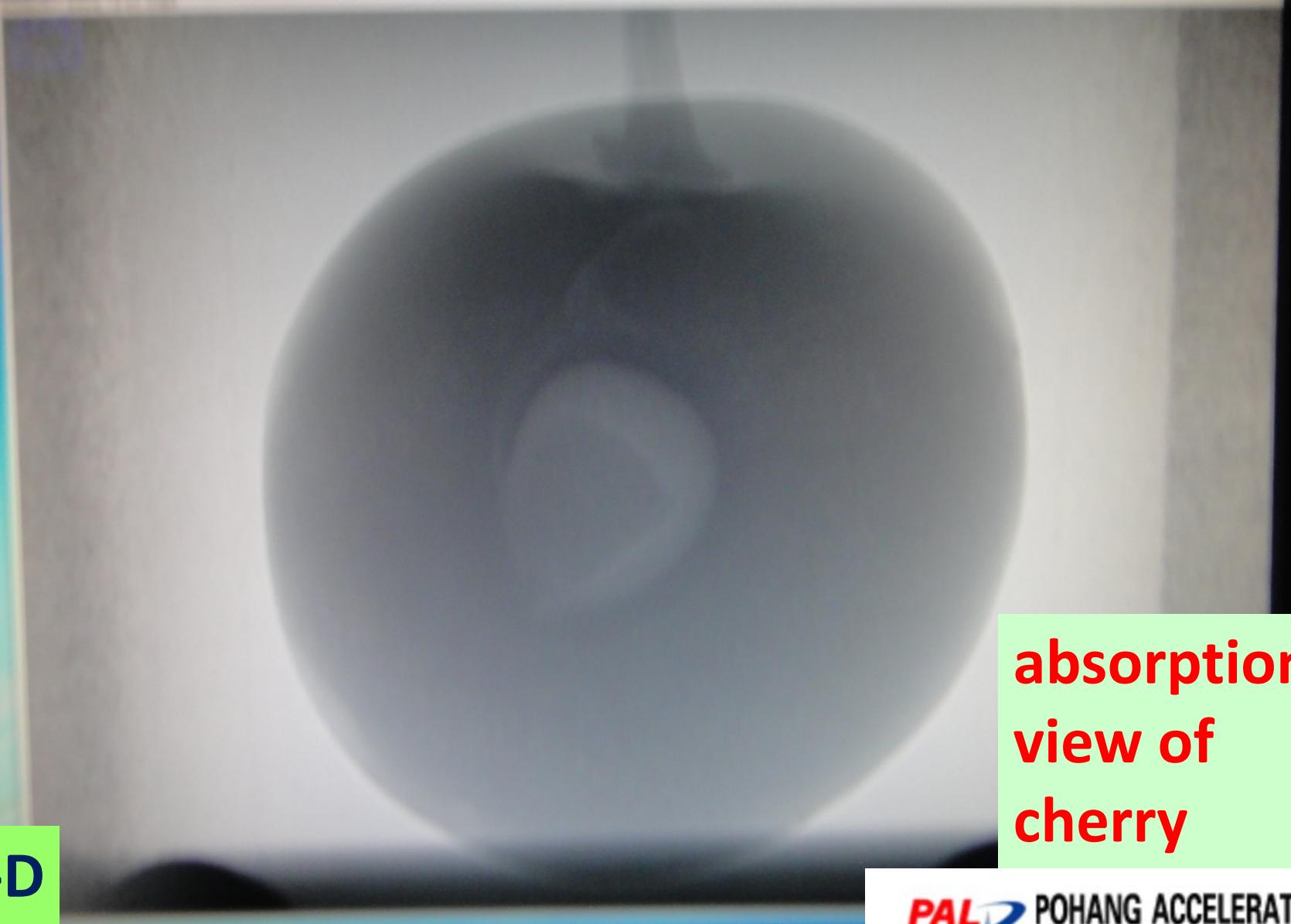
$n=1-\delta$

energy(keV)	Ca
15	$1.4 \times 10^{-6}$
35	$2.6 \times 10^{-7}$

# **2-DIMENSIONAL MED IMAGES**



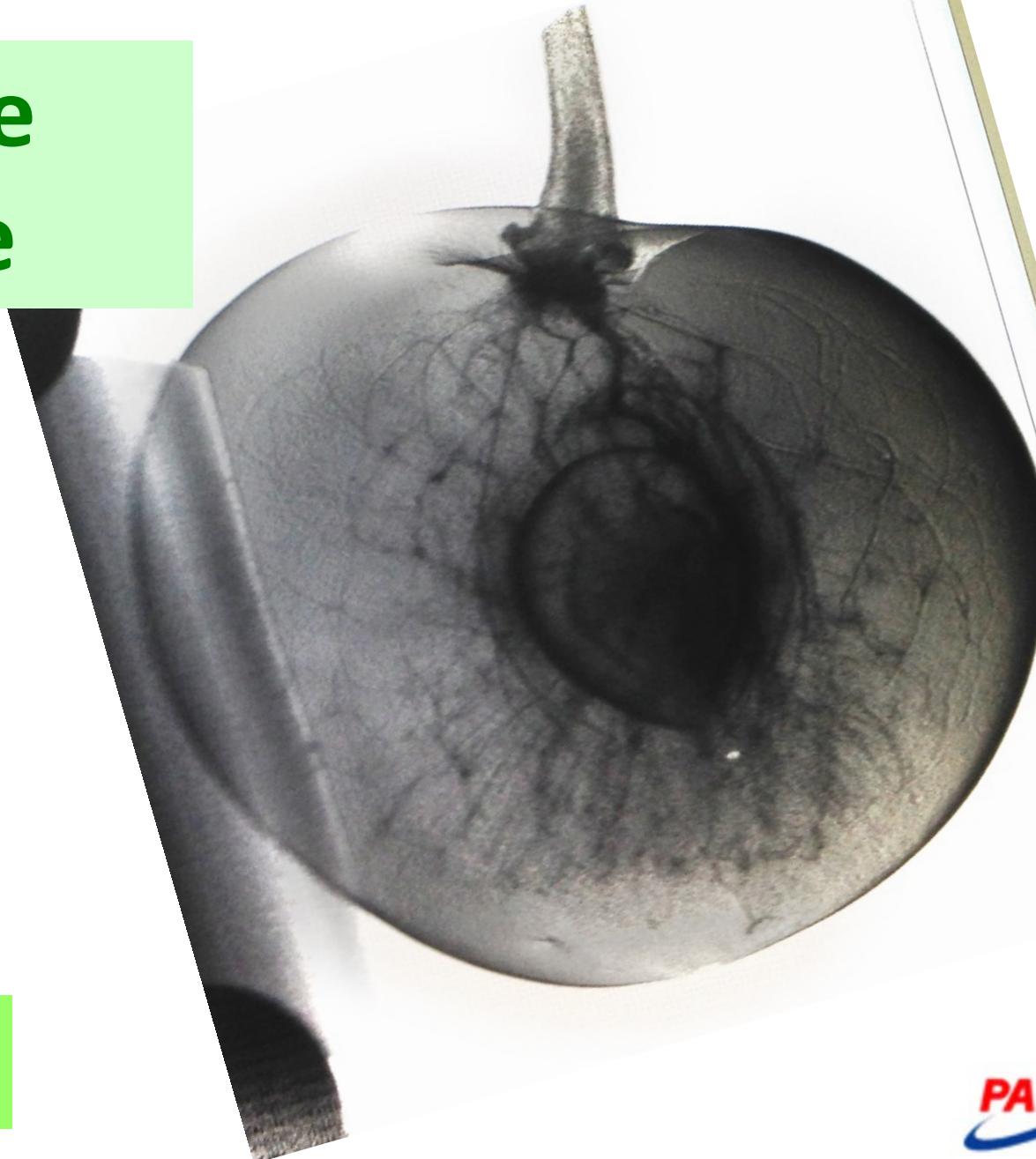
cherry in  
XDFI



2-D

absorption  
view of  
cherry

sieve  
tube



2-D

XDFI view  
of cherry

patella: femor

inner

ligament

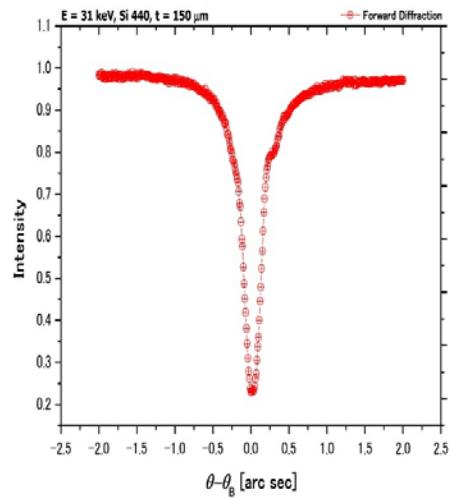
outer

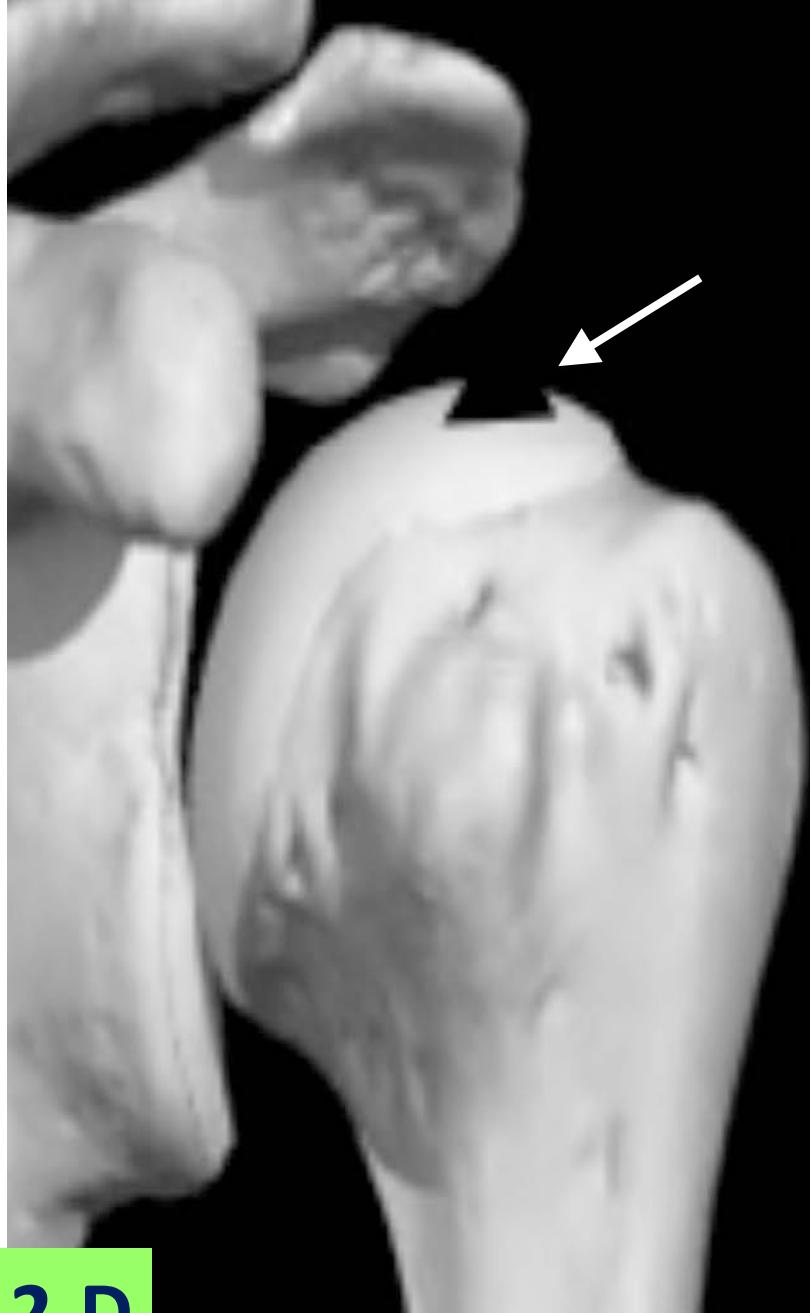
2-D

articular  
cartilage

10 mm

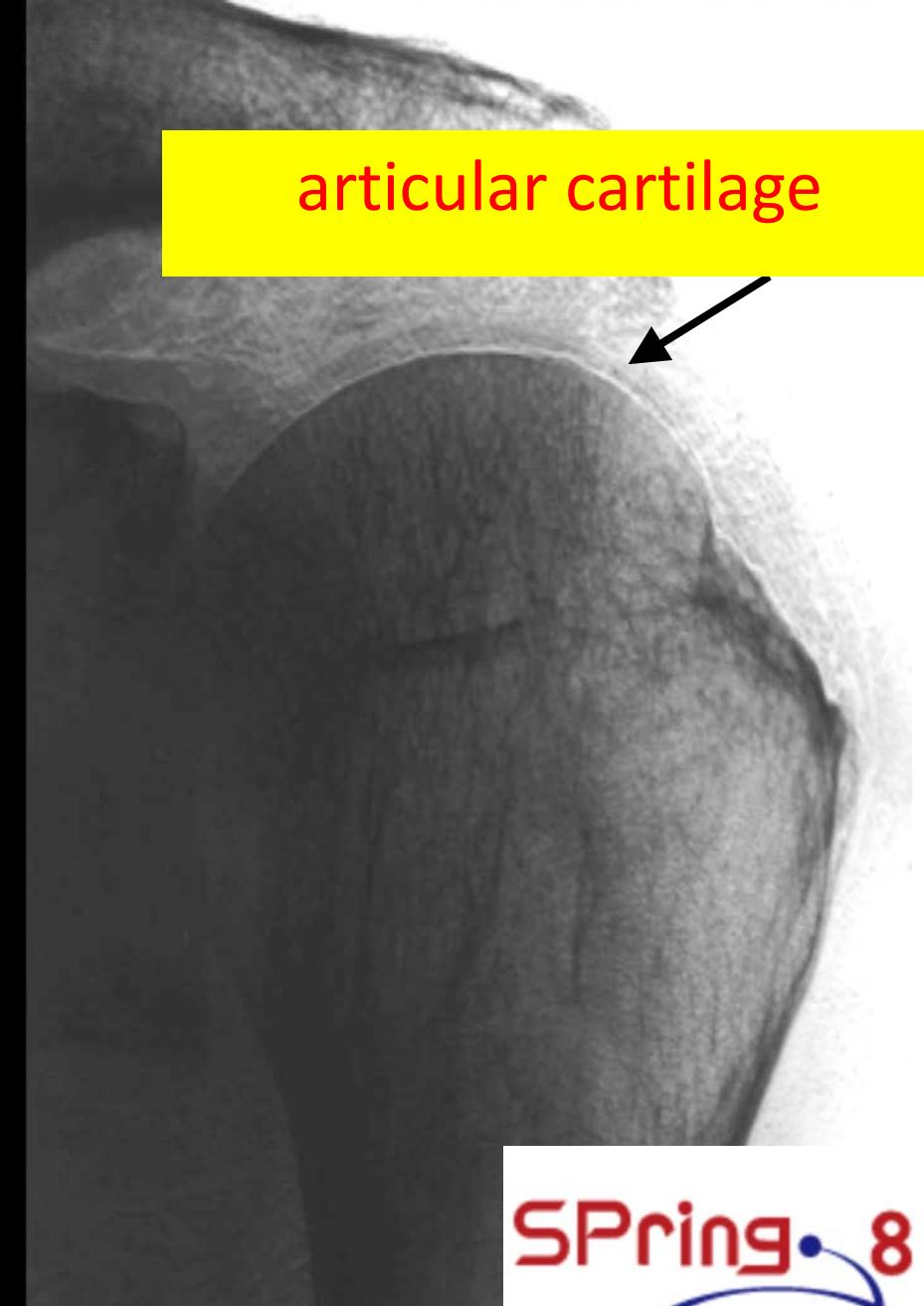
human articular  
cartilage at knee  
taken with XDFI





2-D

human shoulder, cadavar



articular cartilage

SPring-8

diameter  
5mm

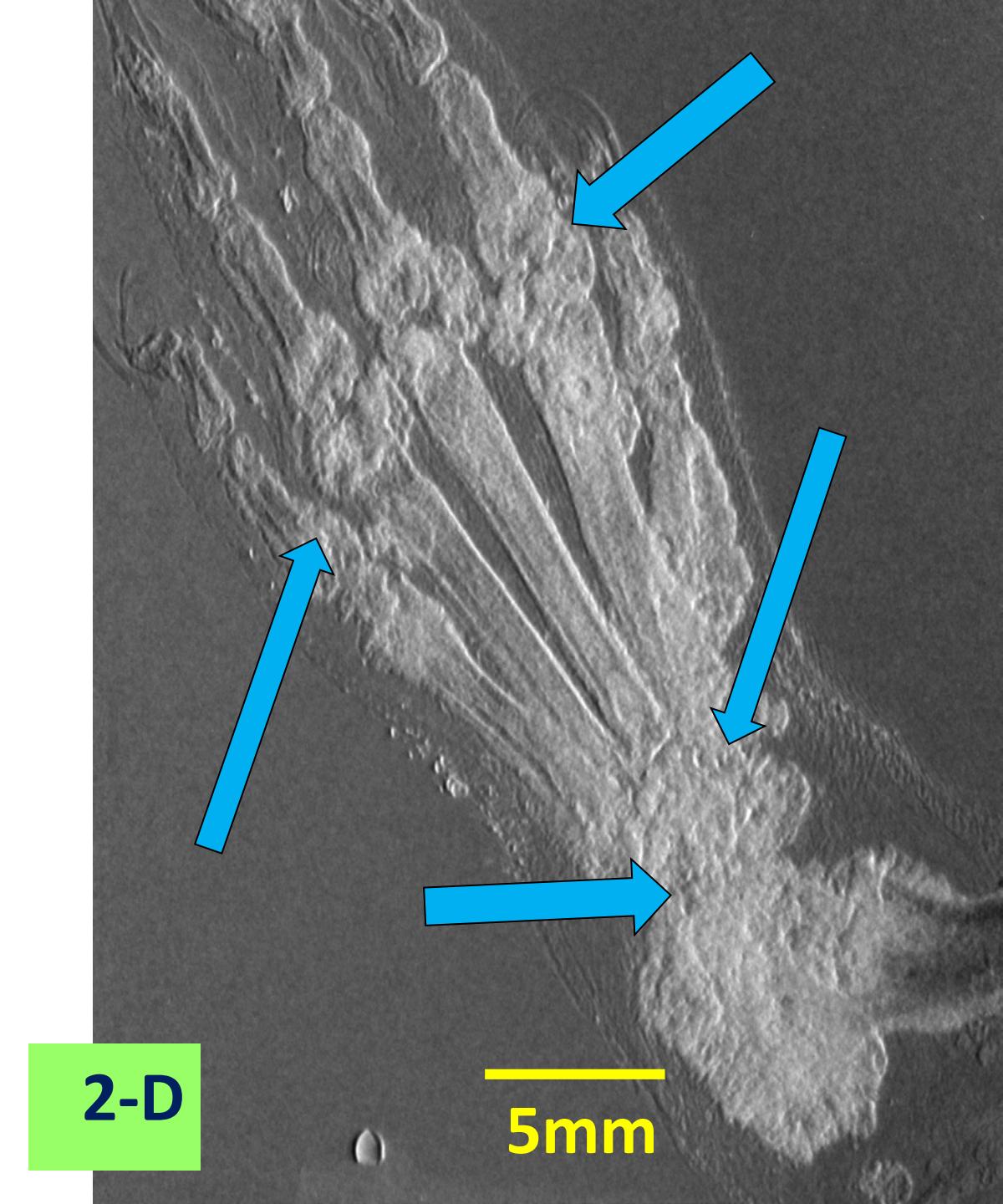
Ca?

5mm

2-D

damage to femoral  
artery  
due to diabetes

grafting with healthy  
artery/vein



2-D

5mm

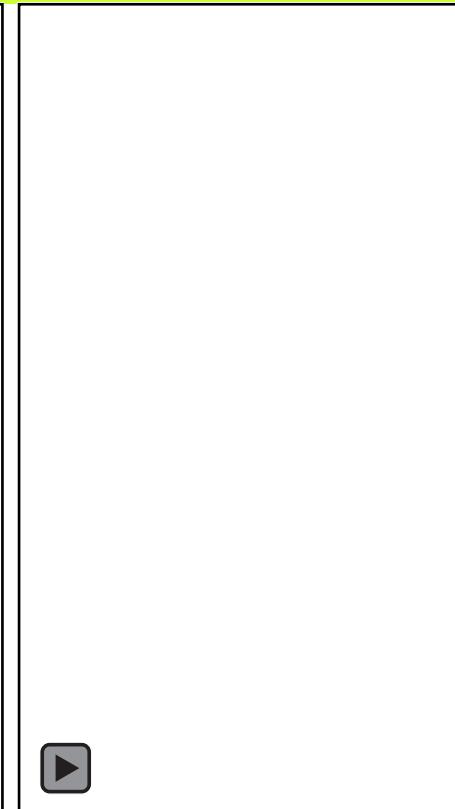
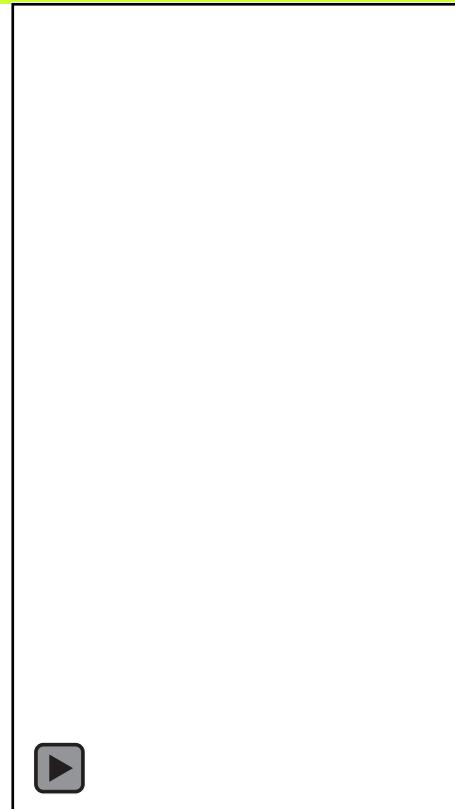
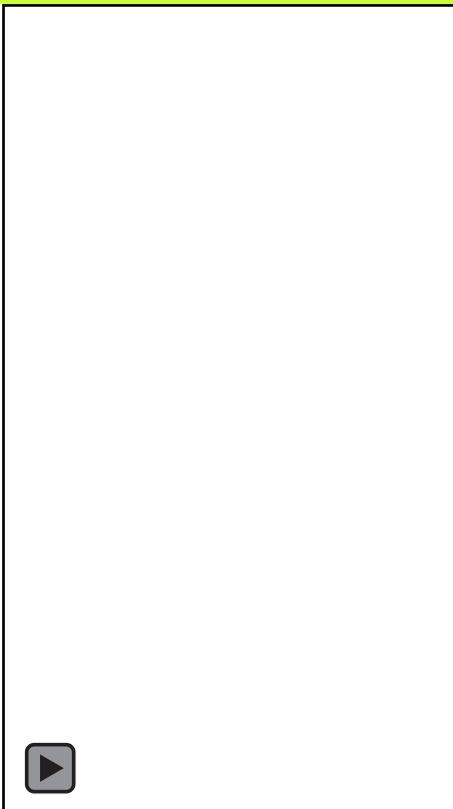
rheumatoid  
arthritis in rat

2 wks later  
after adjuvant  
injection at  
rat right foot

articular  
cartilage

# **3-DIMENSIONAL MED IMAGES**

rat foot 2 wks after adjuvant injected



3-D

abs

XDFI

mod XDFI

eye  
ball  
axial

3-D

Ando et al.  
3029



eye ball  
sagittal

3-D

Gupta  
al 201



globe and optic nerve  
insertion of the same  
specimen. lens; i  
muscle; ciliary bo  
pupil; cornea; op  
nerve sheath; scl  
central retinal art  
nerve fascicles  
retinchoroid; su  
choroidal space

3-D

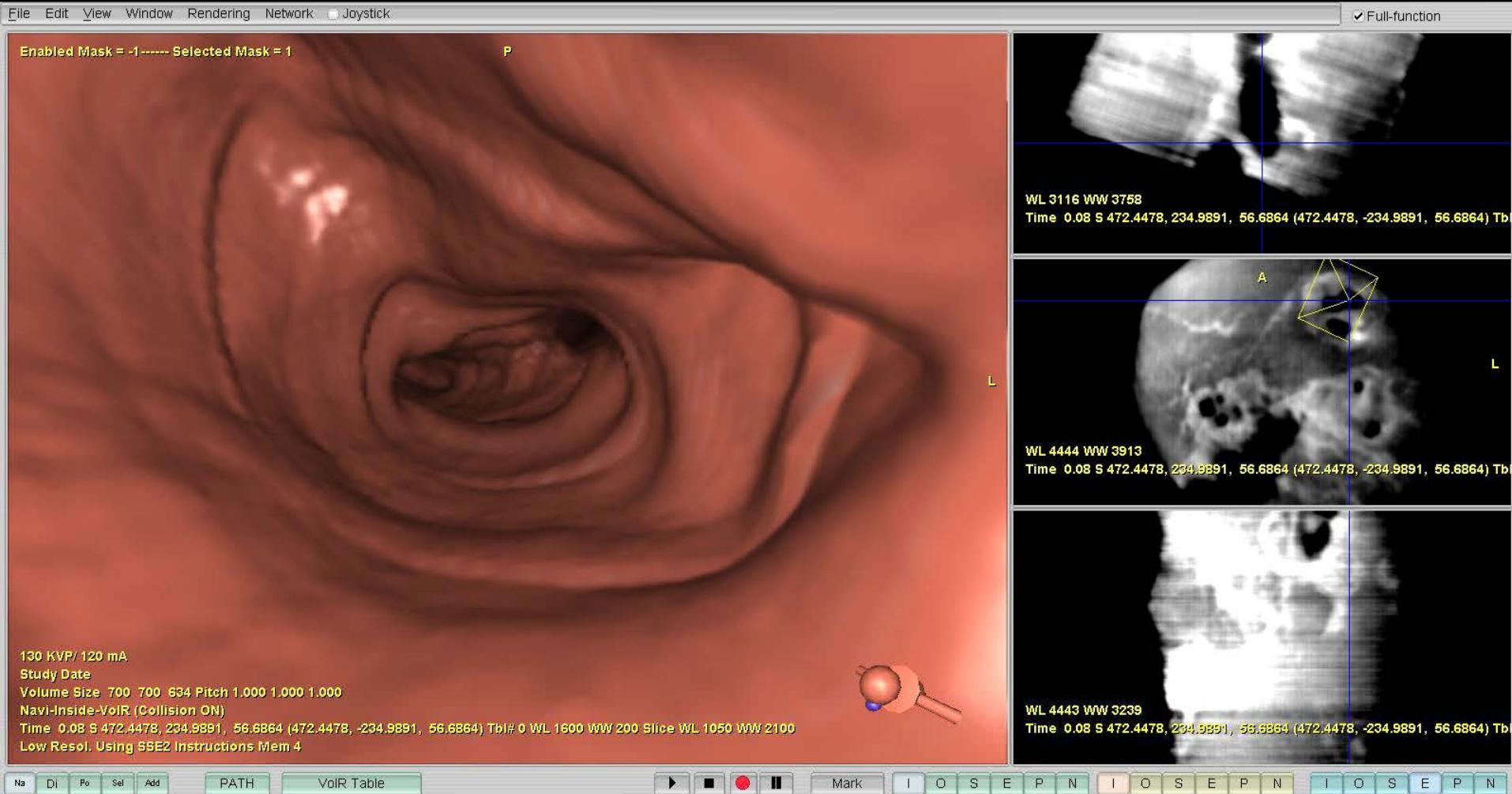
Gupta  
et al  
2012

3-D

DCIS absorption



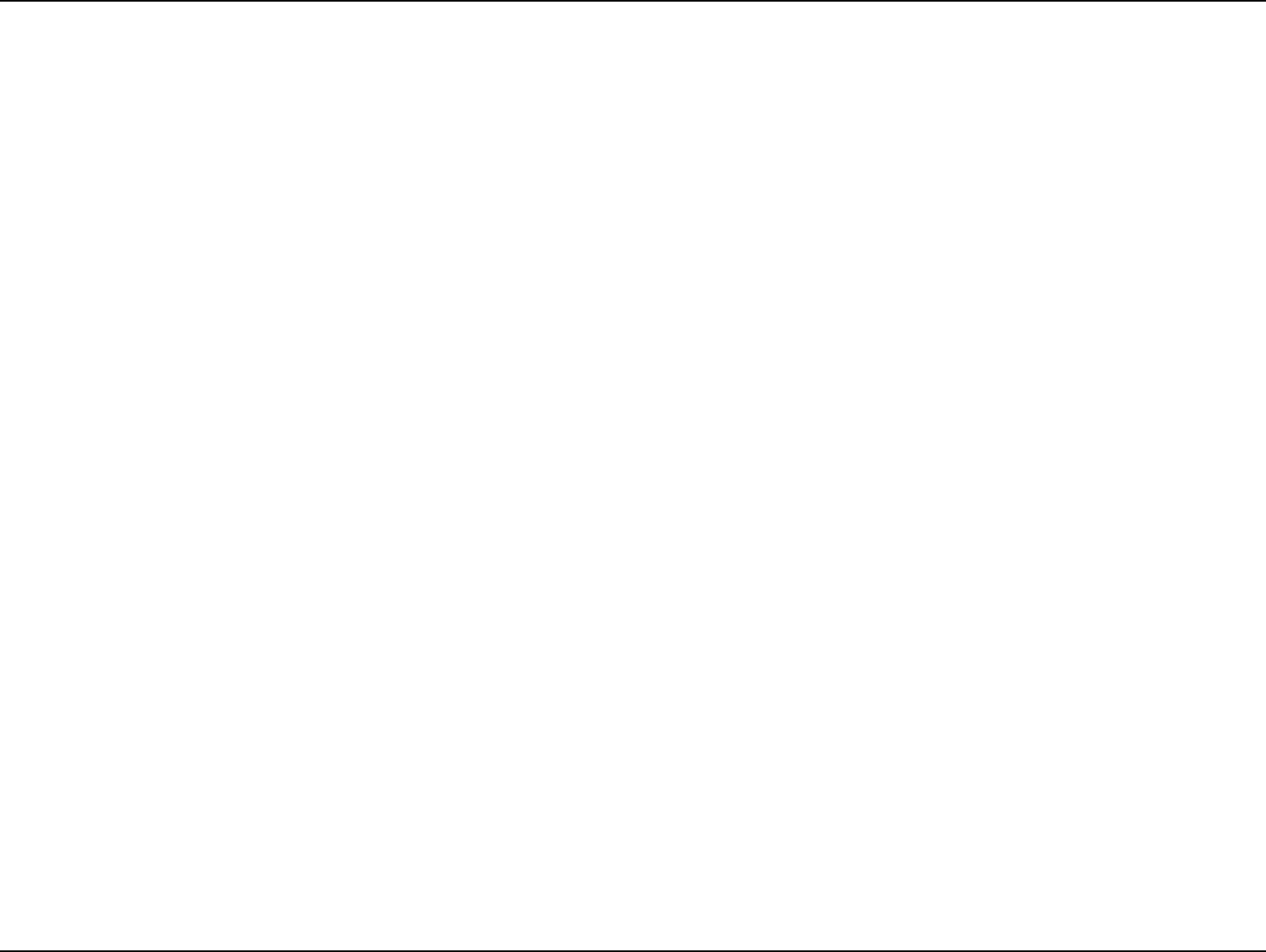




# DCIS endoscopy

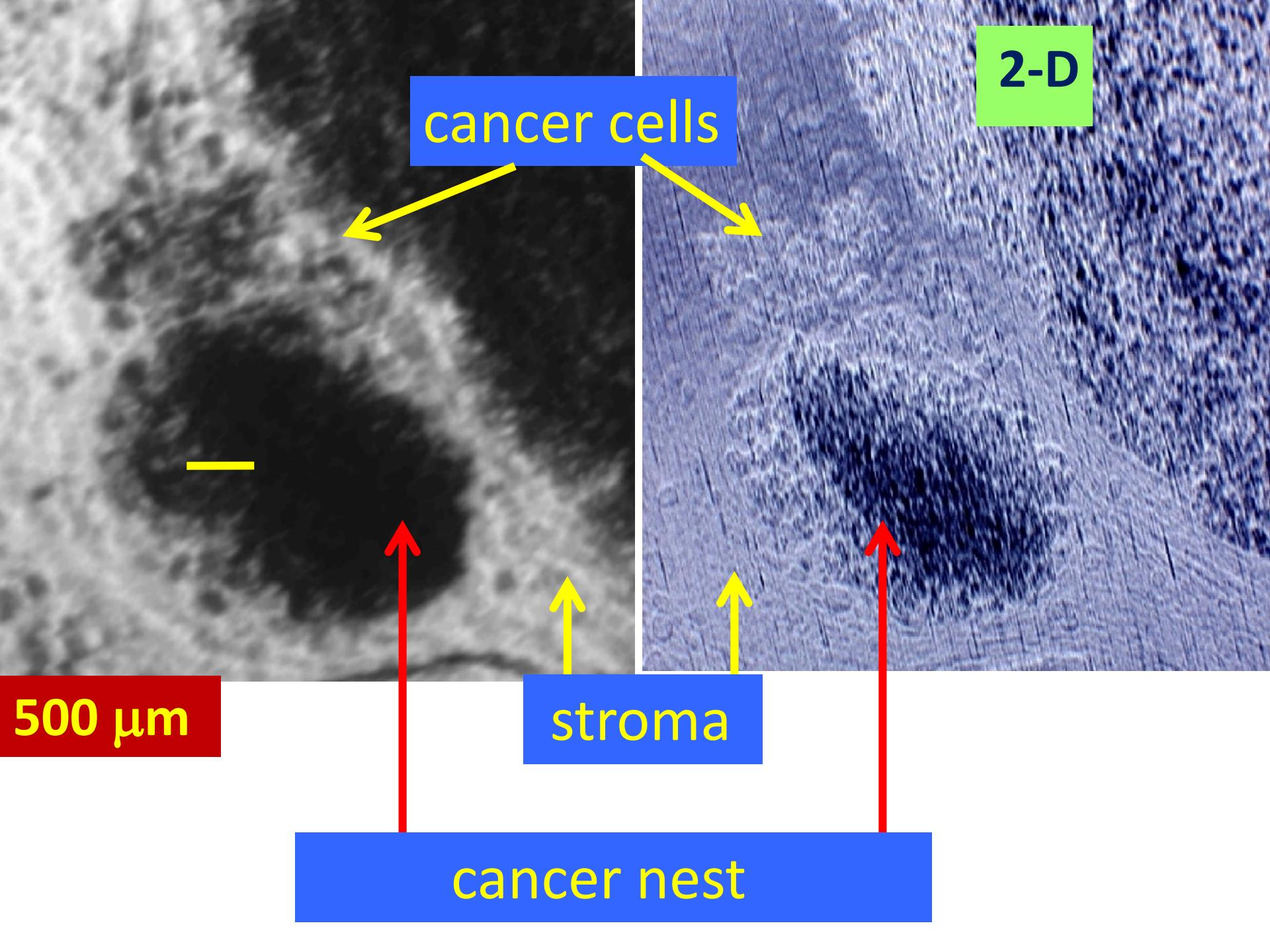
3-D

Ichihara et al Virchows Archiv  
451 (2008) 41-47



# STILL CHALLENGING TOWARDS

- 1) FOV SUITABLE FOR CLINICAL TRIAL  
~ 26 CM X 26CM**
- 2) DIFFICULT TISSUE SUCH AS BRAIN TUMOUR WITH SKULL**
- 3) HIGHER SPATIAL RESOLUTION IN 2D AND CT IMAGE ~ 2-3 MICRON**
- 4) CLINICAL DIRECTION AND PATHOLOGICAL DIRECTION**



2-D

cancer cells

stroma

cancer nest

500  $\mu\text{m}$

# spatial resolution measured by using MTF chart

material:

W  $1.0\mu\text{m}^T$

X-ray energy:

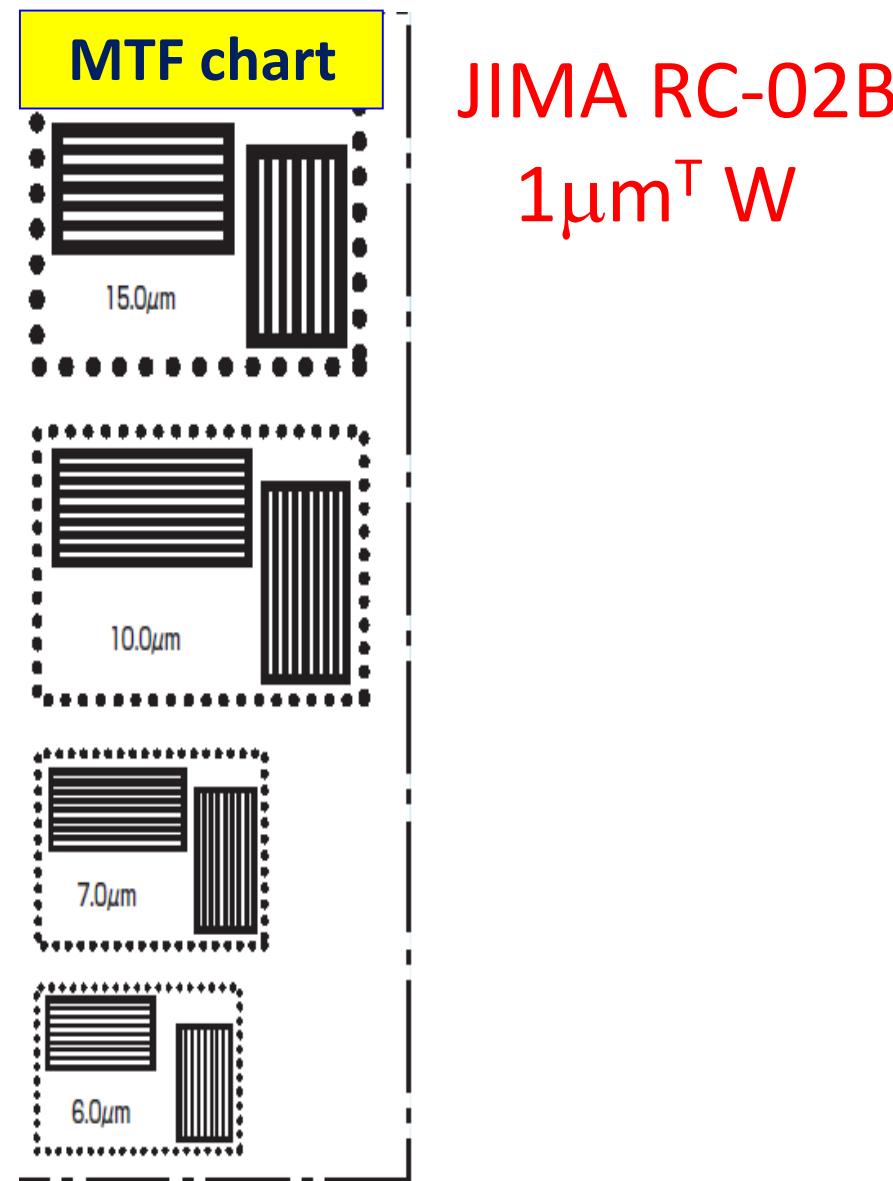
31keV

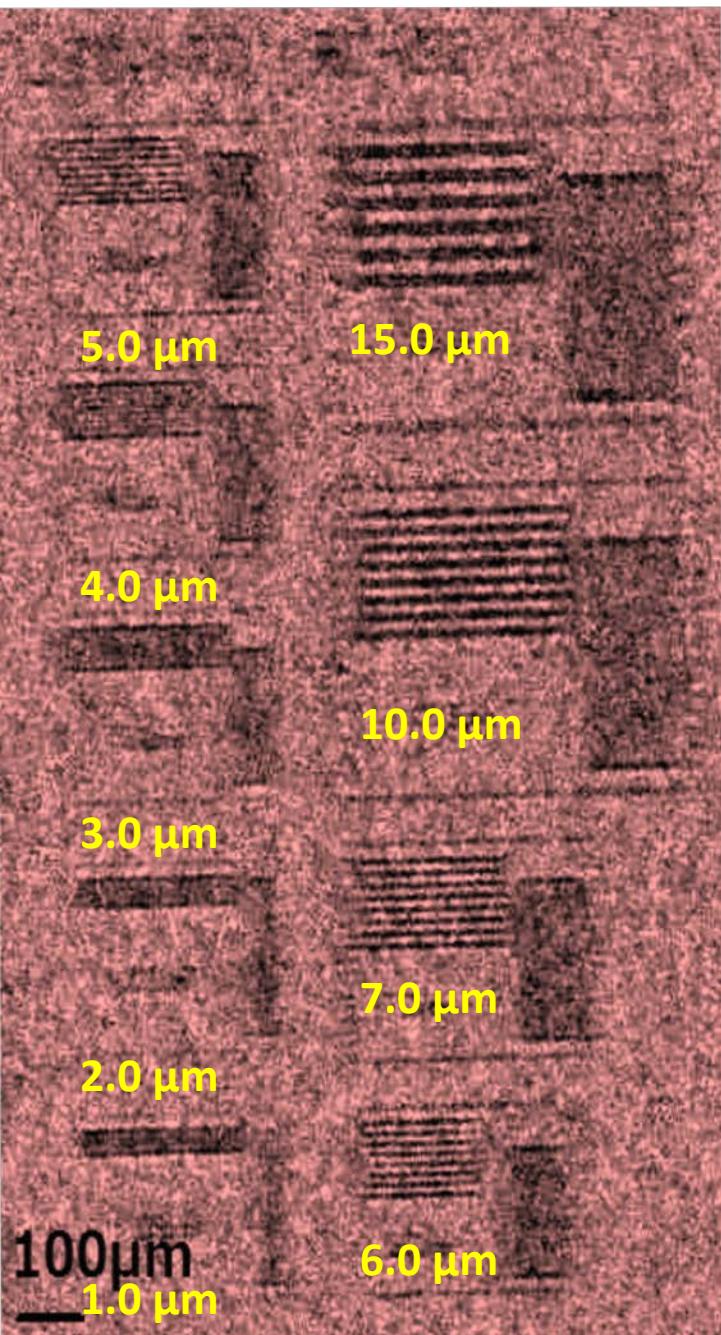
CCD camera:

$3\mu\text{m}/\text{pixel}$

LAA thickness:

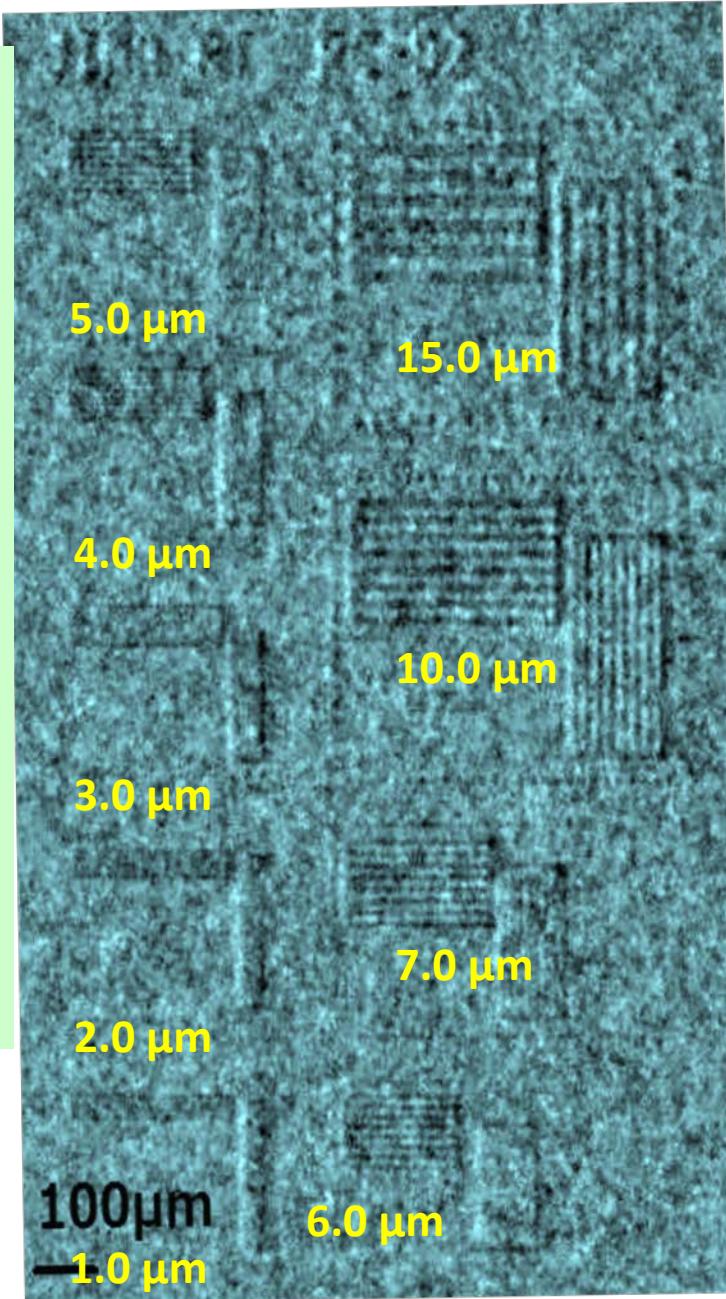
$150\mu\text{m}$





left:  
without MC  
right:  
with MC

31keV  
150 $\mu\text{m}$



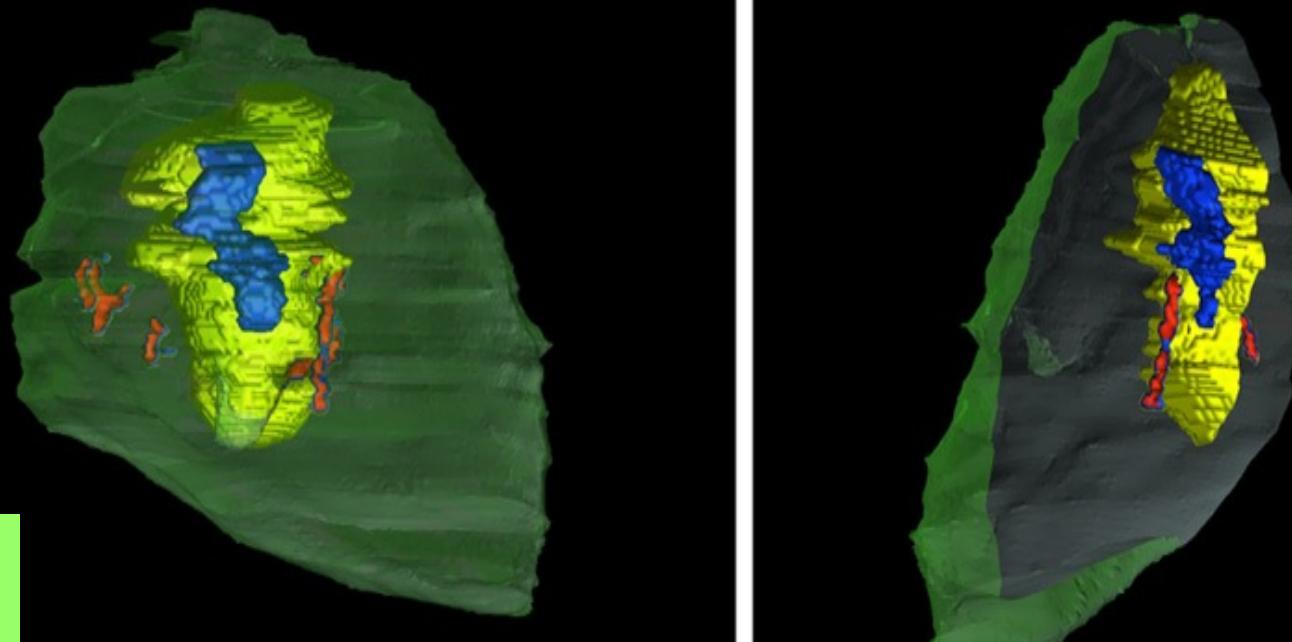
# Requirement for x-ray optics in medical imaging

	FOV	Spatial resolution in 2D	Detector pixel number
Clinical	25cm x 25cm > 10cm x 10cm	25μm < 100μm	100,000,000 > 10,000,000
Pathological	2.5cm(H) x 2.5cm(V)	1μm < 2~3μm < 7.4μm (Photonics Science)	625,000,000 100,000,000 34,000,000

**Thank you very much for the kind  
attention !**

**Looking very much forward to more  
collaboration**

3-D



*rat brain model with skull removed: DEI-CT view delineates, edema (green), viable tumor (yellow), necrosis (blue) and MVP (red) were visualized.*

S.-J. Seo, N. Sunaguchi, M. Ando, K.-H. Choi, H. Kim, W.-S. Chang, K.-H. Kim, J.-K. Kim, PMB 57 (2012) 1251-1262.

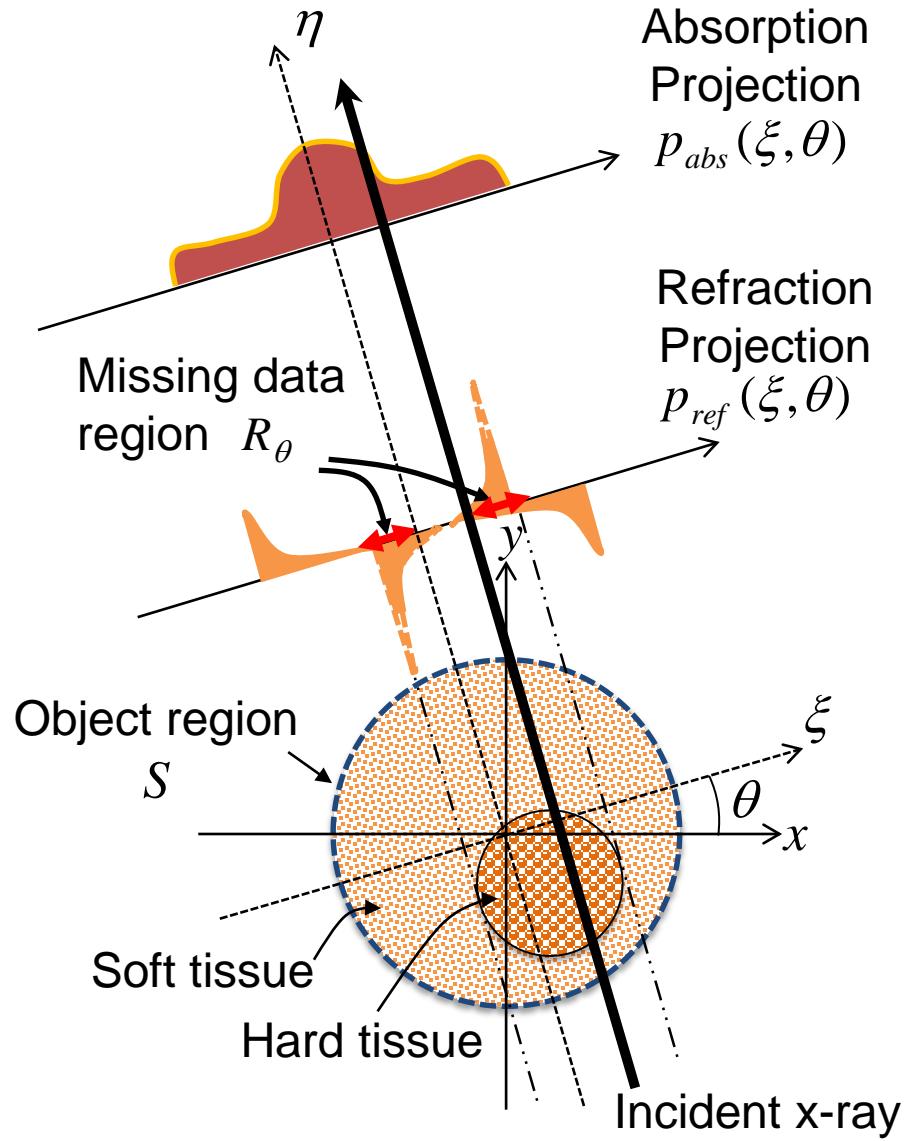
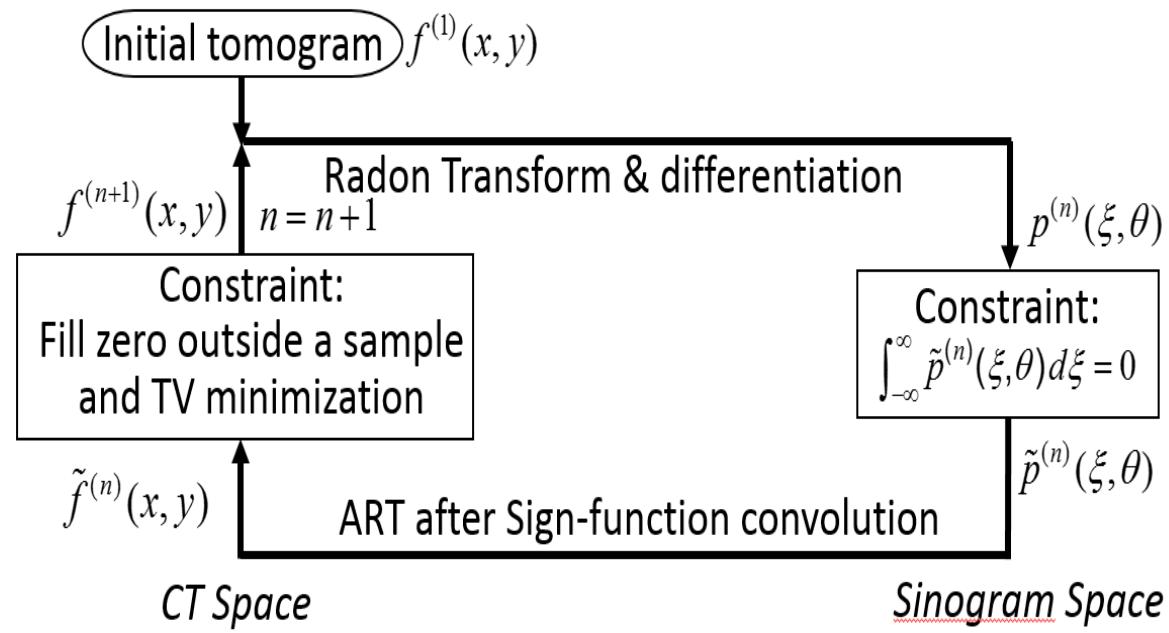
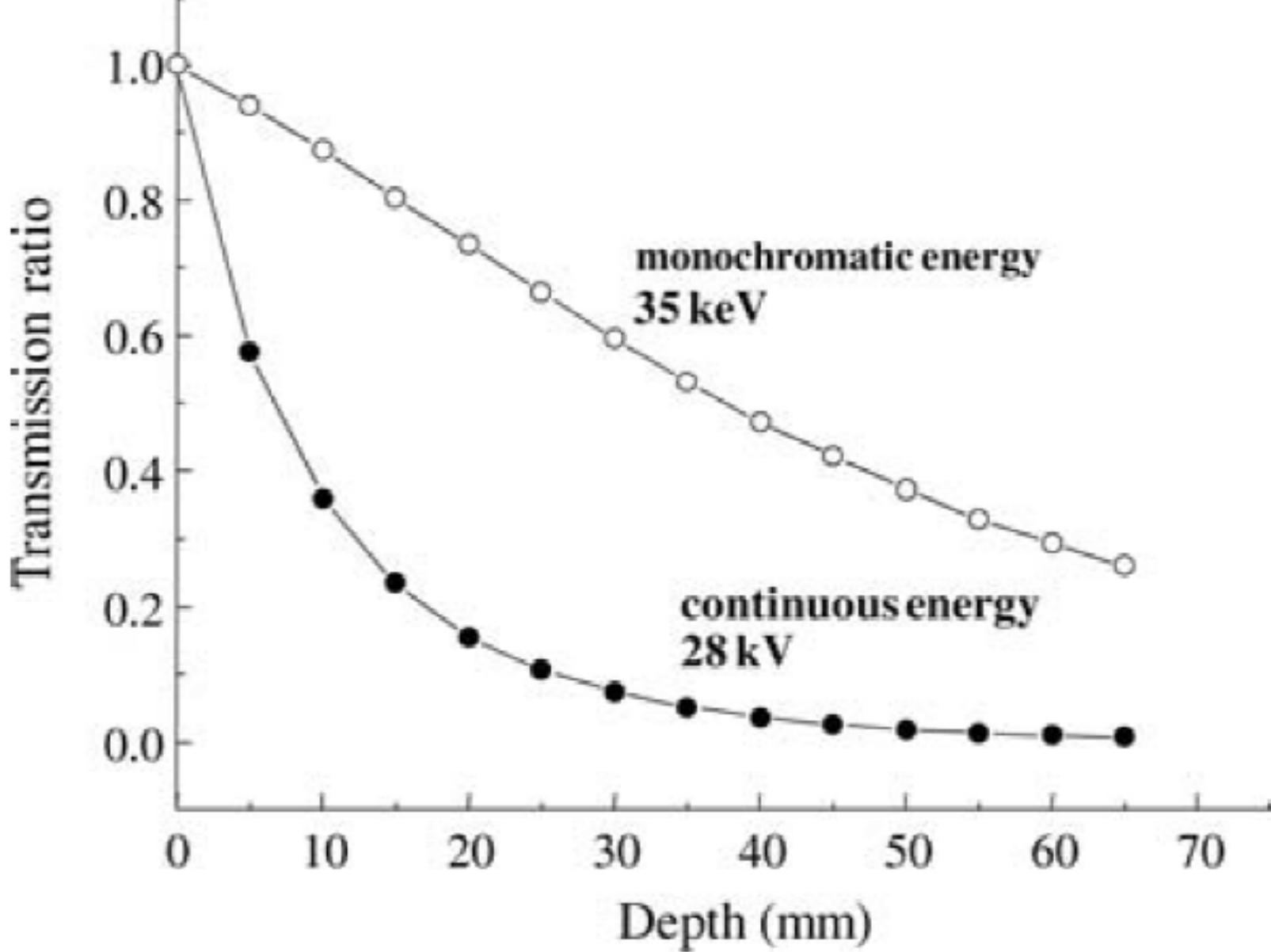
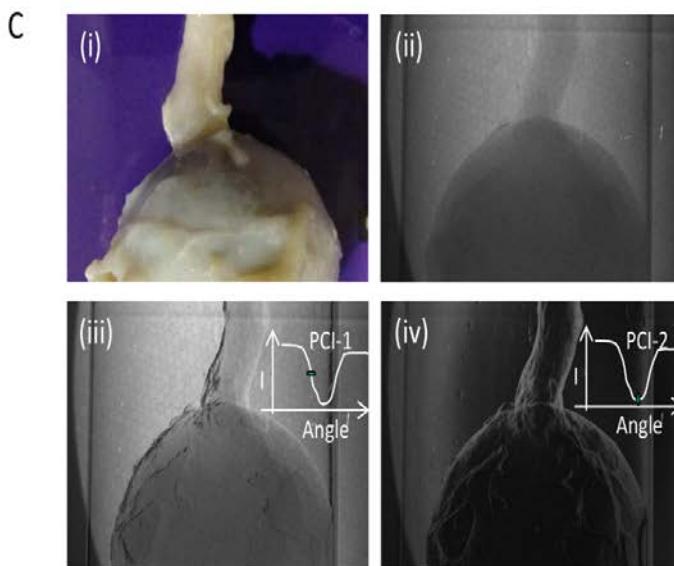
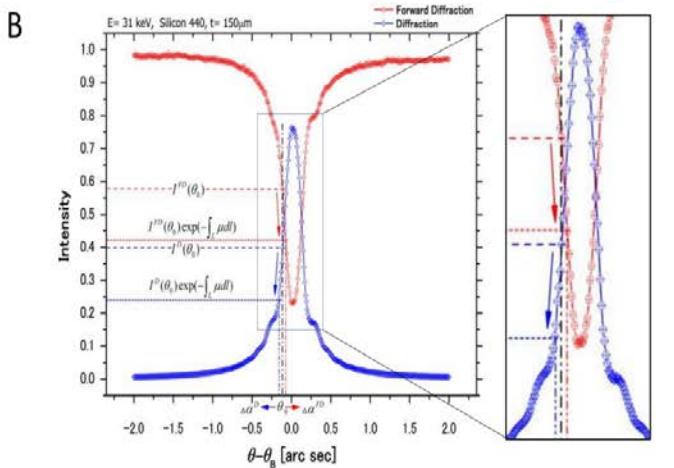
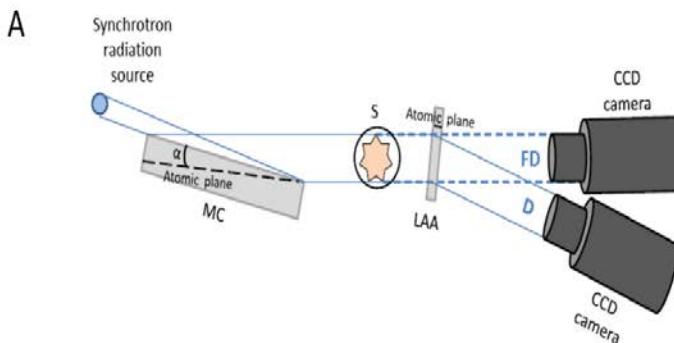


Fig. 4

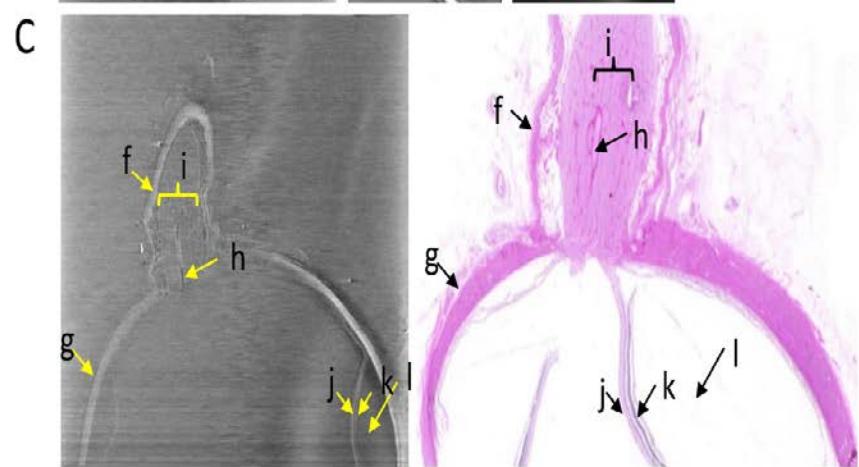
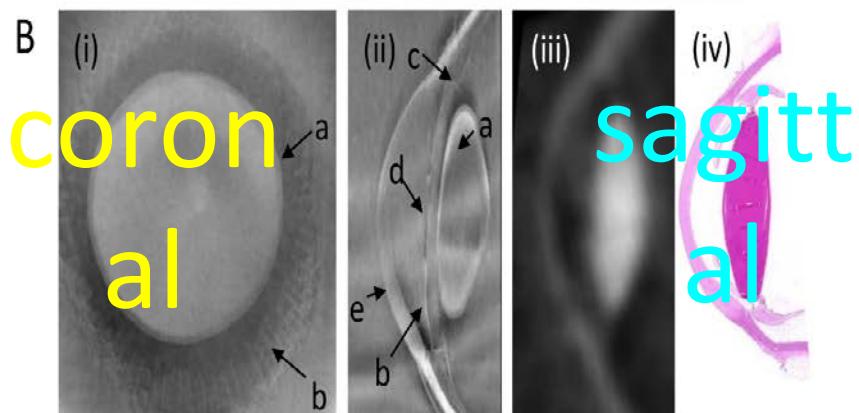
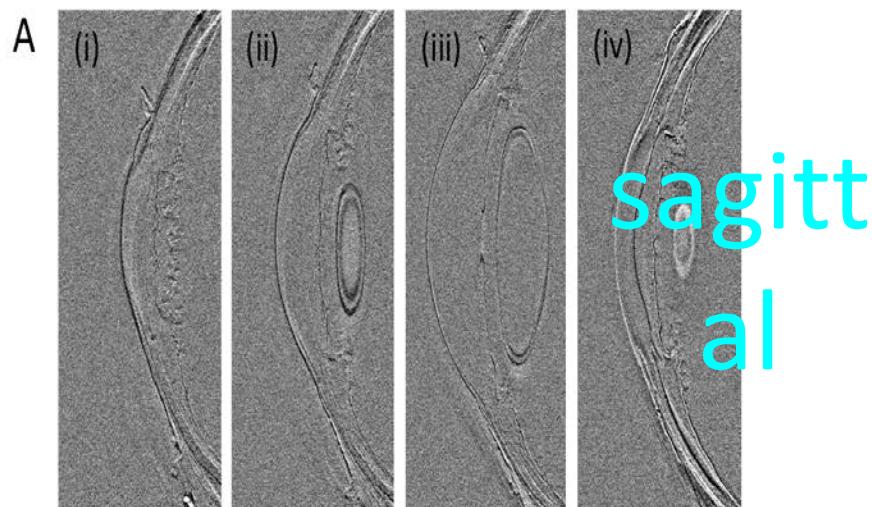




H. Sato, M. Ando, D. Shimao  
Radiation Protection Dosimetry 145 (2011) 1-3.



- Experimental system and rocking curve for the Laue-case angle analyser. (A) Experimental system. MC: Bragg-type monochromator-collimator; LAA: Laue-case angle analyser. (B) Rocking curve for the 150- $\mu\text{m}$ -thick LAA used in this study; the arrows show the displacement of operating points on the dark and bright field images due to tissue absorption. (C) Projection images of an eye specimen showing the posterior globe and optic nerve insertion: (i) photographic image, (ii) diffraction or bright field image, (iii and iv) forward diffraction or dark field images obtained at two different angles.



- Anterior and posterior chambers of an eyeball imaged with XDFI. (A) Tomographic slices (sagittal plane) of the differential phase map of an anterior chamber ((i)-(iv)). (B) Refractive index images in the coronal (i) and sagittal (ii) planes, a matched absorption image from conventional MDCT (iii), and a histological H&E slide showing this specimen (iv); the difference in the imaging and histological appearance are due to tissue retraction during histological processing and sample preparation.

(C) Globe and optic nerve insertion of the same specimen. a: lens; b: iris muscle; c: ciliary body; d: pupil; e: cornea; f: optic nerve sheath; g: sclera; h: central retinal artery; i:

# refraction angle: $\alpha$

$$\alpha =$$

$$r_e h^2 c^2 / E^2 2\pi \int \partial n(r) / \partial x dz$$

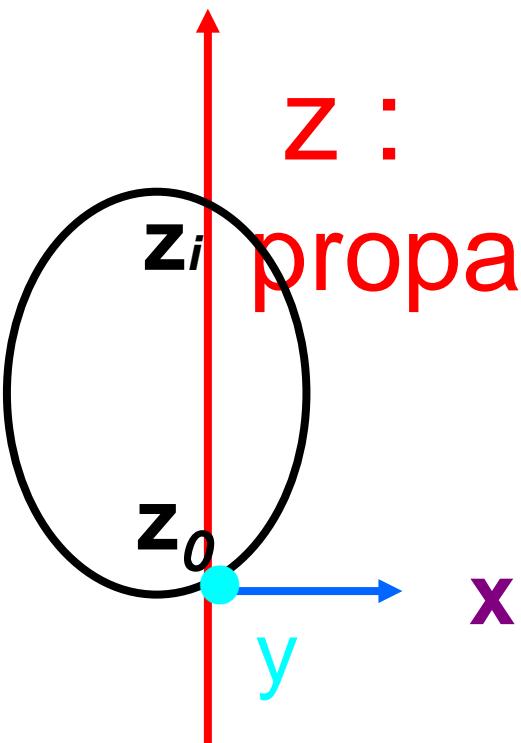
$$= 1 - \delta$$

objekt

$z_i$

$z_0 \quad n$

$z :$   
propagation



phase grating made  
by  
Professor Guo  
Jinchuan

20 $\mu\text{m}$



201411

07

silicon

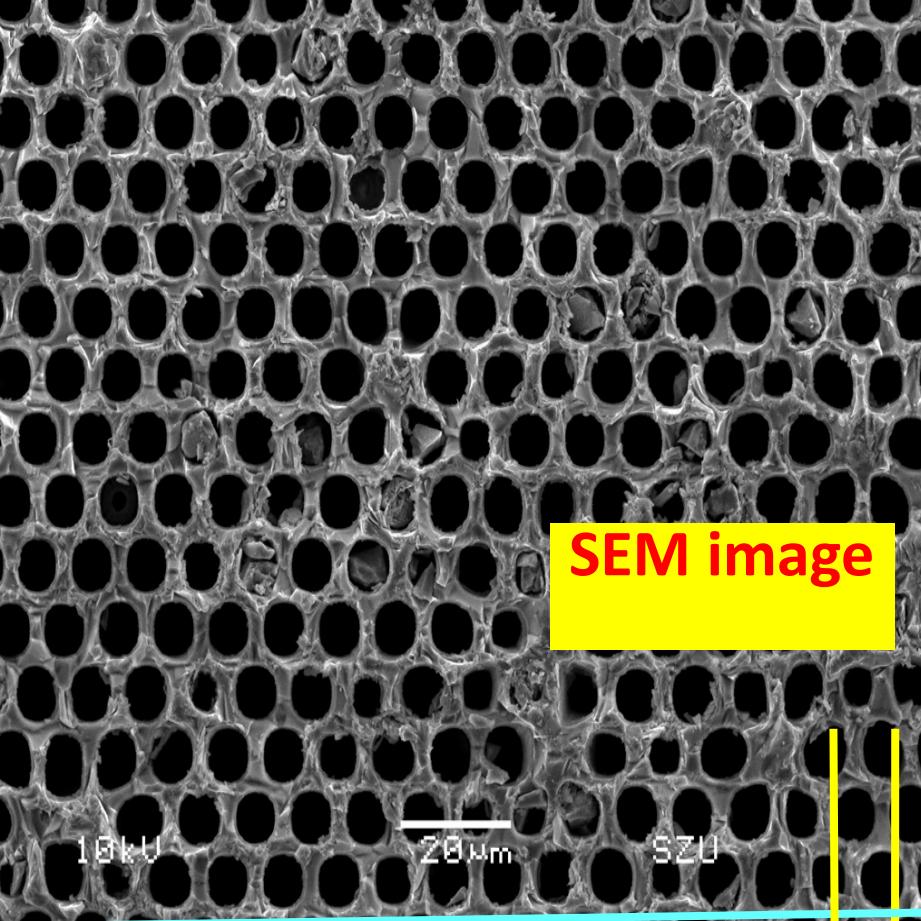
no crystal

100 $\mu\text{m}$

250 $\mu\text{m}$

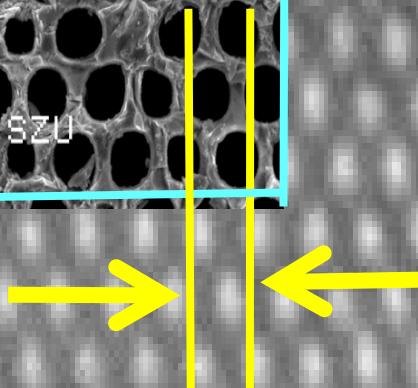
direct view with 35keV X-rays

detector made by  
Professor Guo  
Jinchuan

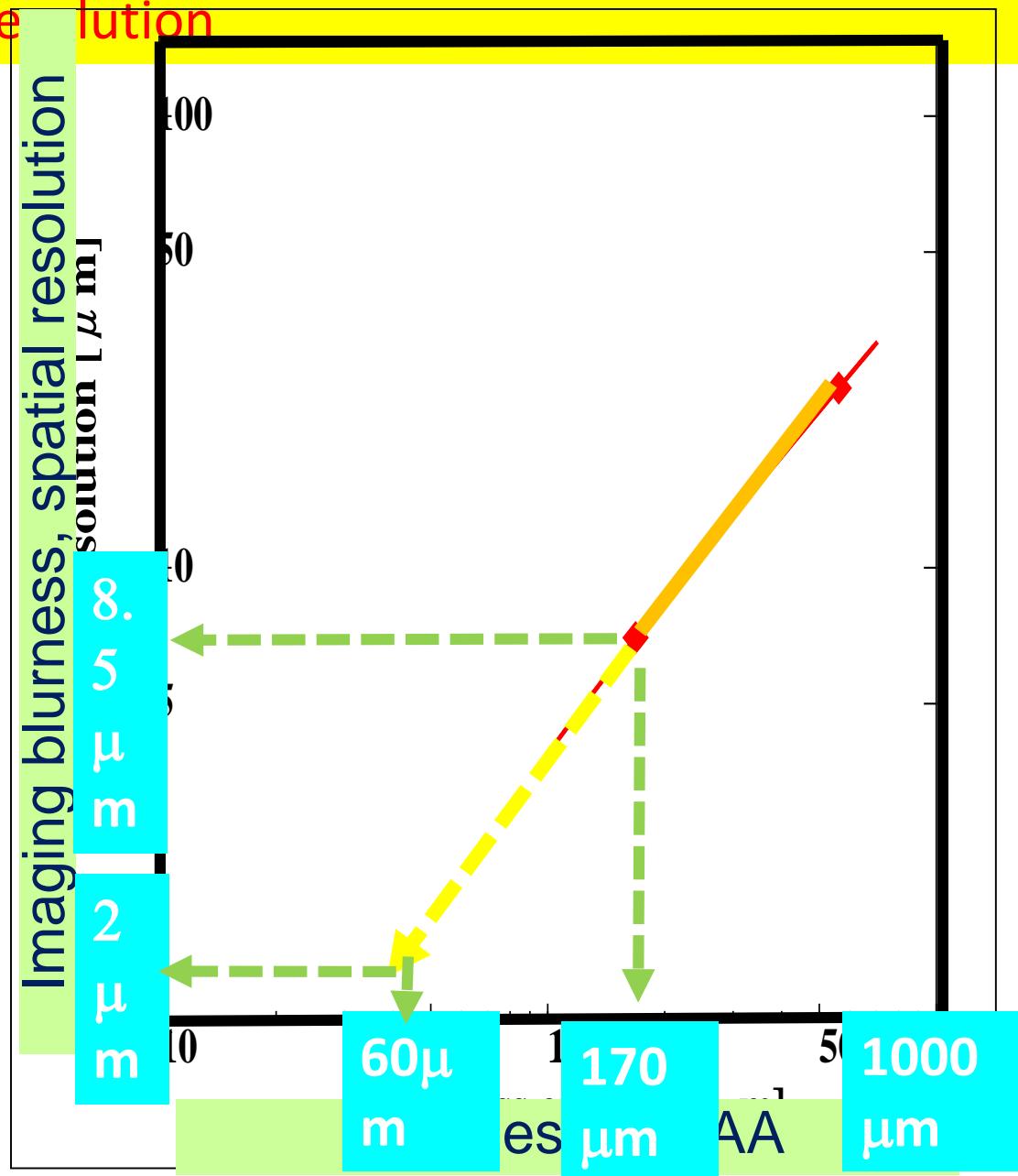


X-ray  
image

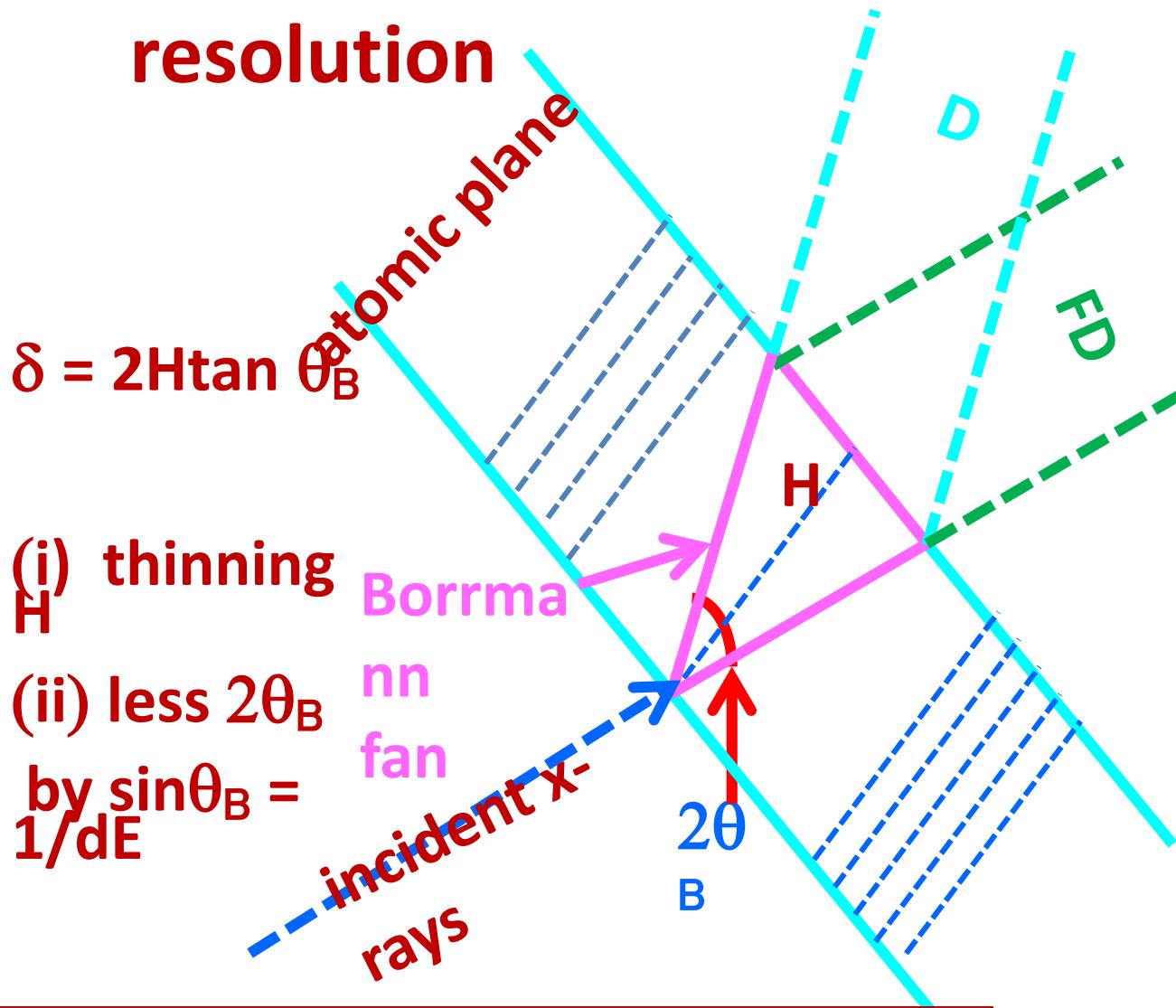
201411  
07



## LAA thickness and spatial resolution



# towards micrometer spatial resolution



approaching to  $2-3\mu\text{m}$  spatial resolution

# Towards future

Needing development

(b)pathology : high  
resolution 3D像

(c)clinical : large FOV

Thank you for listening

measurement of spatial resolution  
without x-ray optics in a hutch BL6C

**20141112**

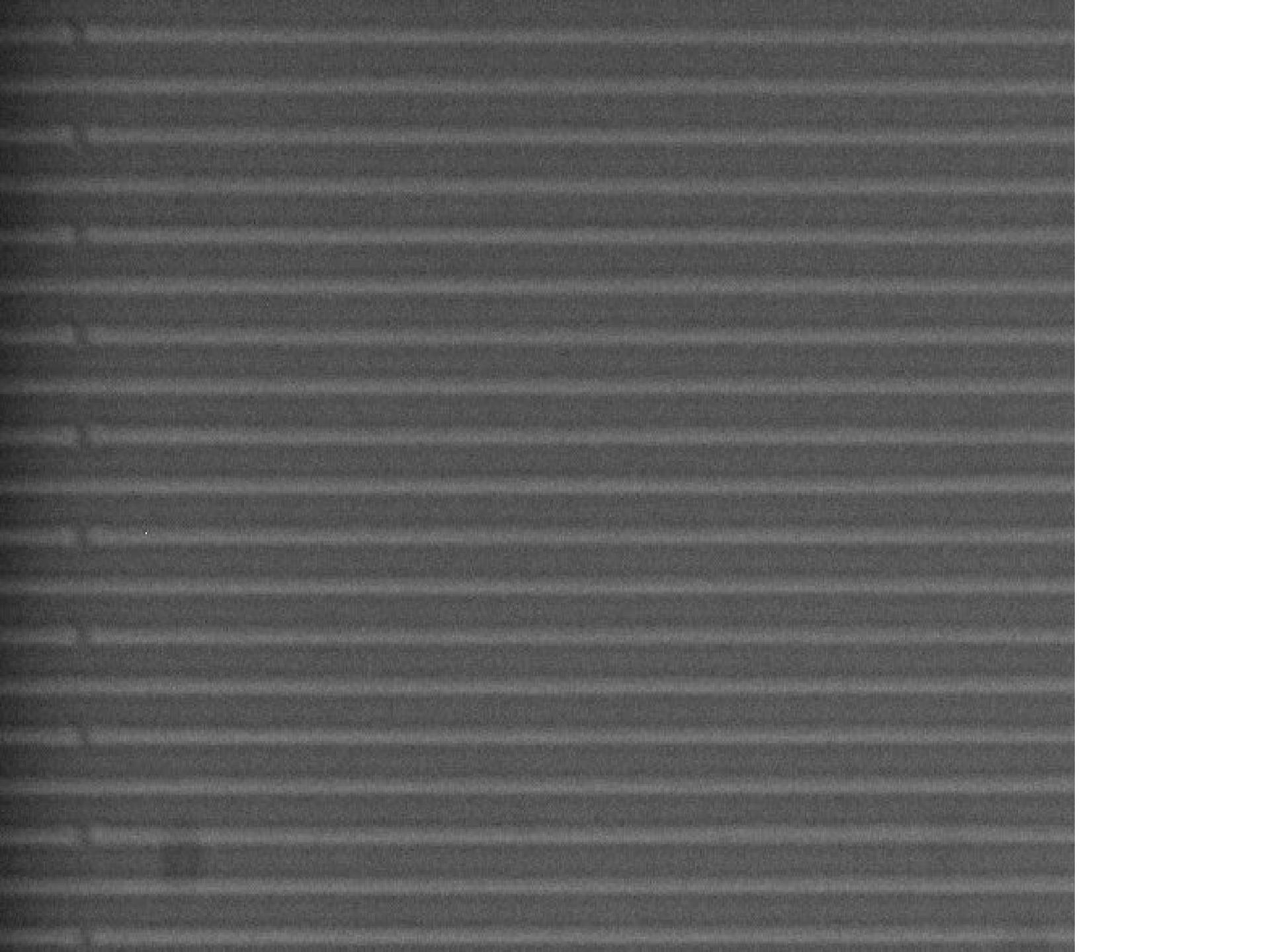
**X-ray image of MTF  
chart  
made of 1 $\mu$ m thick W**

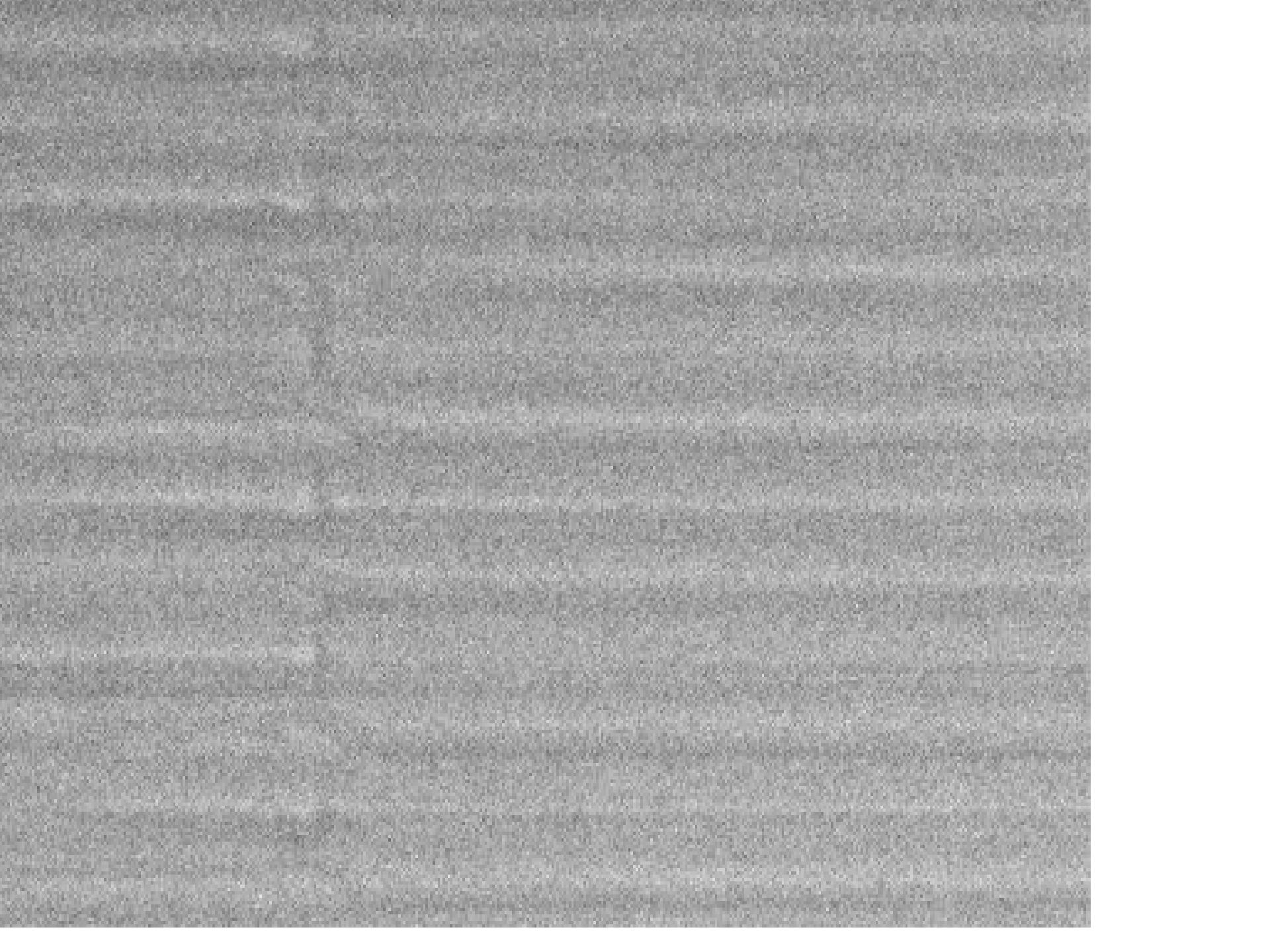
**99.7% transmission**

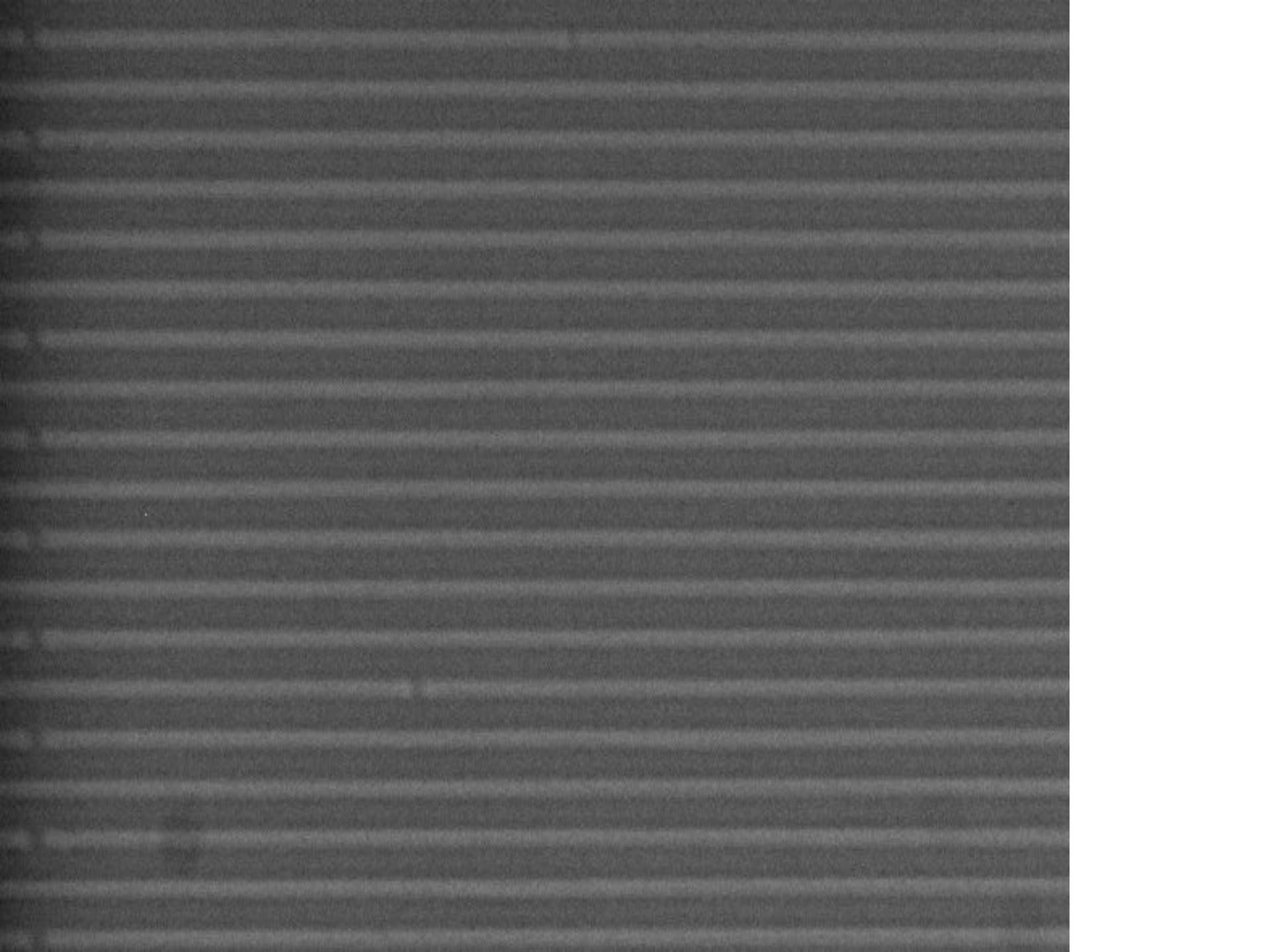
vertical polarization at BL14 by 5T  
horizontal magnetic field at  
Photon Factory

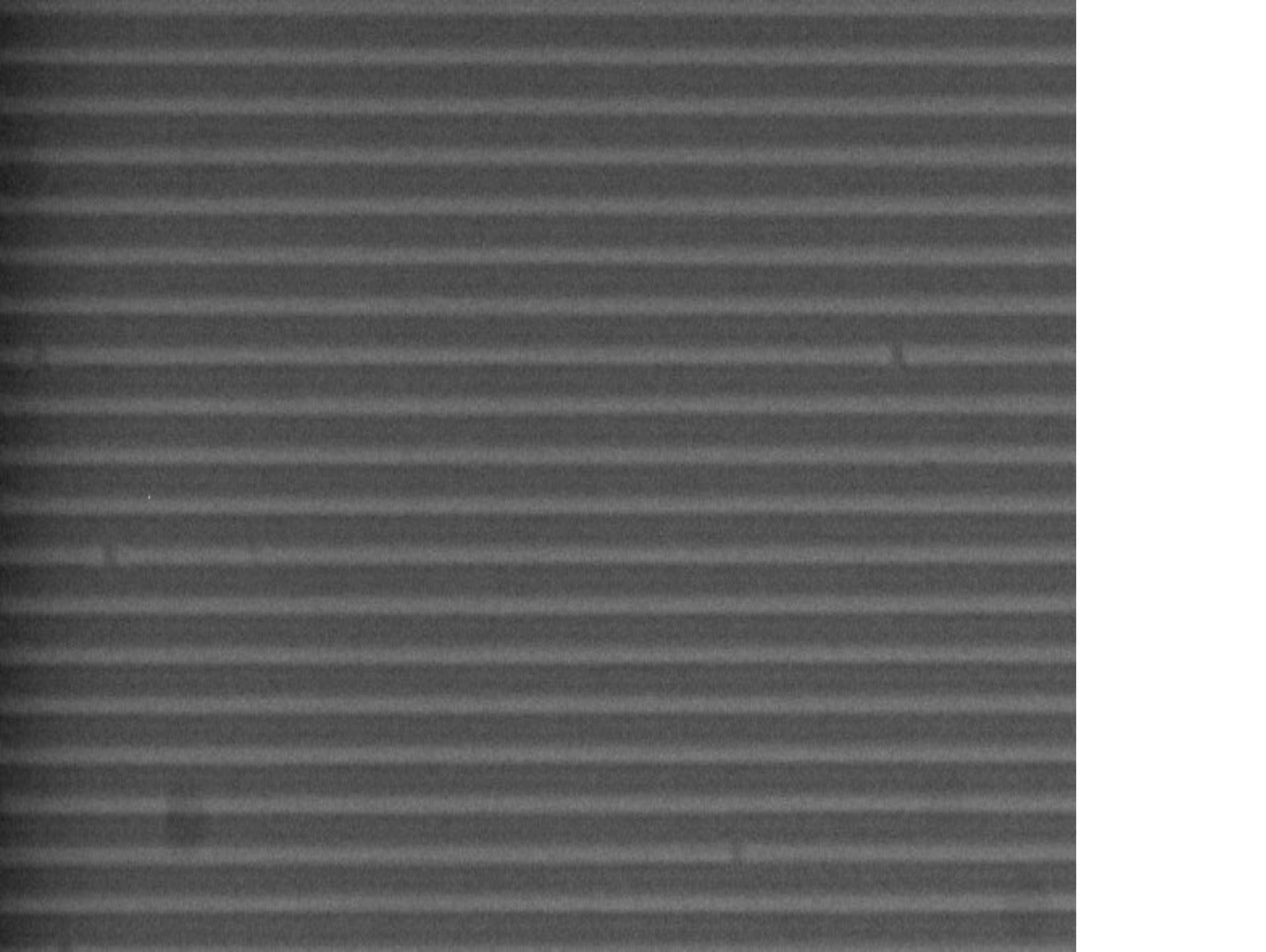
To: in Korea  
Pohang Accelerator Laboratory  
VACTRON Co., Ltd.  
In relation to Leading Industry  
Development for Economic Region Program  
by the Ministry of Trade, Industry & Energy  
(MOTIE),  
Korea Institute for Advancement of  
Technology (KIAT)  
Daegyeong Institute for Regional Program  
Evaluation (DGIRPE)

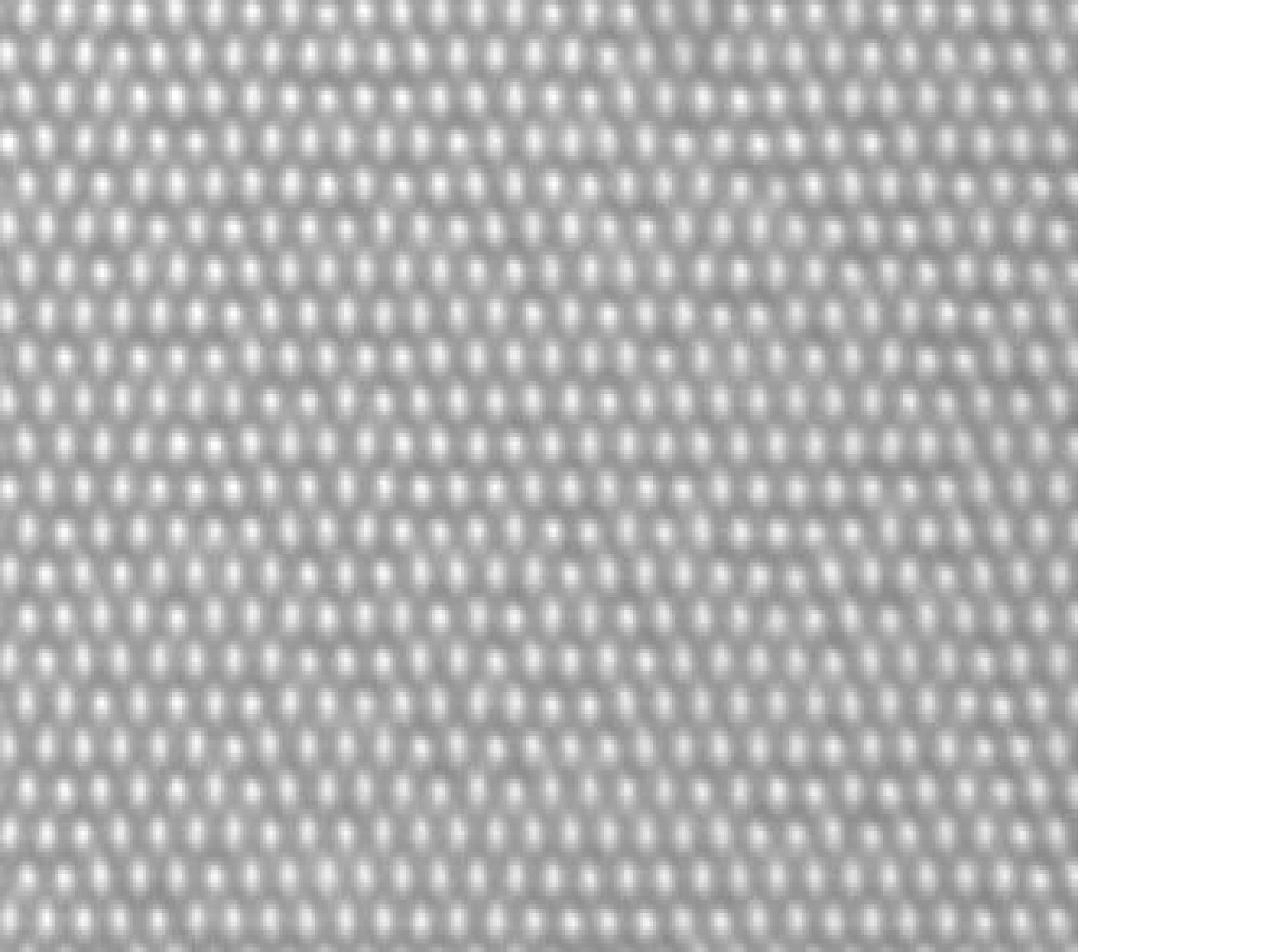
To: in Japan  
MEXT (Ministry of Education, Culture and

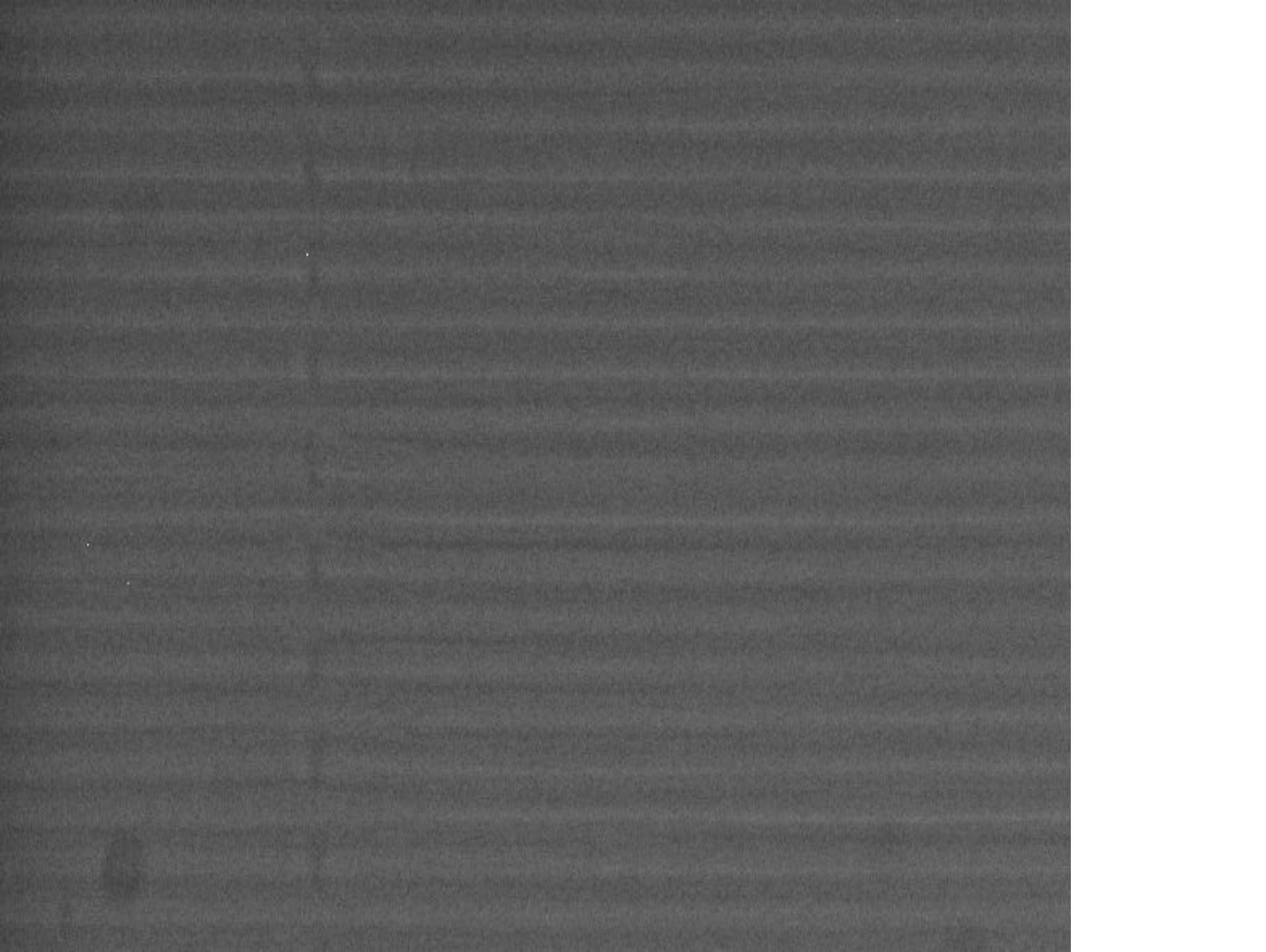












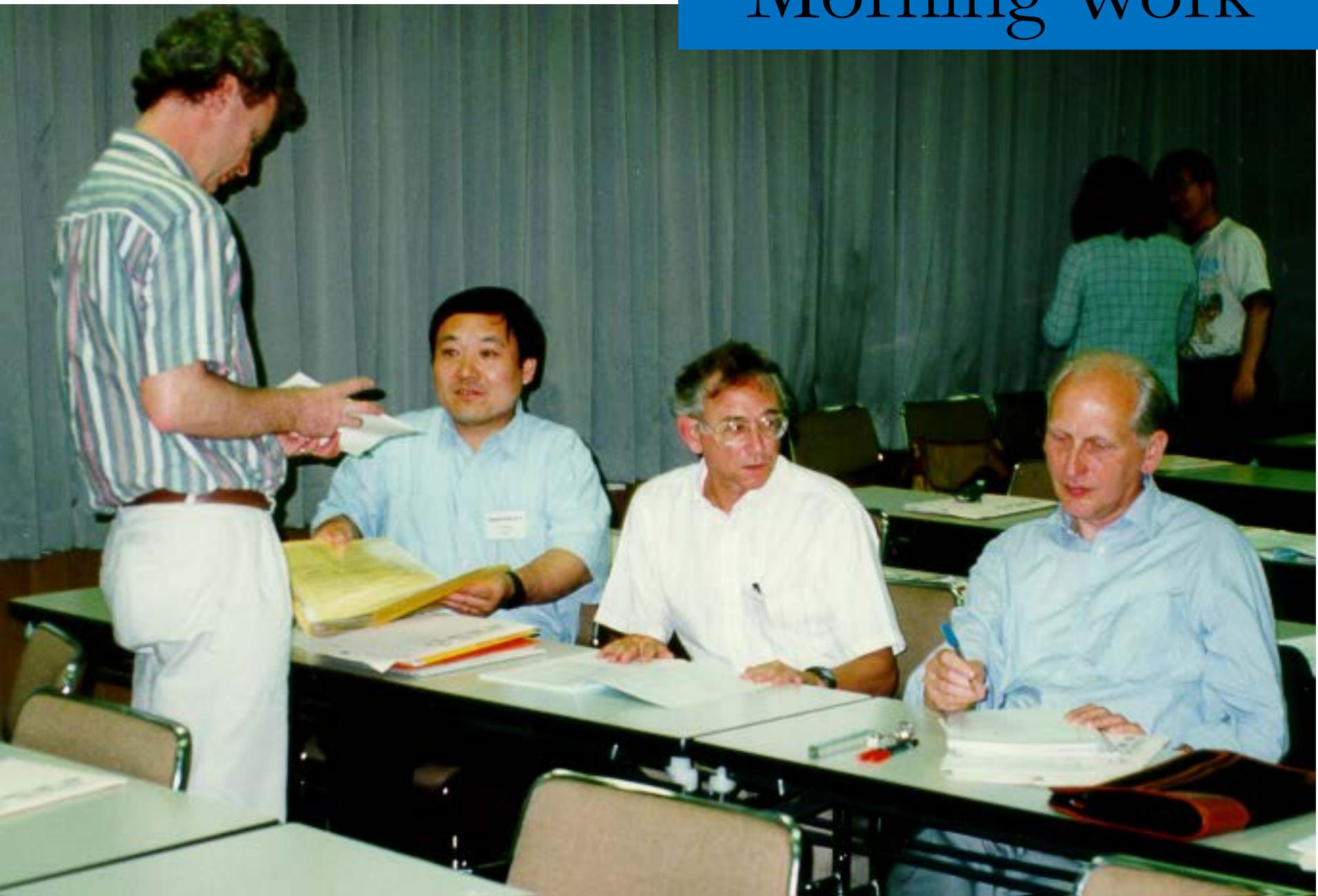
“It's not French wine  
but.....”



# Haga in 1997



# Morning Work

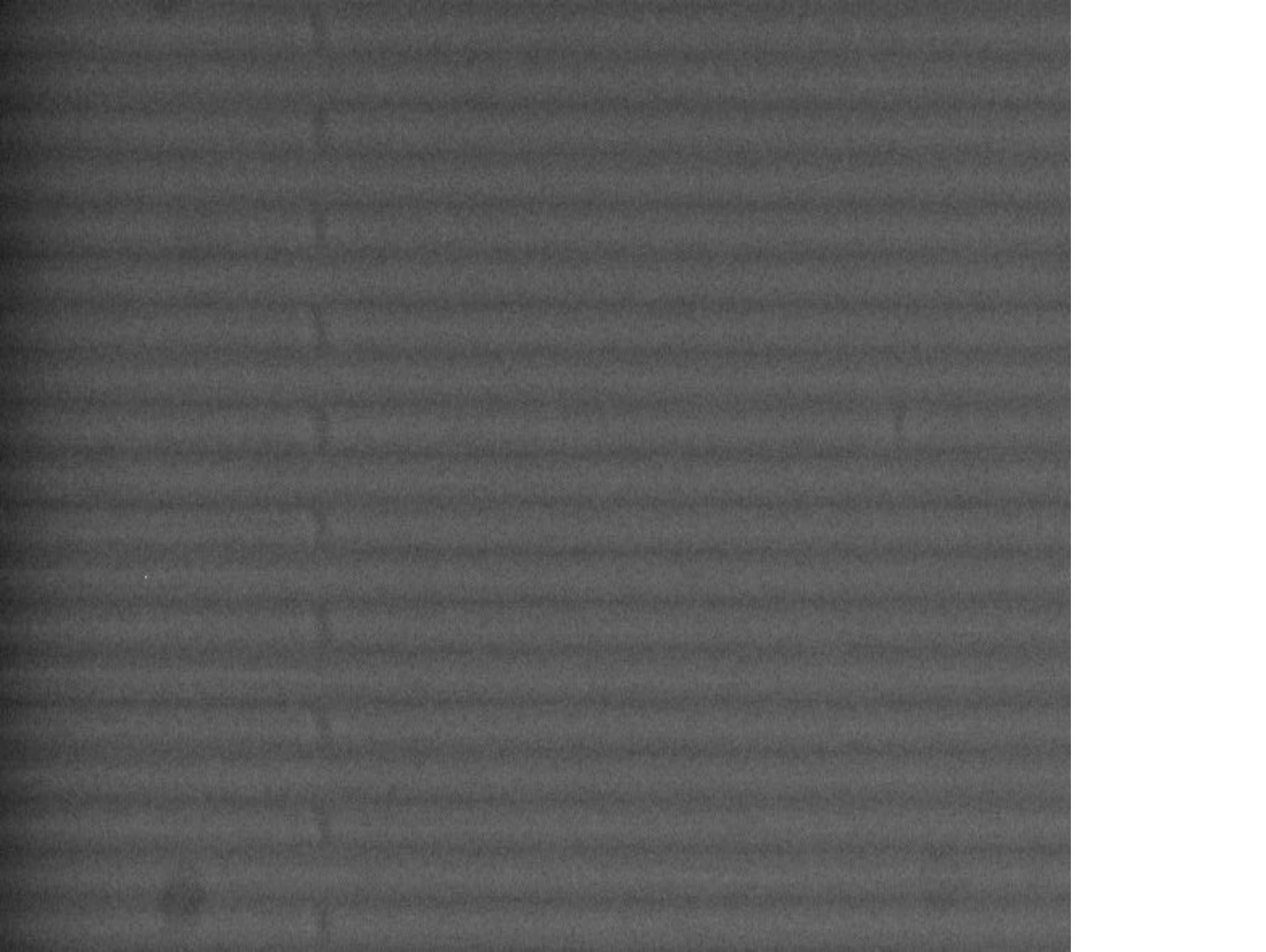


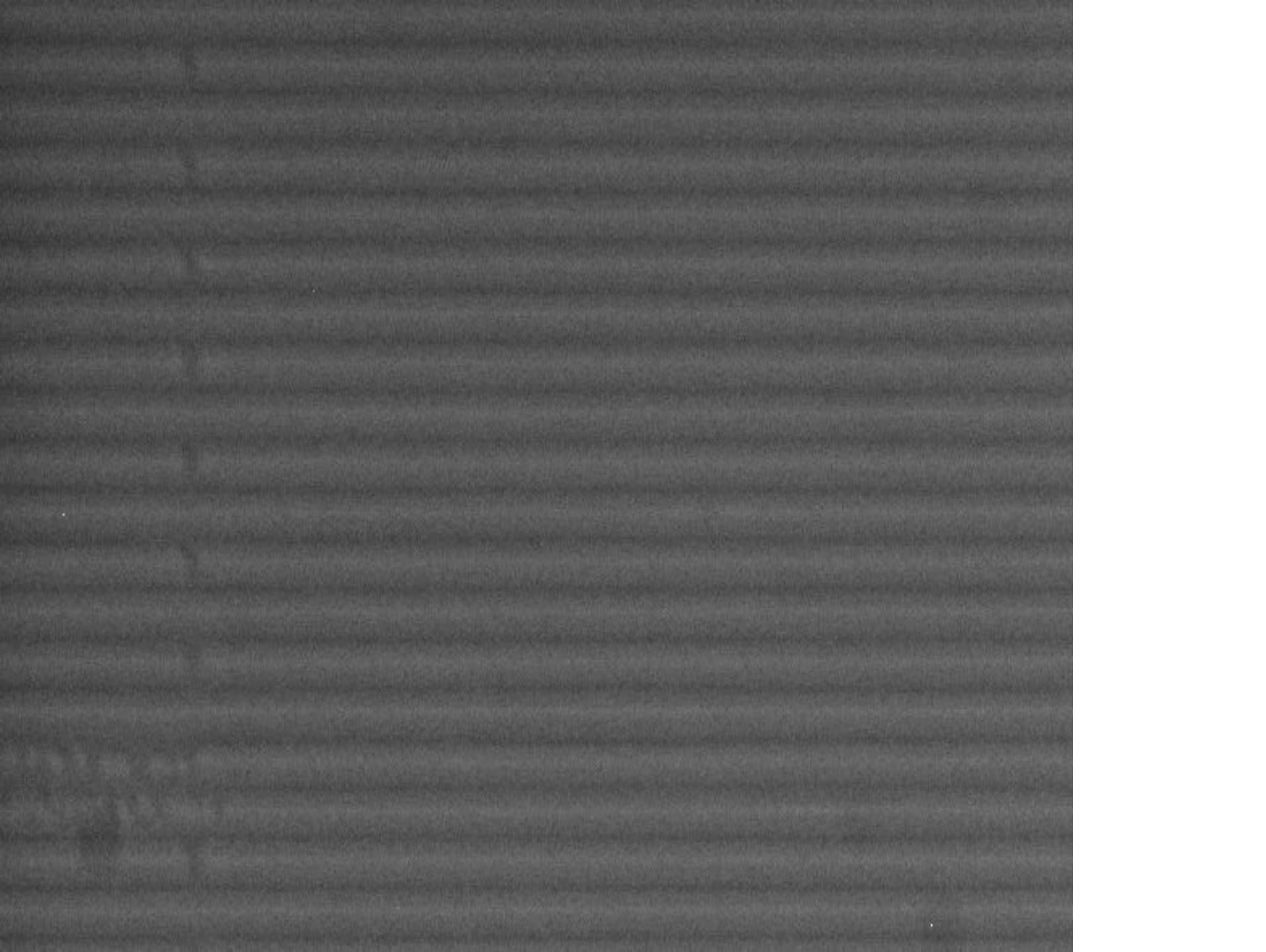
# Afternoon Work

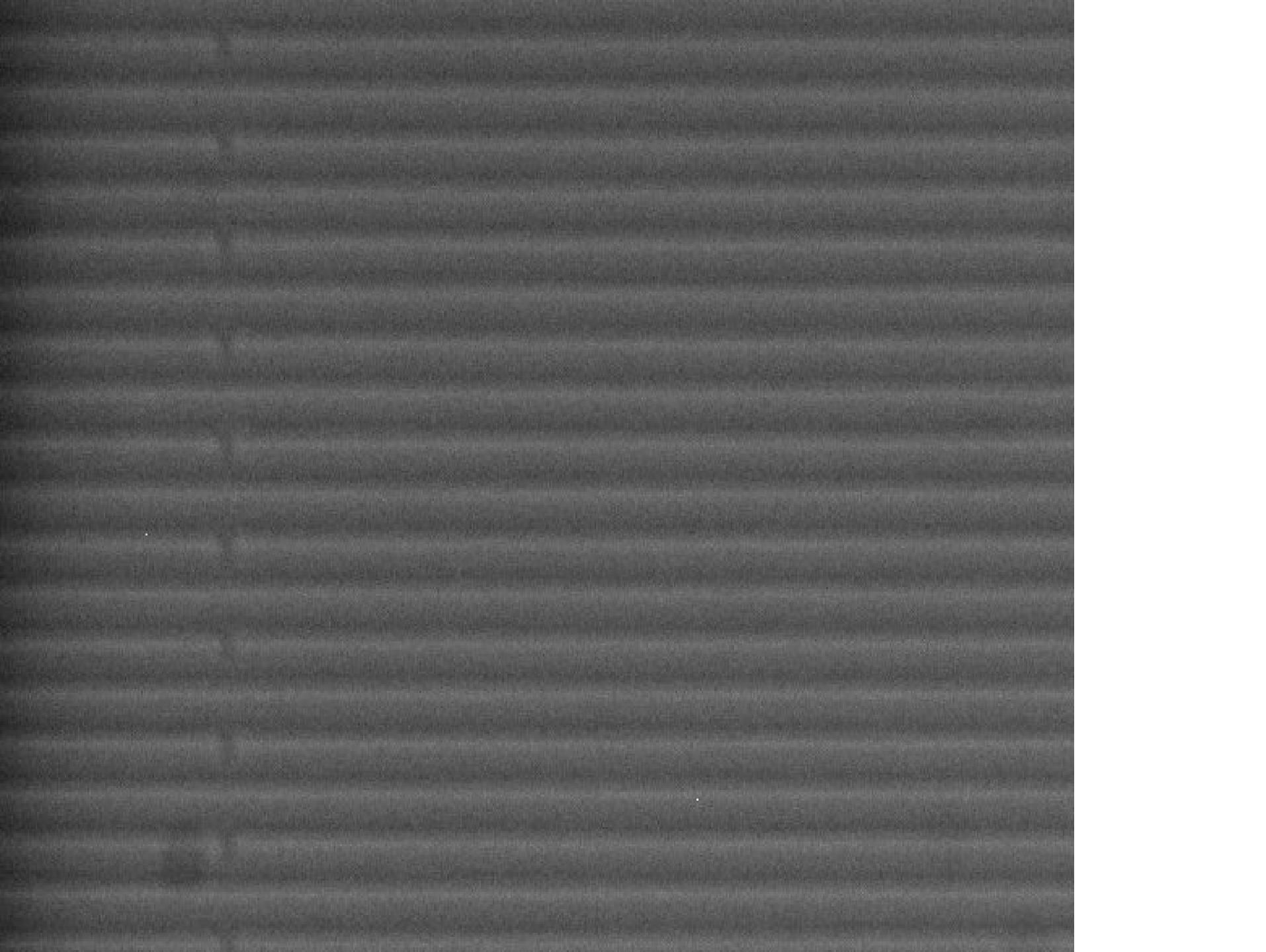


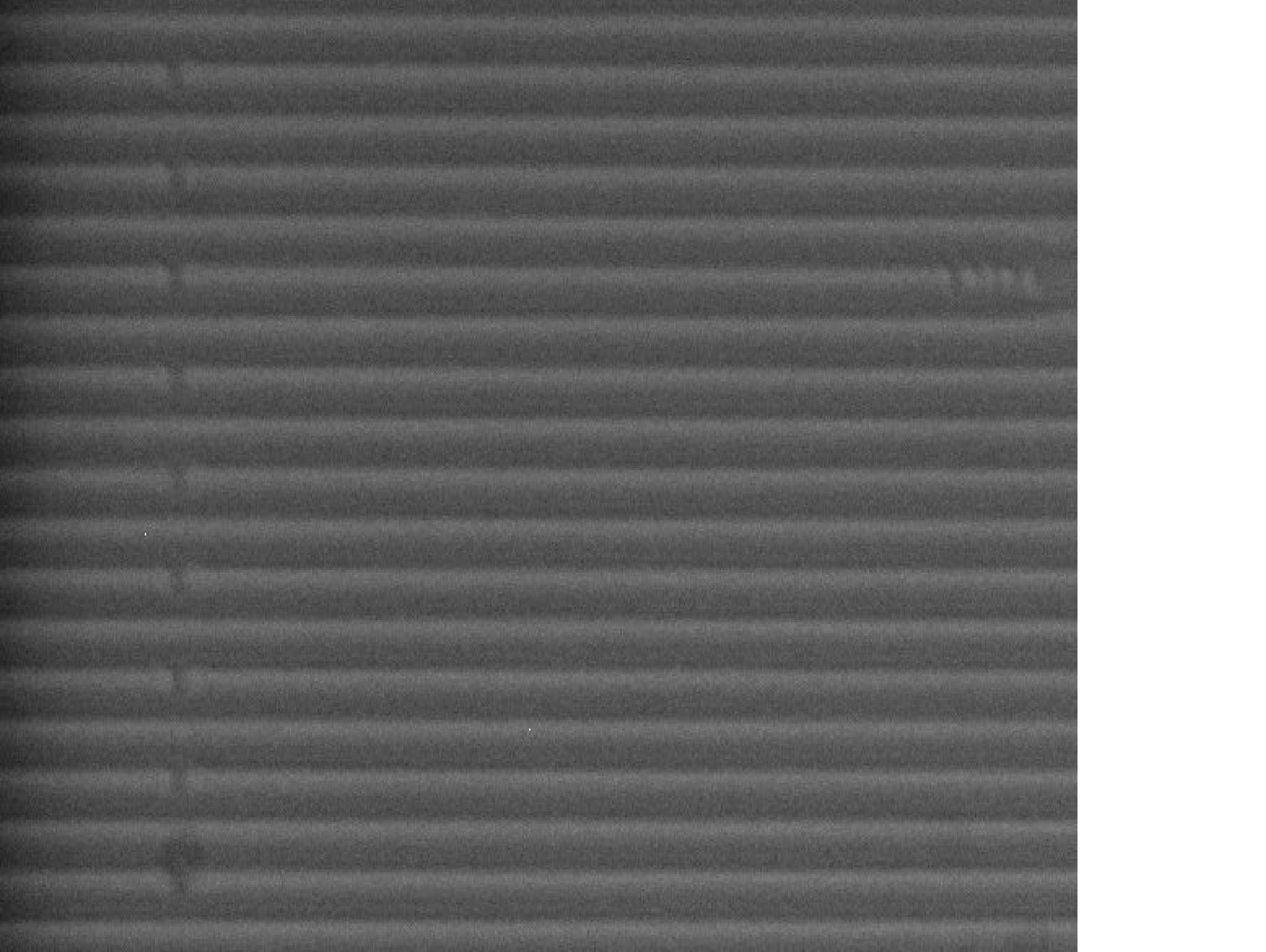
# Evening Work

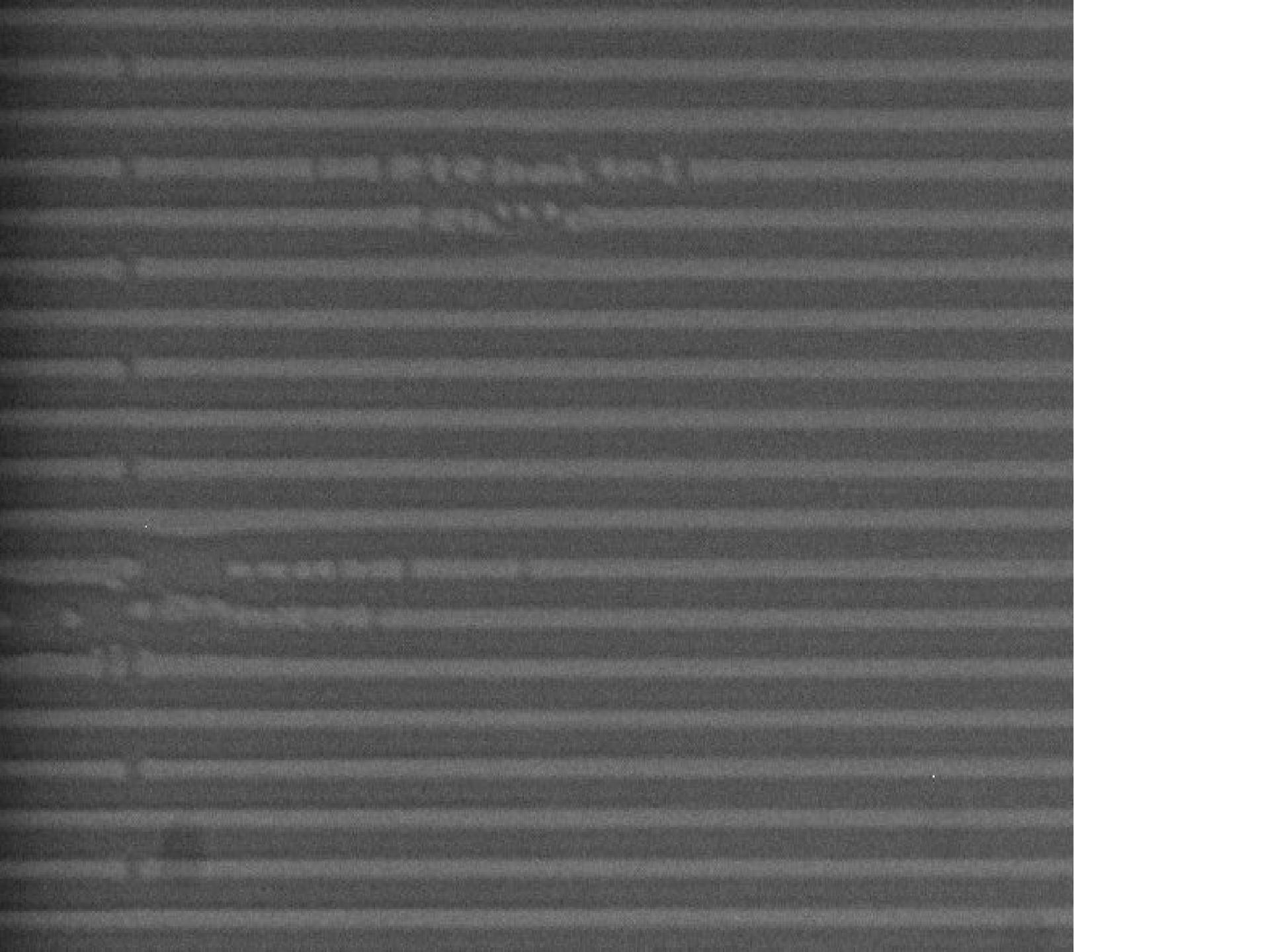


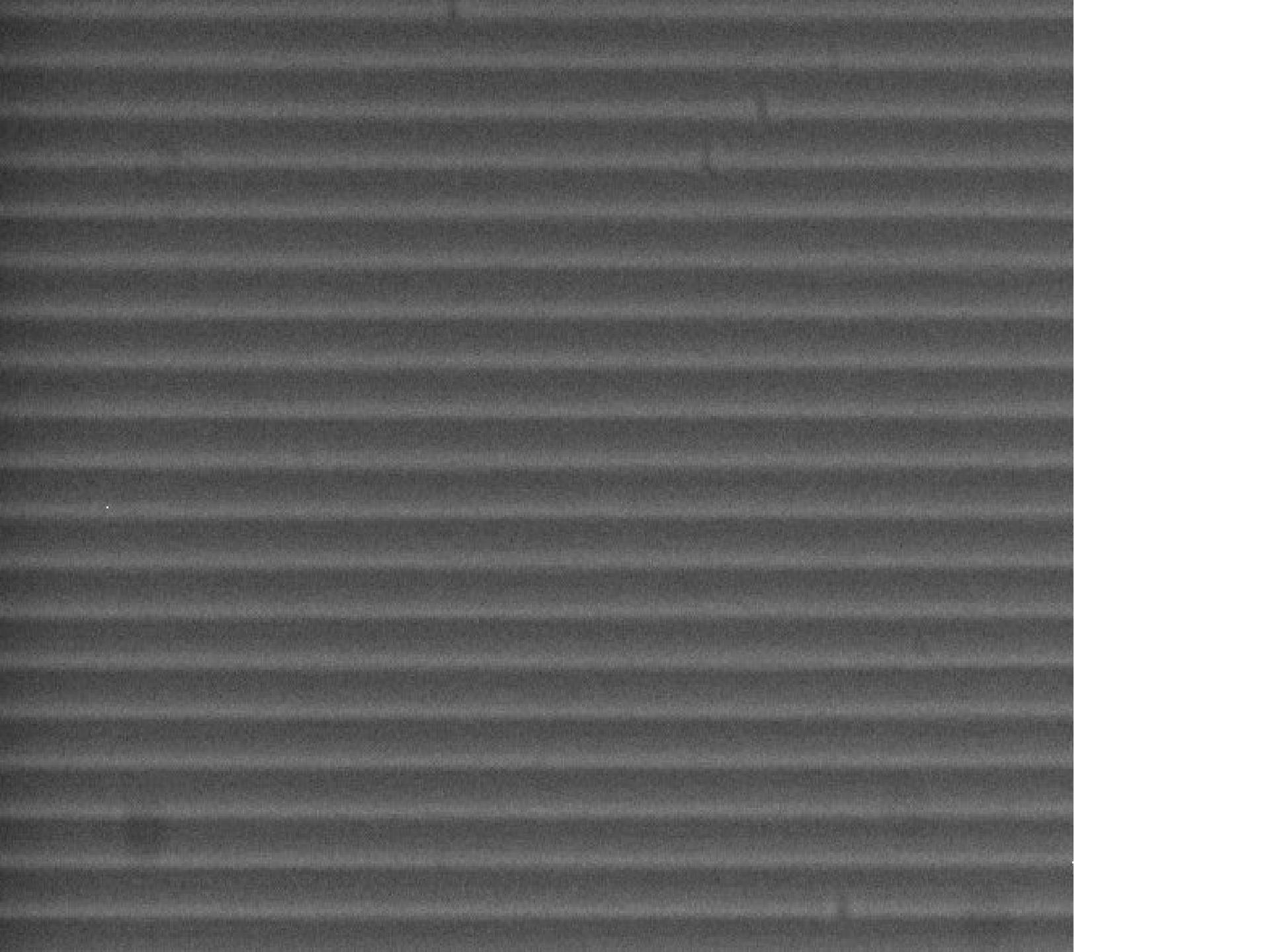


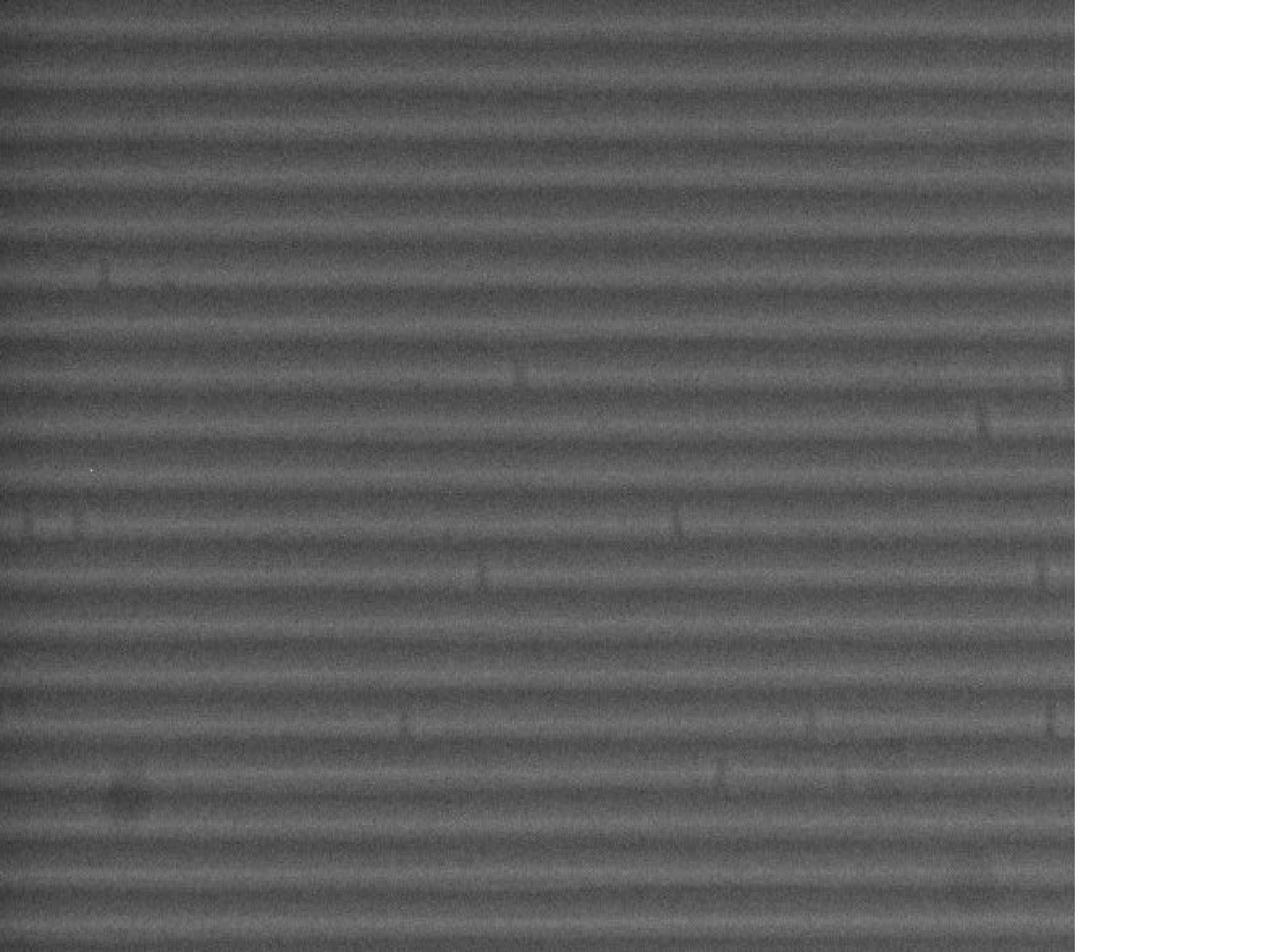




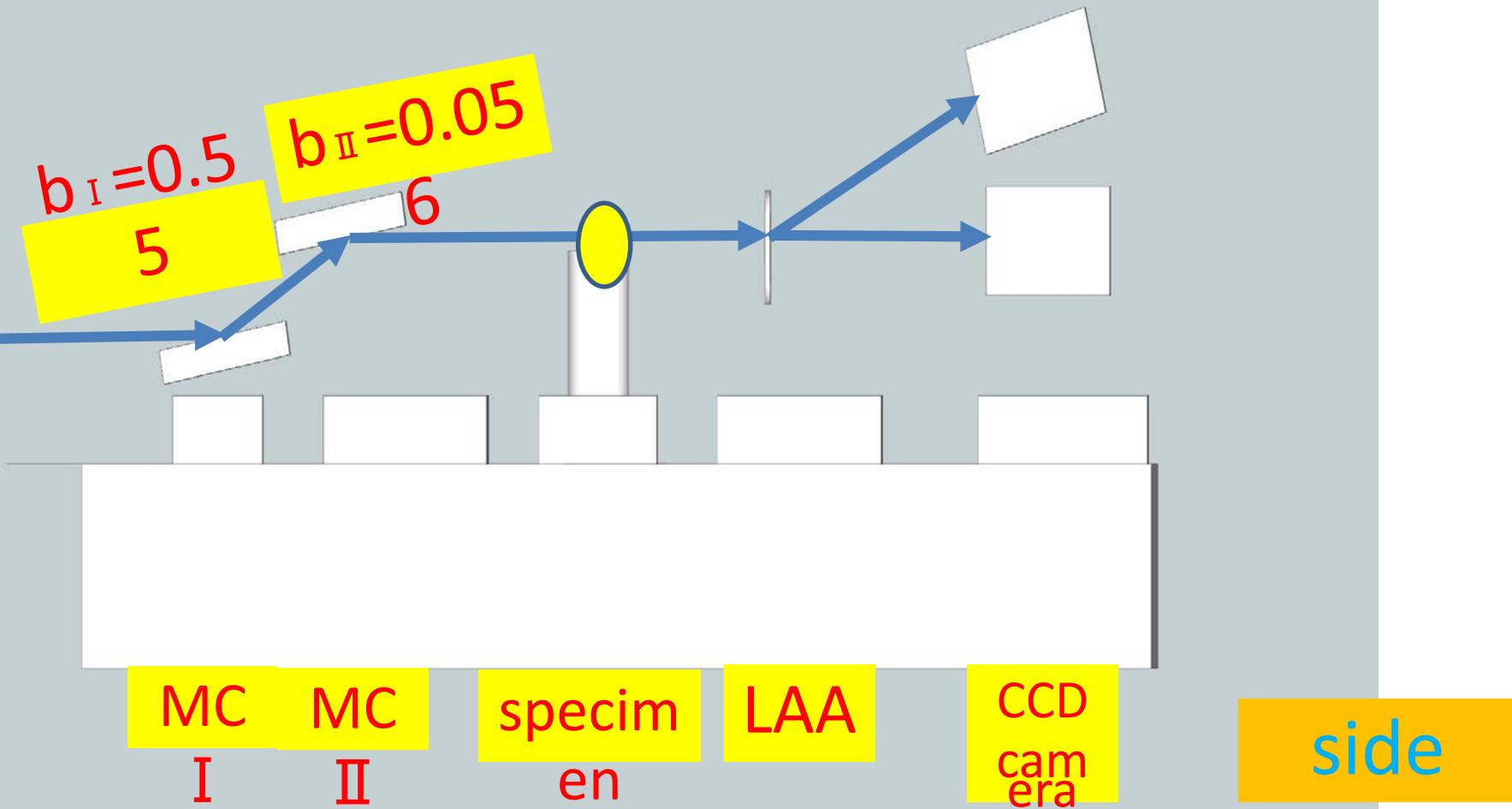






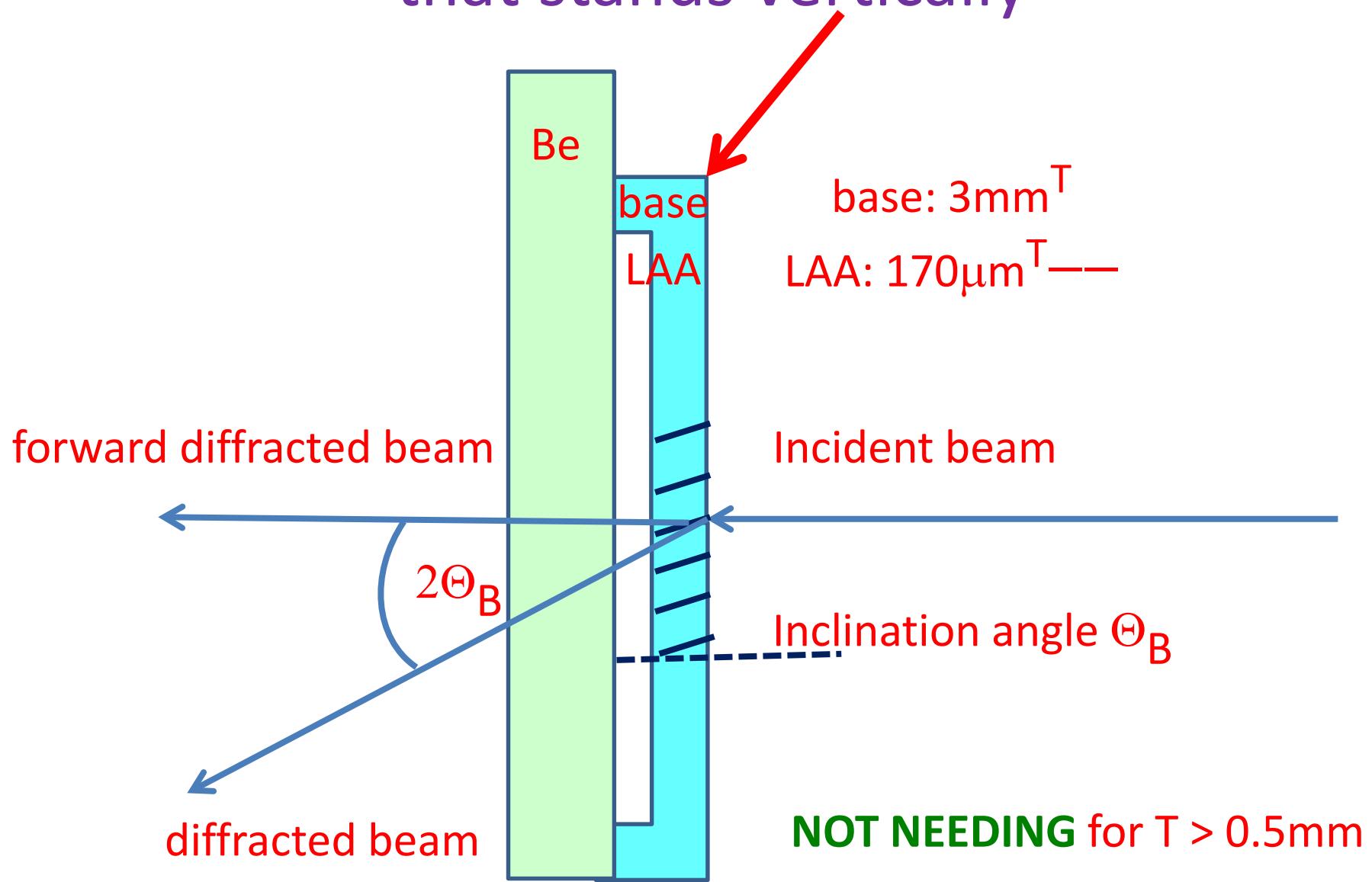


# Monochromator-collimator in an experimental hutch

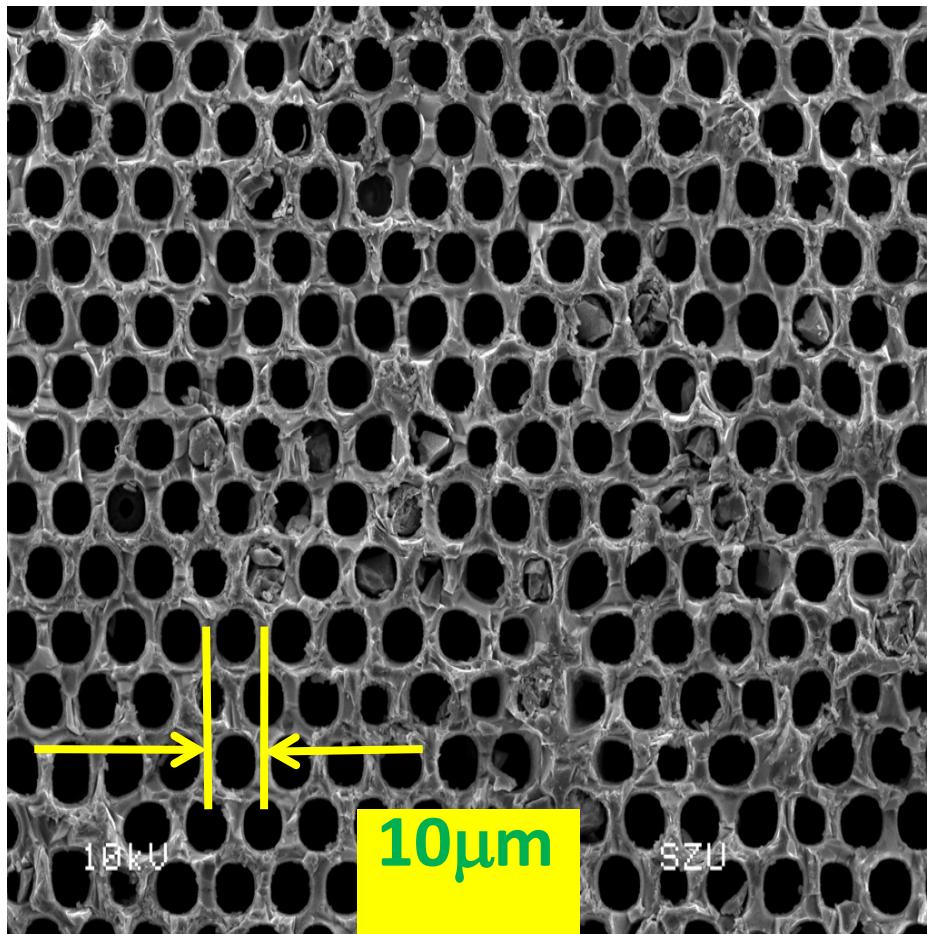


Designed by Dr. DW

# Asymmetric LAA that stands vertically



detector made by  
Professor Guo  
Jinchuan



# Daigo in 1992



Mrs. Rubenstein serenading Ed!



phase grating made  
by  
Professor Guo  
Jinchuan

20 $\mu\text{m}$



201411

07

silicon

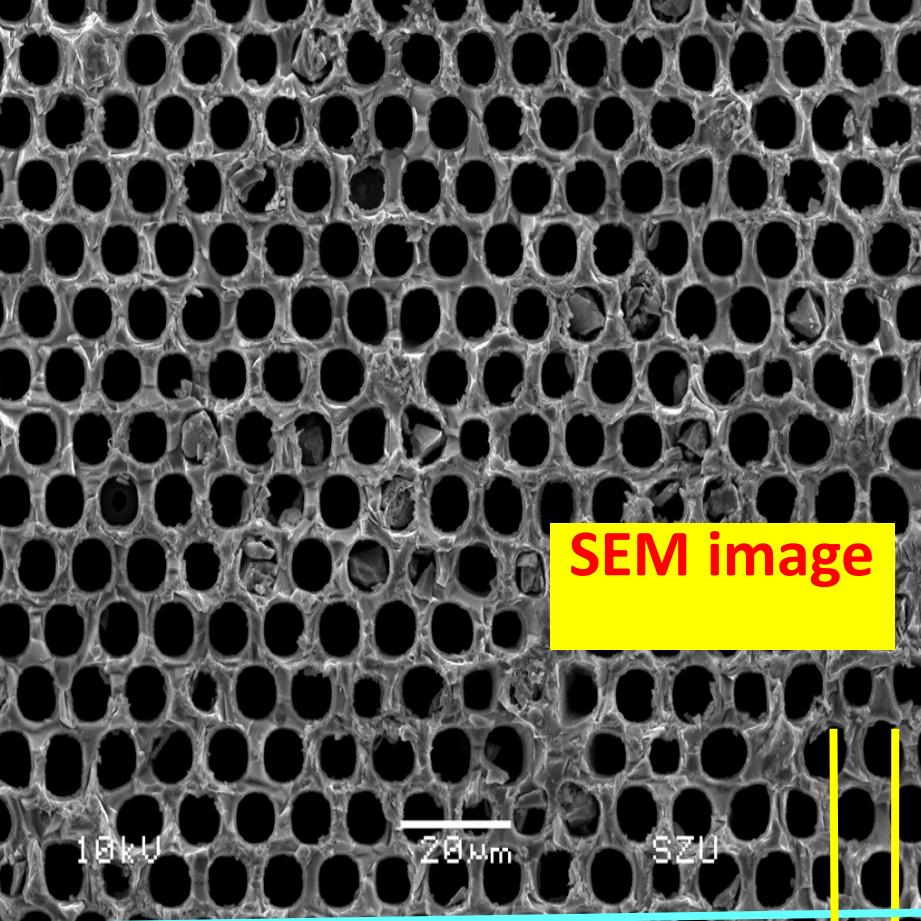
no crystal

100 $\mu\text{m}$

250 $\mu\text{m}$

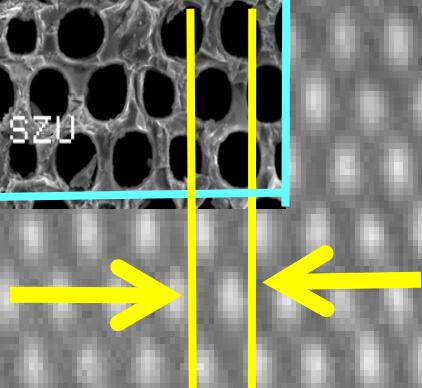
direct view with 35keV X-rays

detector made by  
Professor Guo  
Jinchuan

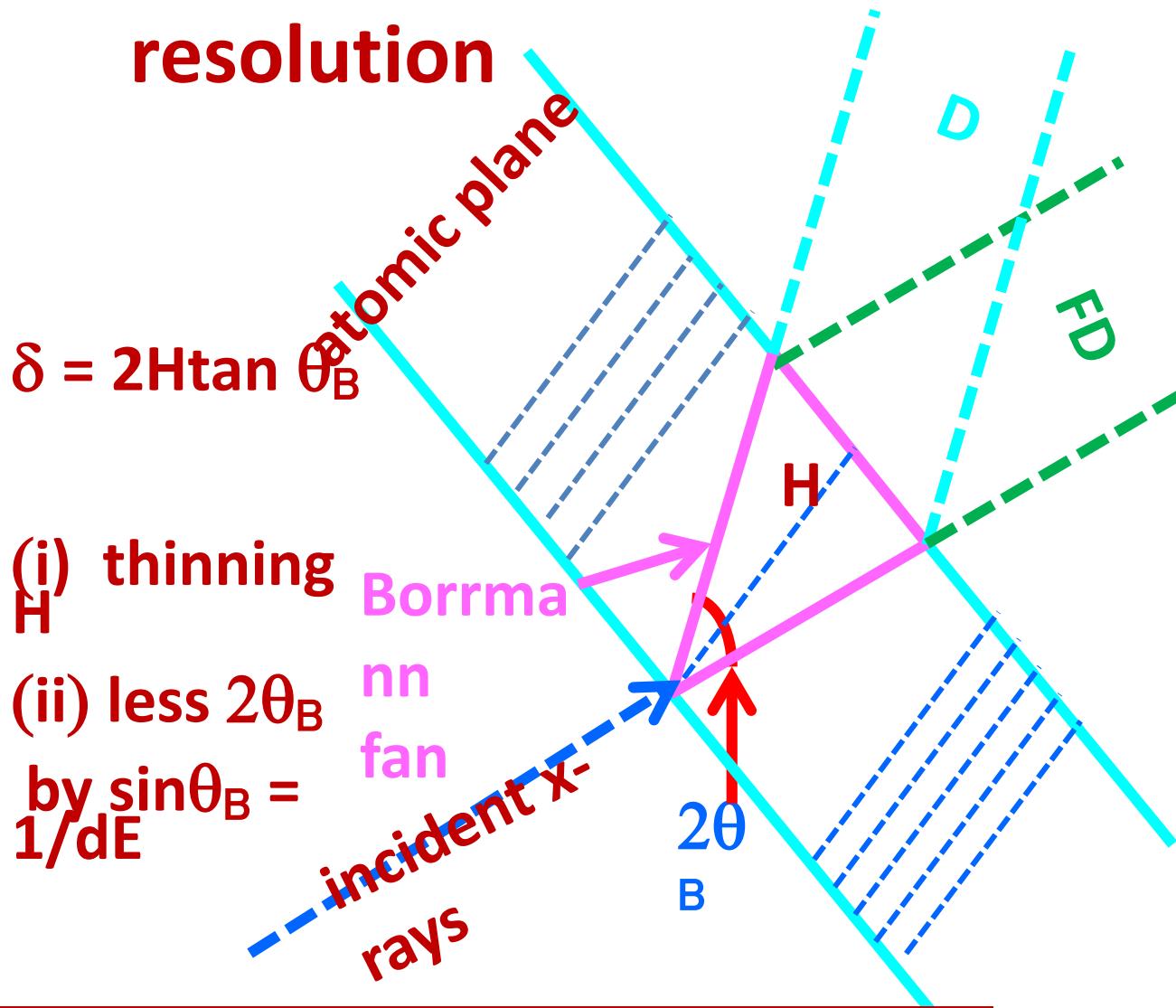


X-ray  
image

201411  
07



# towards micrometer spatial resolution



approaching to  $2-3\mu\text{m}$  spatial resolution

JIMA RC-02B

$1\mu\text{m}^T$  W

left:

without MC

right:

with MC

31keV

150 $\mu\text{m}$

15 $\mu\text{m}$

10 $\mu\text{m}$

7 $\mu\text{m}$

15 $\mu\text{m}$

10 $\mu\text{m}$

7 $\mu\text{m}$

thank

- Organizing committee, Professor Kulipanov,  
Old friends
- Special thanks to Professor Vazina
- Today