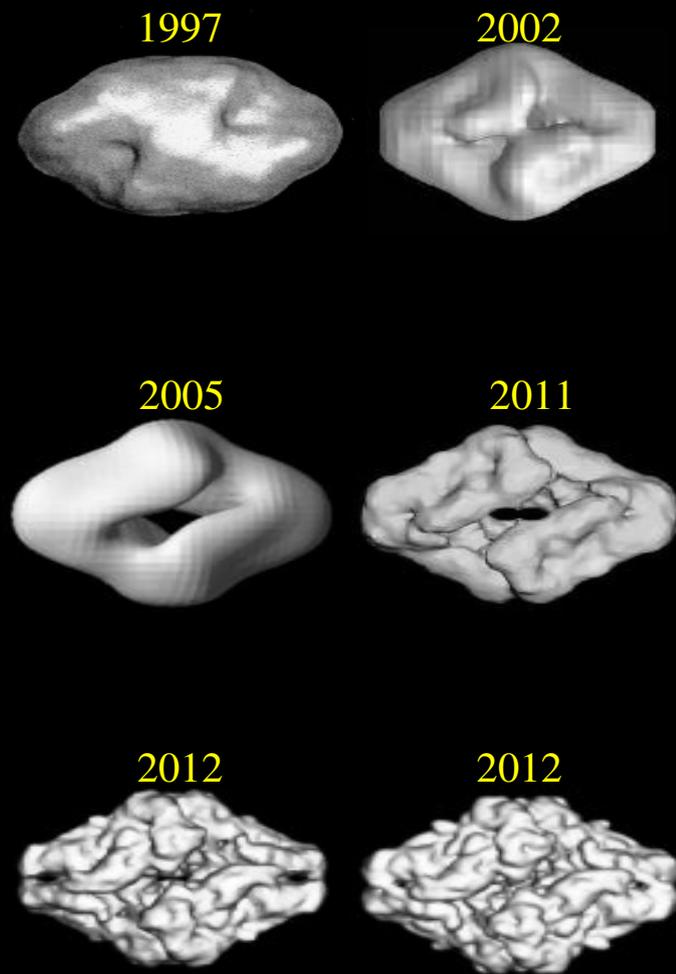
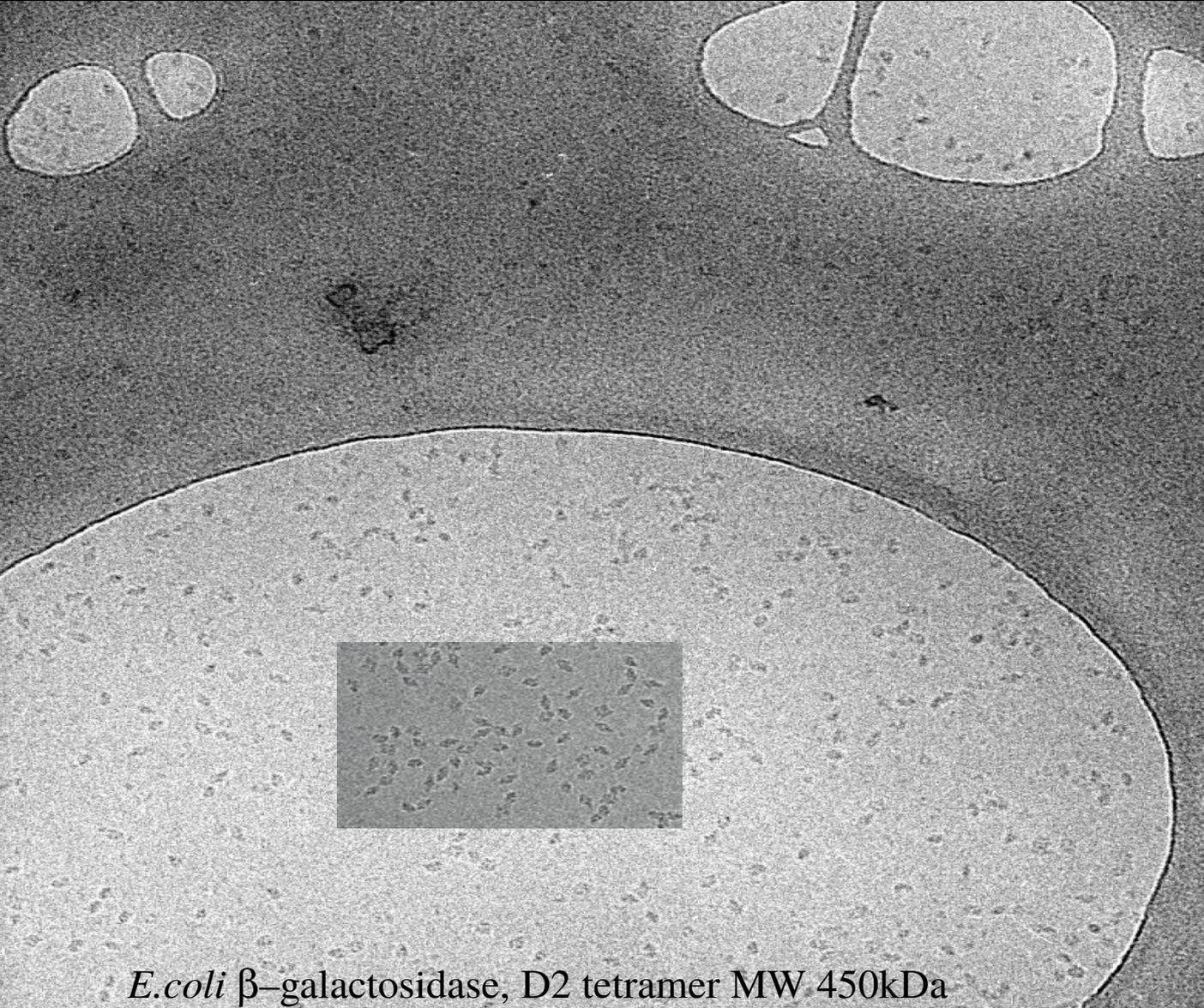


# Direct detector technology

Greg McMullan  
MRC-LMB  
Cambridge, U.K.

MRC

Laboratory of  
Molecular Biology



*E. coli*  $\beta$ -galactosidase, D2 tetramer MW 450kDa

# Change is not all due to detectors

- Microscopes have improved.
- Reconstruction programs have improved.
- Improved computing allows things to be done correctly
- Lots more data
- Improved samples
- Expectations

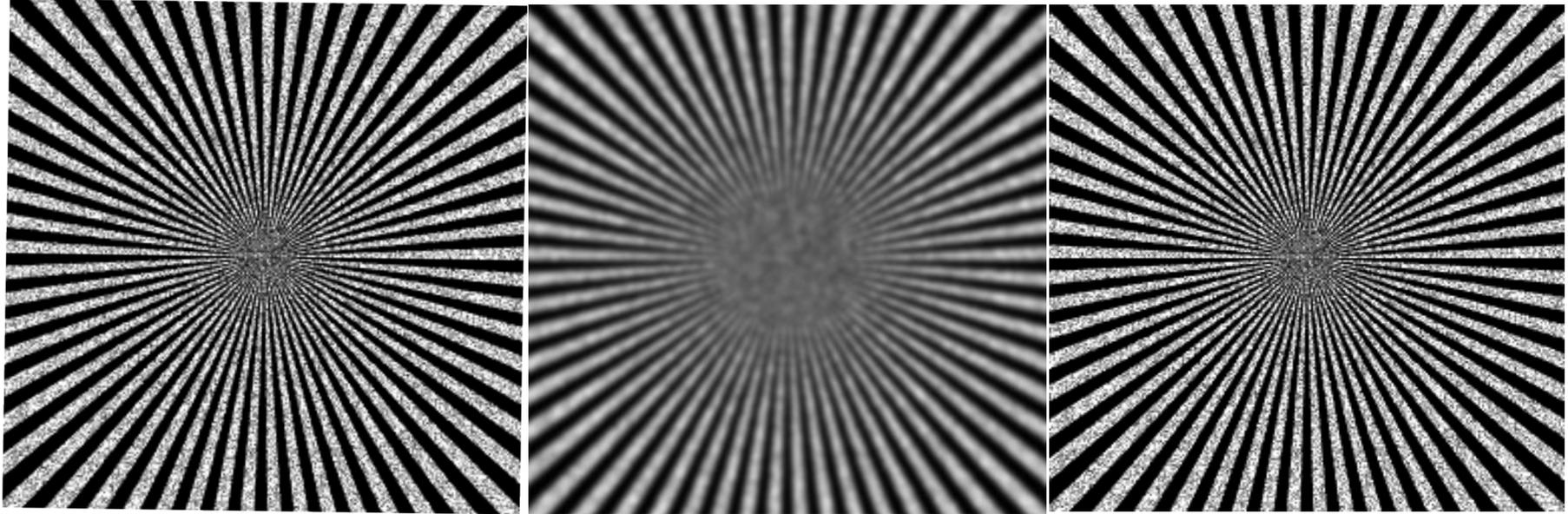
# Detecting electrons

- Should be easy
  - 300 keV versus  $1/40$  eV room temperature
- Electrons are light and interact strongly:
  - Only lose small amount of energy each time
  - Long tracks
  - Random direction change

# MTF/DQE

- DQE is the important property
- Unlike CCD detectors drop in MTF doesn't lose information (acts as low pass filter)

# MTF: Poor MTF is not necessarily a problem



Original

MTF blur

Sharpened

# Problems getting high DQE

- Variable energy deposition (Landau noise)
- Long trajectories with energy loss concentrated at the end as it slows down

# Energy loss from high energy particles

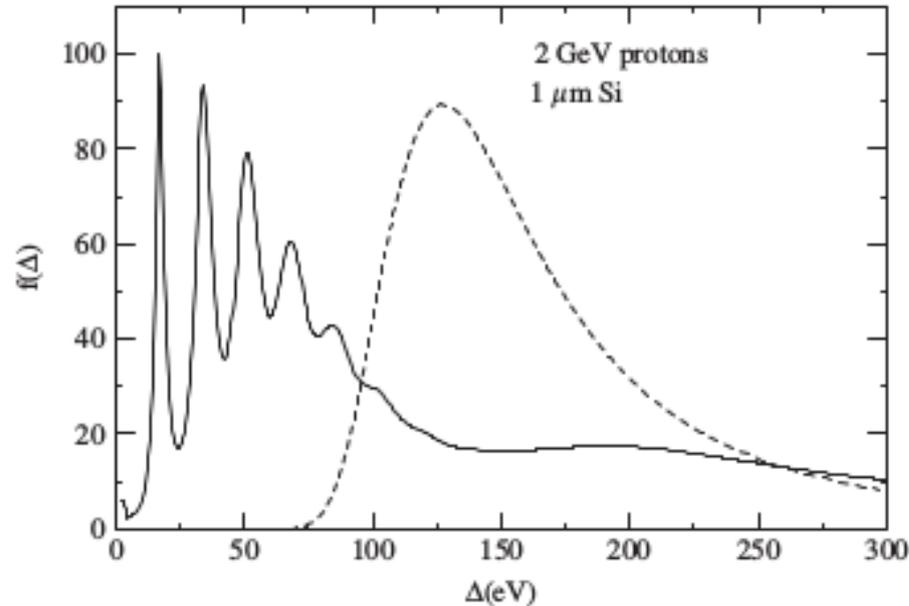
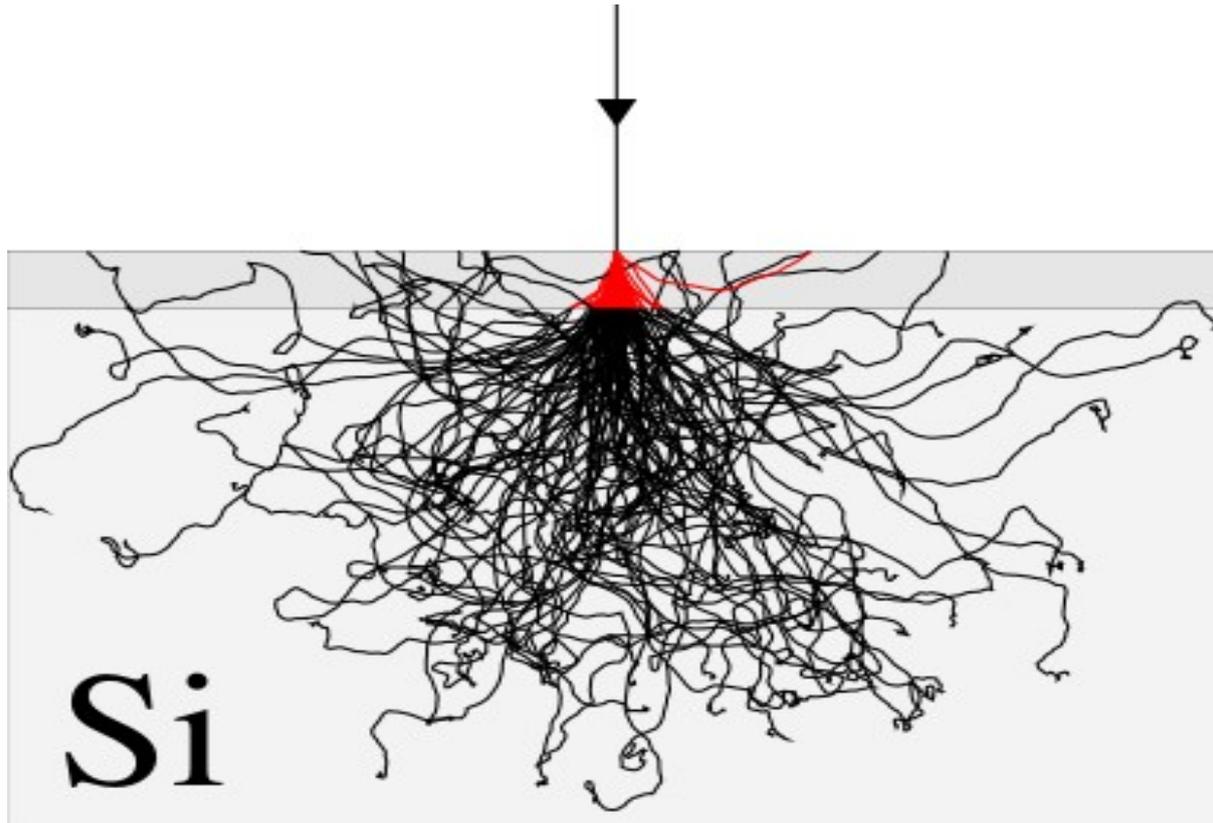


Fig. 2. Straggling in 1  $\mu\text{m}$  of Si, compared to the Landau function. The Bethe-Bloch mean energy loss is  $\langle \Delta \rangle = 400 \text{ eV}$ .

# 300 keV electron trajectories in silicon



Monte Carlo simulation

# Two problems:

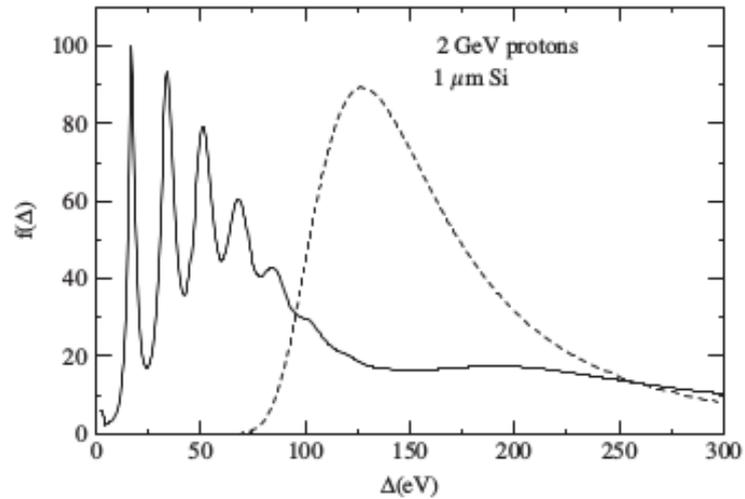
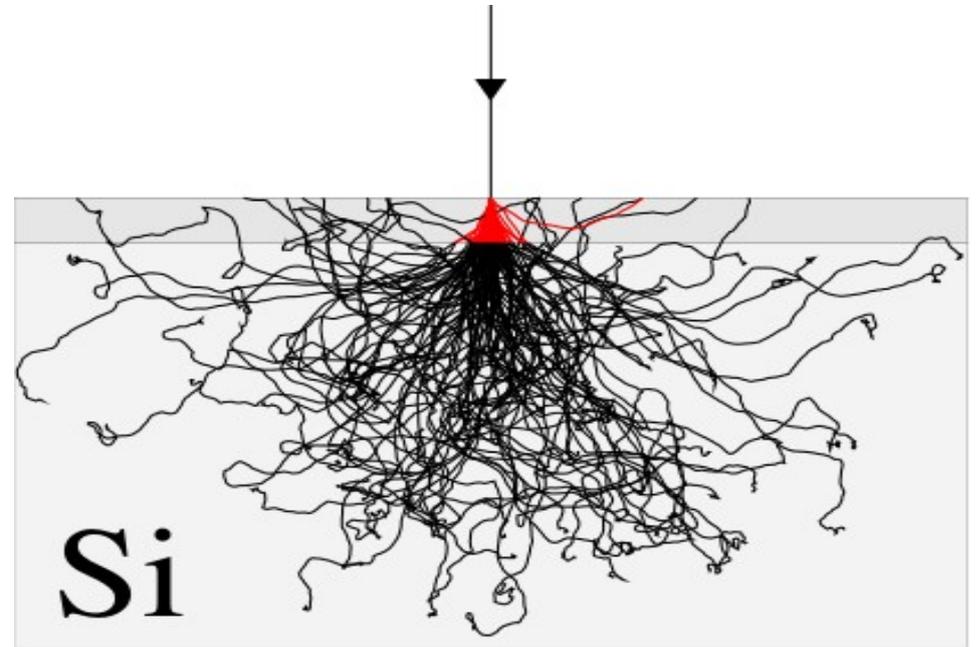
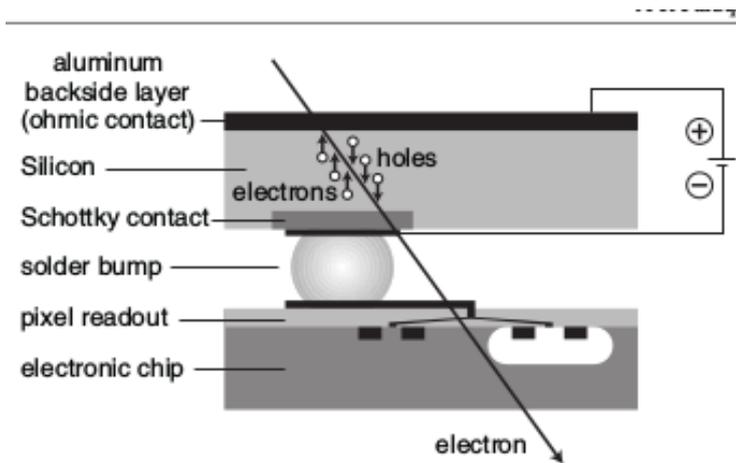


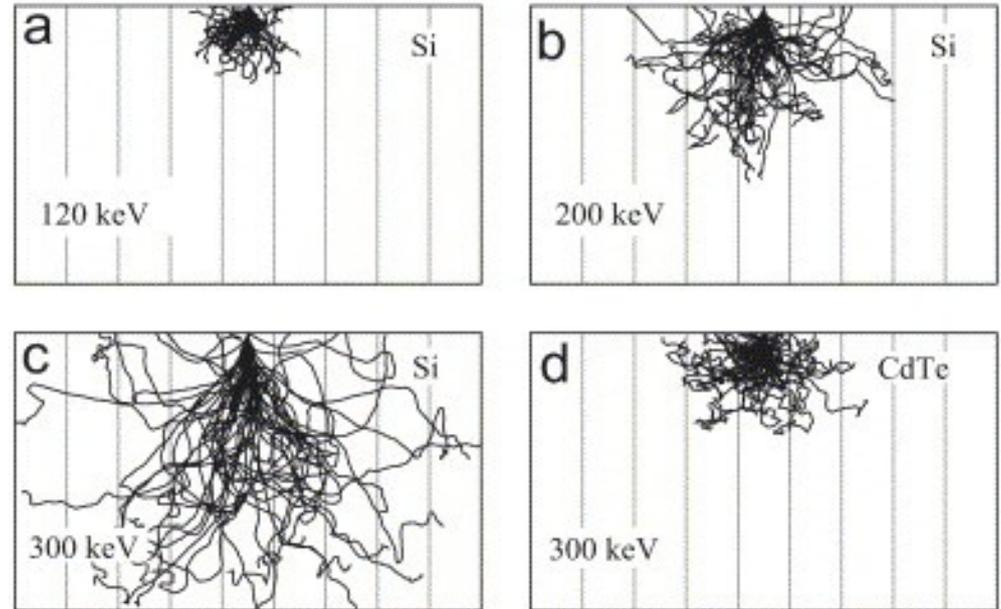
Fig. 2. Straggling in 1  $\mu\text{m}$  of Si, compared to the Landau function. The Bethe-Bloch mean energy loss is  $\langle \Delta \rangle = 400 \text{ eV}$ .



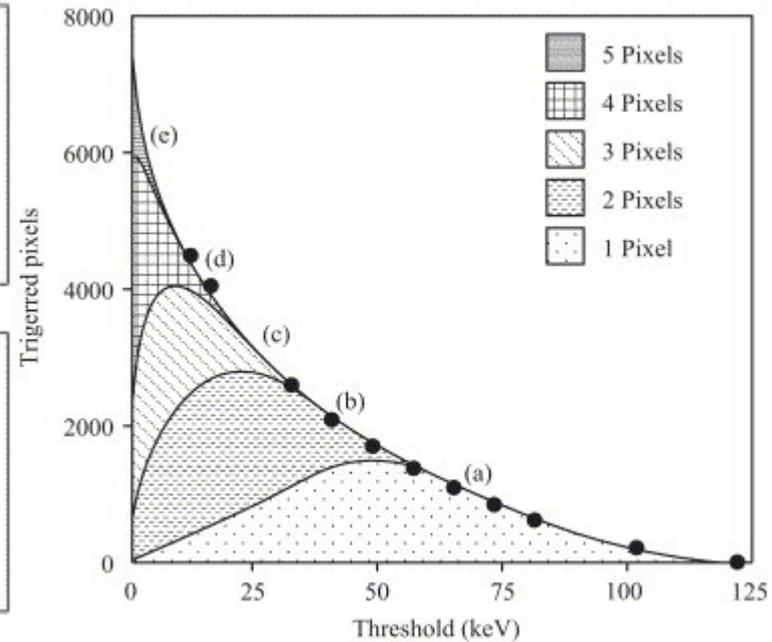
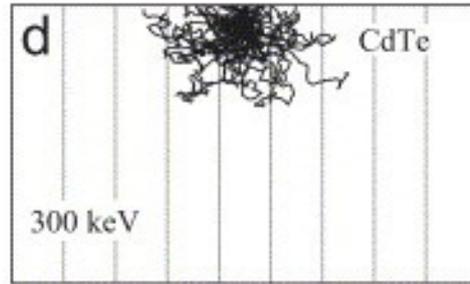
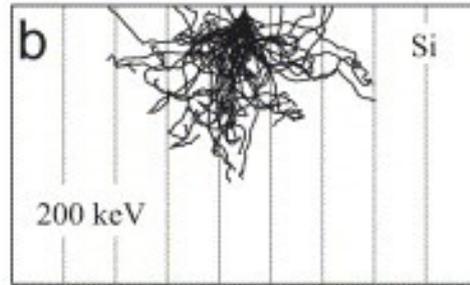
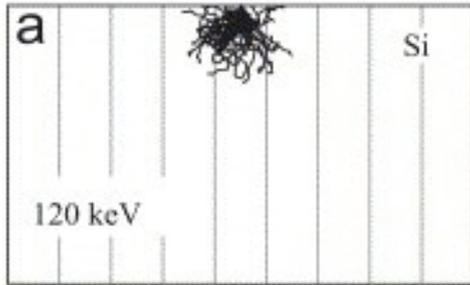
# Solution 1: Hybrid pixel detectors



**Figure 4.** Schematic diagram of a single Medipix2 pixel with the sensor and readout separated by a bump bond. The aluminium layer forming the ohmic contact is only a fraction of  $1\ \mu\text{m}$ , making it more convenient for the detection of low energy electrons.



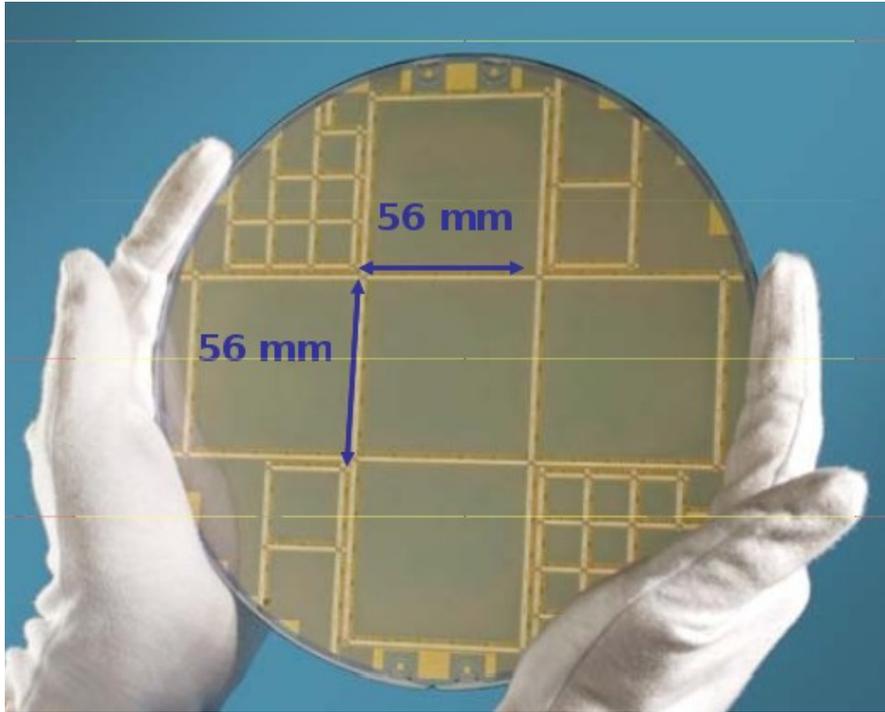
# Problems with solution 1:



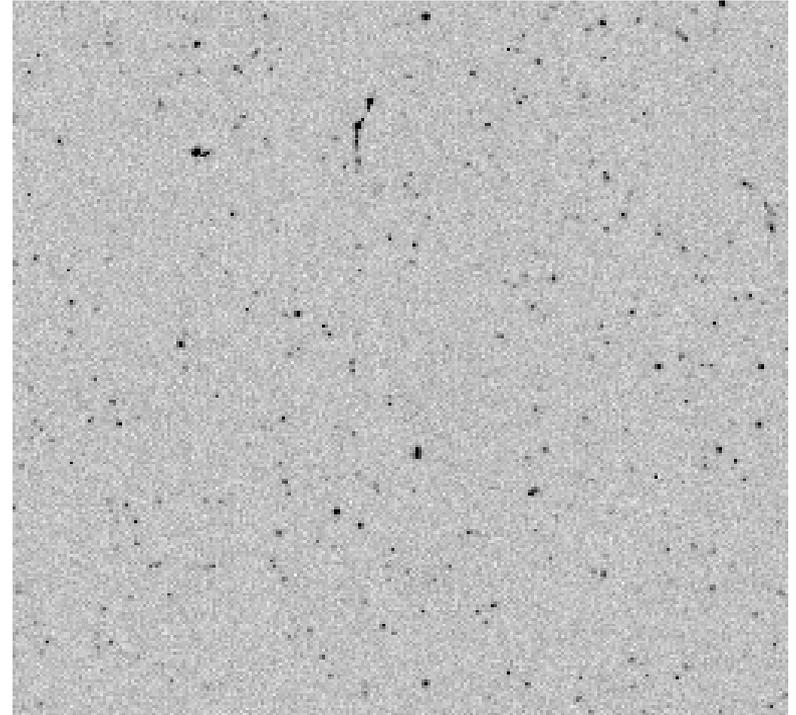
# Hybrid pixel detectors

- Very fast
- Radiation hard
- Medipix III has event processing
  - Winner takes all.
- Several companies are in process of introducing them for EM.
- Best suited for low energy (  $< 100$  keV)
- Medipix4 in the future

# Solution 2: Backthinned MAPS detectors

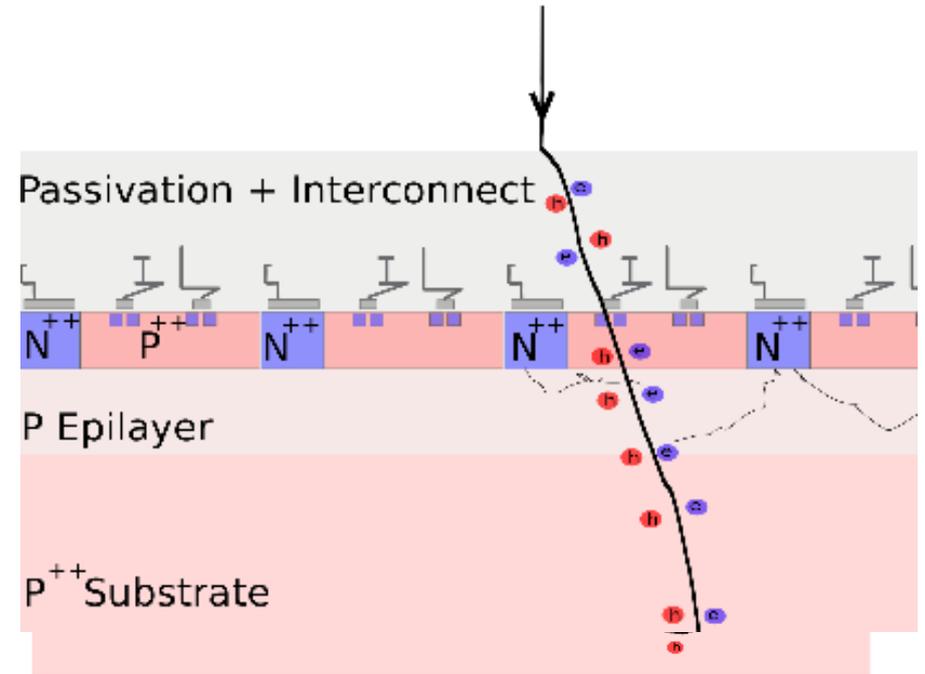
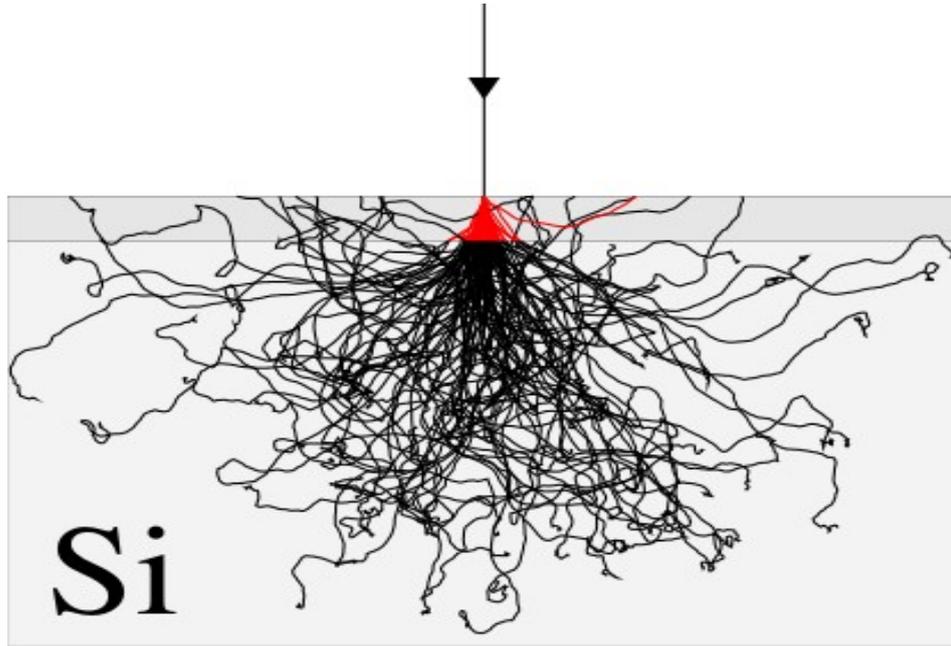


“Standard” CMOS technology

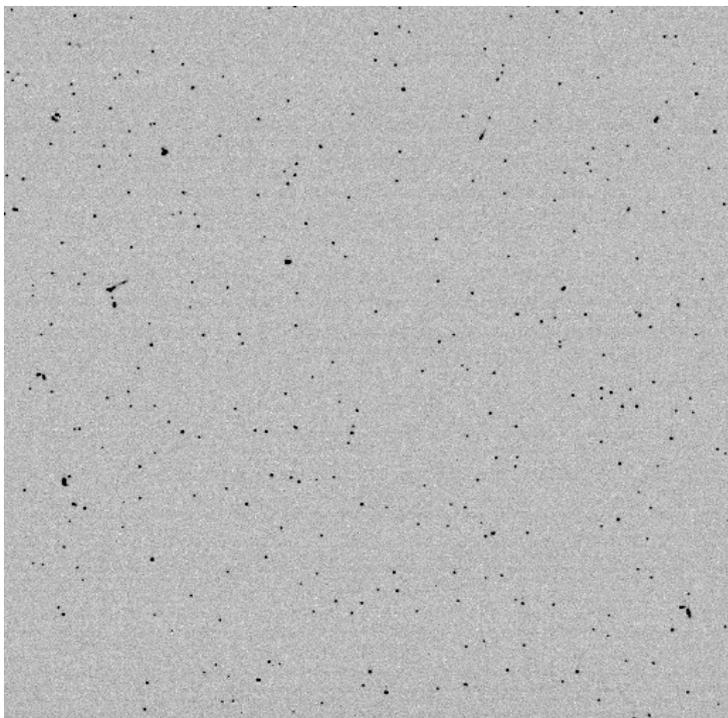


Large format with high sensitivity for electrons

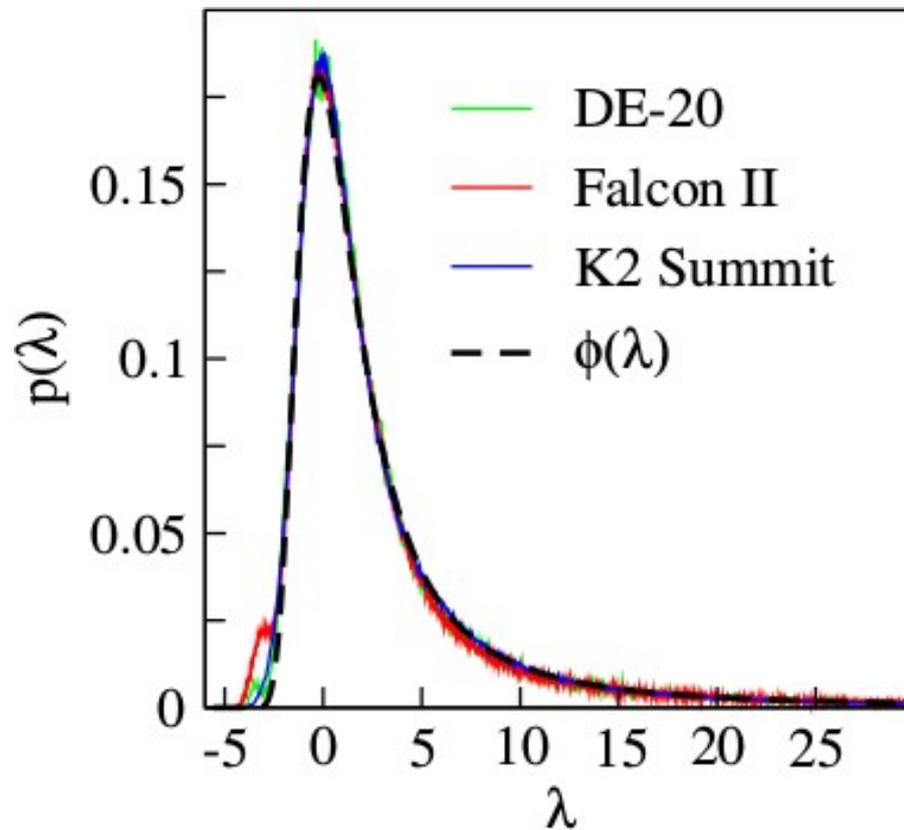
# Solution 2: Backthinning reduces track problem. (strong backscattered events)



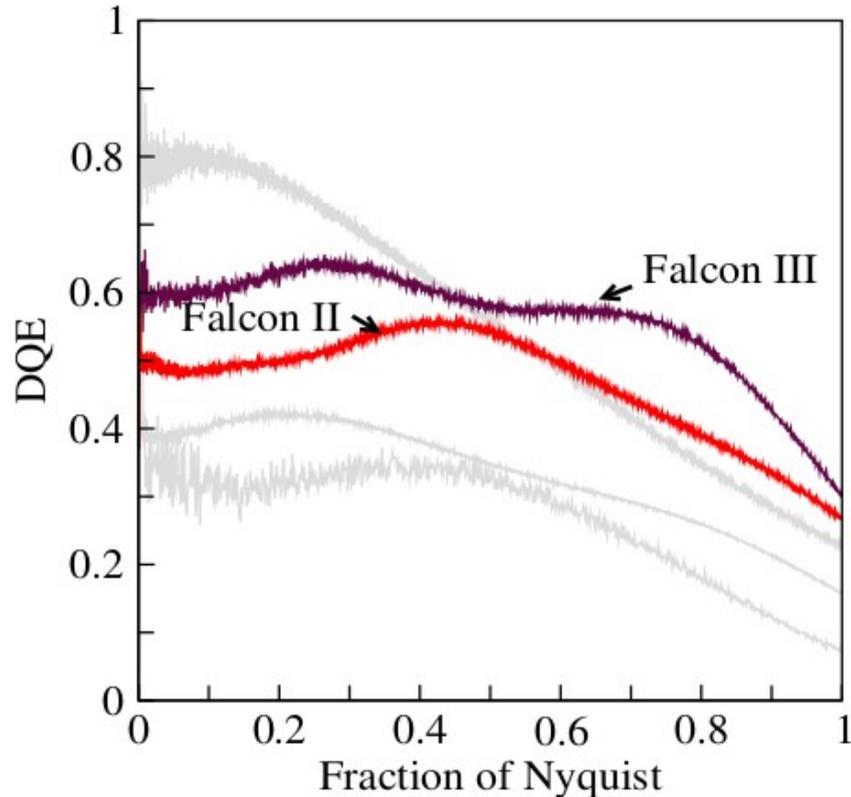
# Still have Landau noise problem



*Ultramicroscopy* **147** (2014) 156-163



# Landau noise limits integrating DQE



- DQE  $\sim 0.6$  is the limit of an integrating detector
- Better performance at Nyquist than current counting detectors
- Faster data acquisition

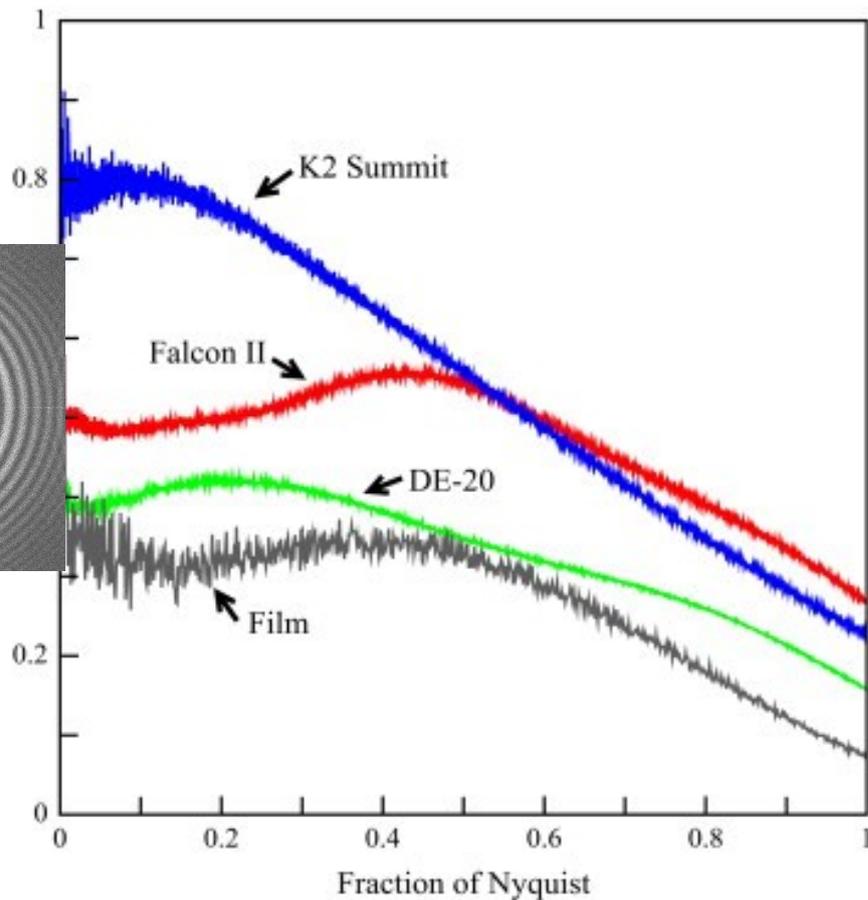
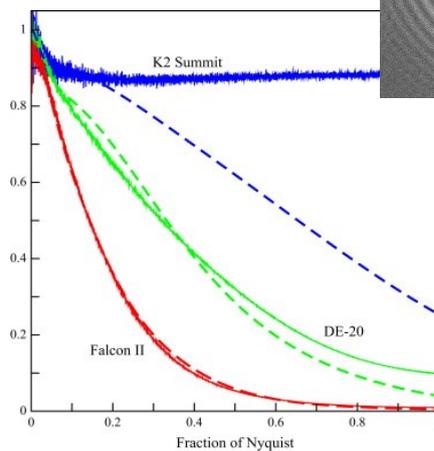
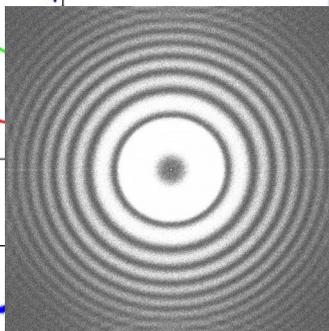
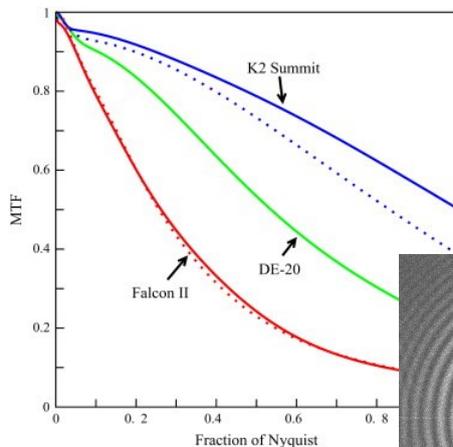
# Future is counting

- Anything to improve images helps
- Gatan K2 currently counts
- DE and FEI are introducing products
- Two problems:
  - Even K2 is not fast enough
  - Performance at Nyquist frequency is poor

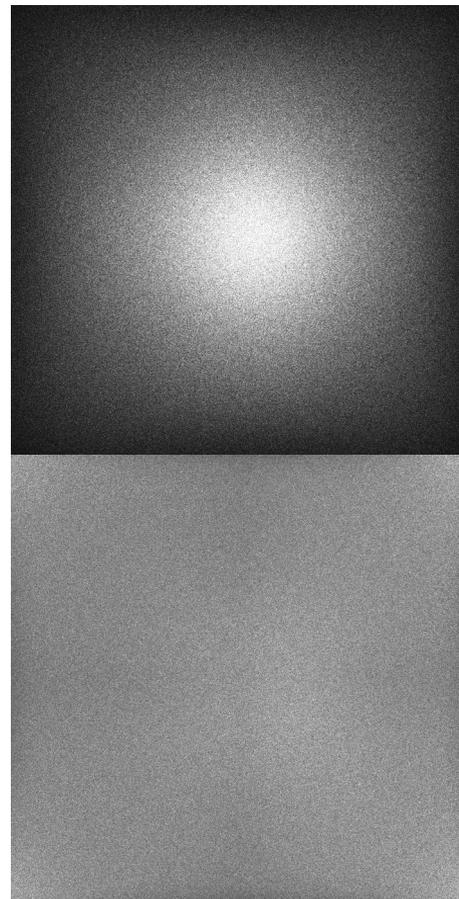
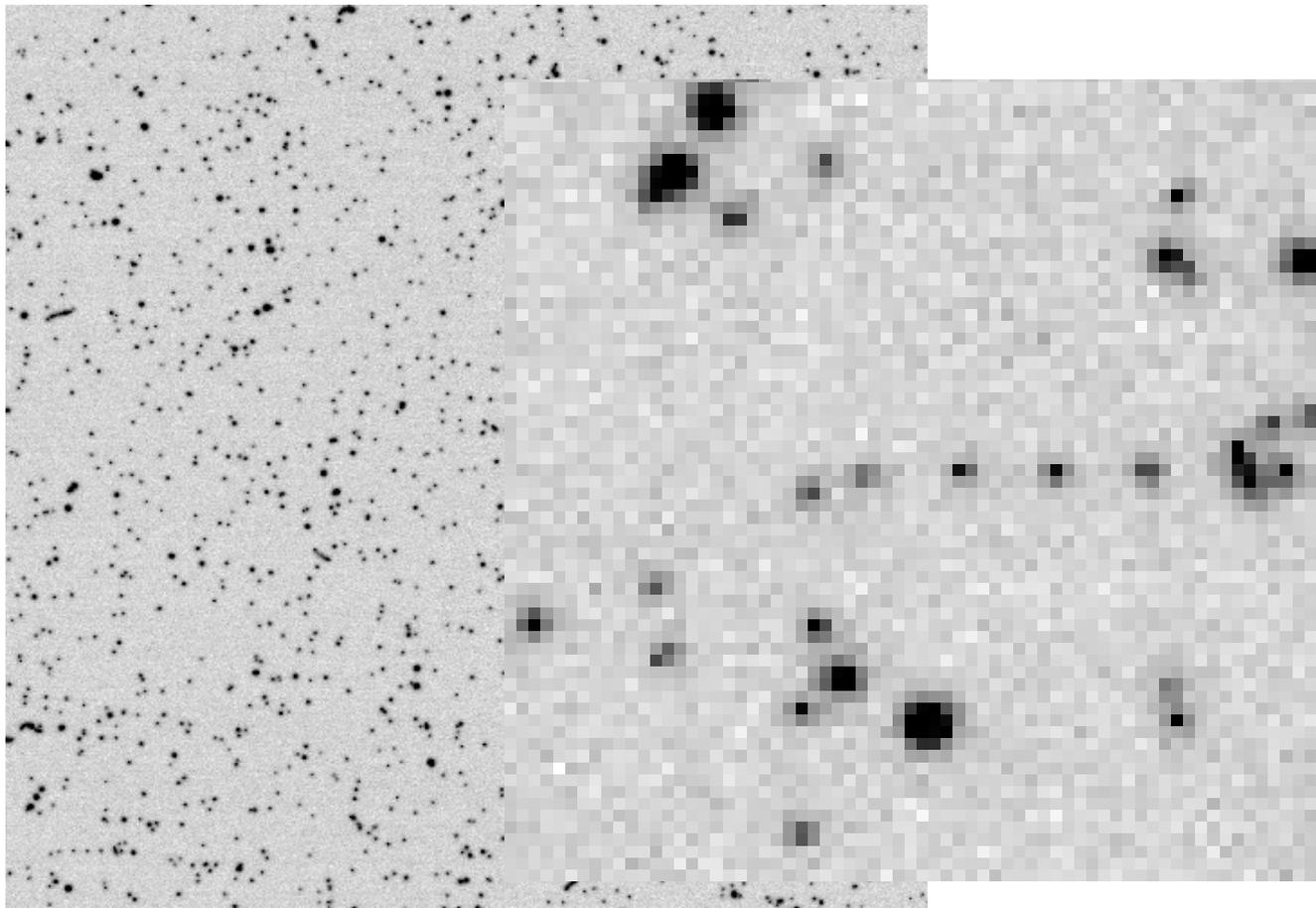
# Speed is important

- Can not tell if an event is 1 or 2 electrons
- Can not tell with neighbouring pixel events if this results from 1 or 2 electrons
- Need 1 electron per 100 pixels in a frame
- Long exposures need drift correction
- Long exposures slow down data acquisition

# DQE $\sim$ MTF<sup>2</sup>/NPS

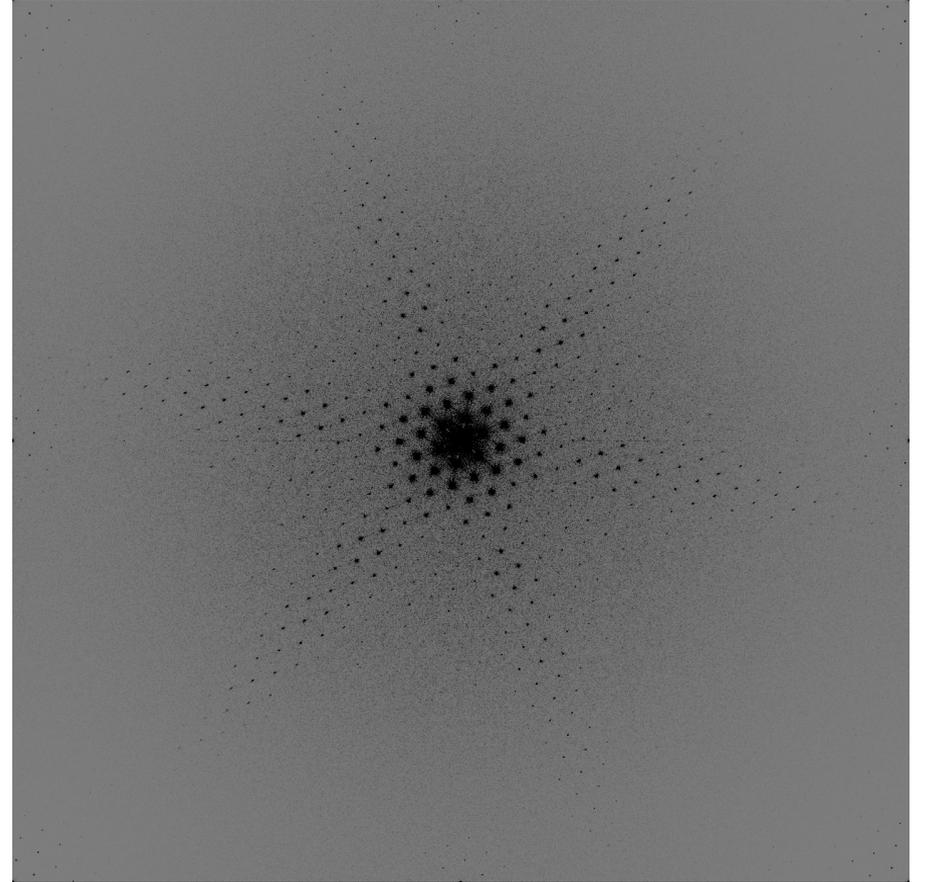
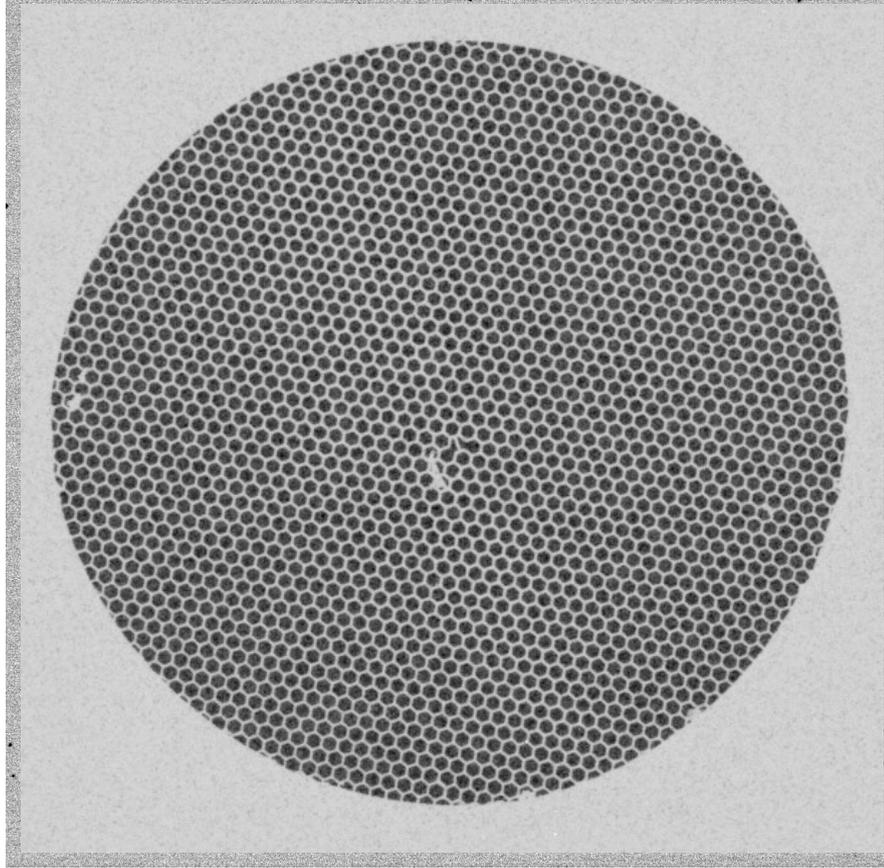


# Improving Nyquist frequency performance



Inferred incident electron position within pixel

# Showing improved performance



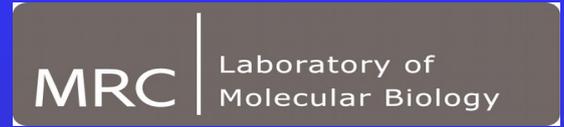
# Direct detectors

- Better images
- More images
- Movie processing
- Optimal exposures
  - CTF determination
  - Particle picking
  - Radiation damage weighting

# Future is Counting (but)

- Currently Gatan K2 summit is the only detector fast enough but still too slow.
- Nyquist frequency performance is too low. Getting the best DQE requires high magnification with small field of view
- Better performance is possible
  - when and how much will it cost

# Thanks



- Richard Henderson
- Wasi Faruqi
- Vinoth Kumar
- Jake Grimmett
- Toby Darling
- ...

Help from

