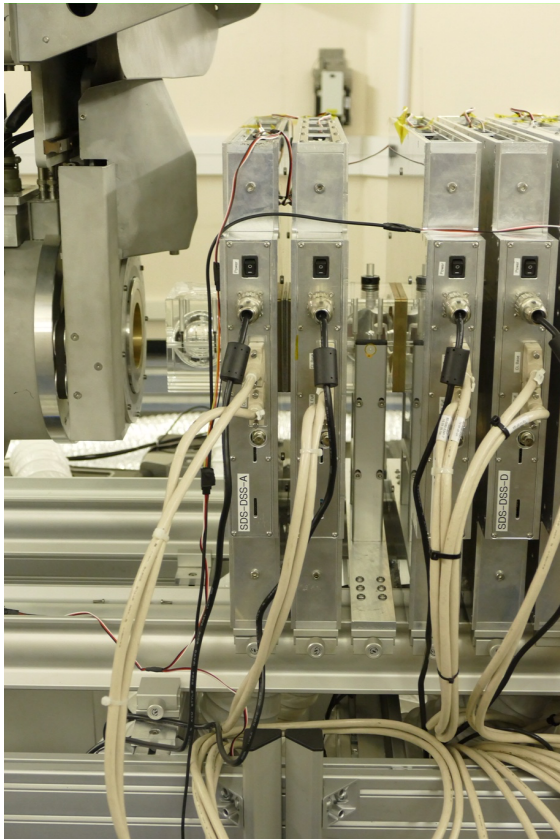


# Monte Carlo simulations for imaging in proton therapy

Michela Esposito, Tony Price, Jon T. Taylor, Chris Waltham, Sam Manger,  
Ben Phoenix, Gavin Poludniowski, Stuart Green, Philip M. Evans,  
Philip P. Allport, Spyros Manolopoulos, Jaime Nieto-Camero and Nigel M. Allinson

# Proton Radiotherapy Verification and Dosimetry Applications

– integrated platform for proton therapy imaging and dosimetry



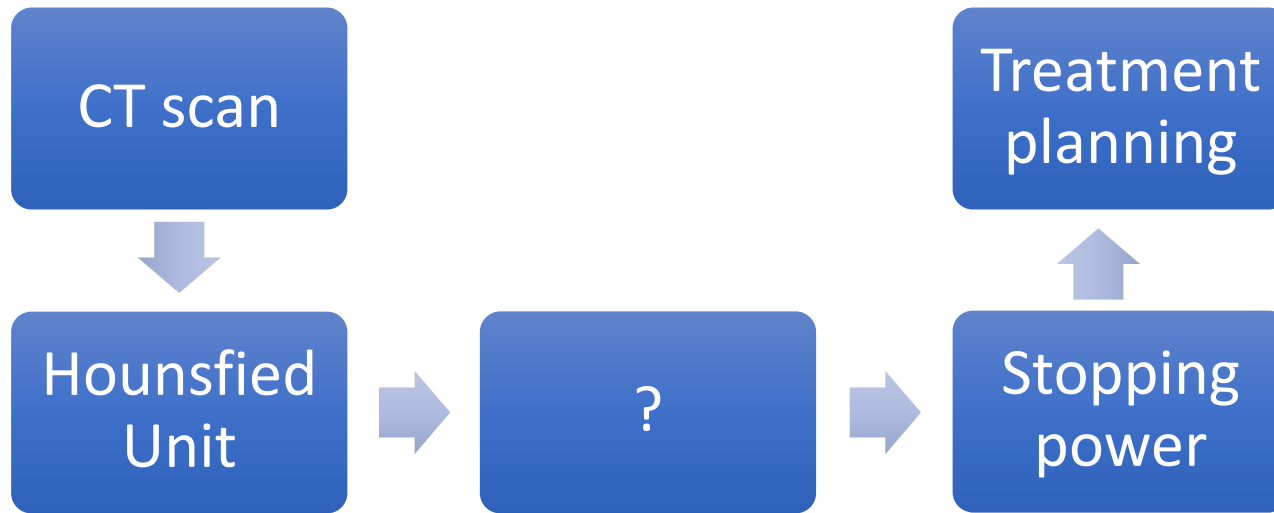
- University of Lincoln
- University of Birmingham
- University of Liverpool
- University of Surrey
- University of Cape Town
- University of Warwick
- Karolinska University Hospital, Sweden
- University Hospital Birmingham NHS Foundation Trust
- University Hospital Coventry and Warwickshire NHS Trust
- National Research Foundation (NRF) - iThemba LABS, SA
- United Lincolnshire Hospitals NHS Trust
- The Christie NHS Foundation Trust
  
- ISDI: Image Sensor Design and Innovation Ltd
- aSpect Systems GmbH
- Elekta AB (Publ)
- Advanced Oncotherapy Plc

Funded by

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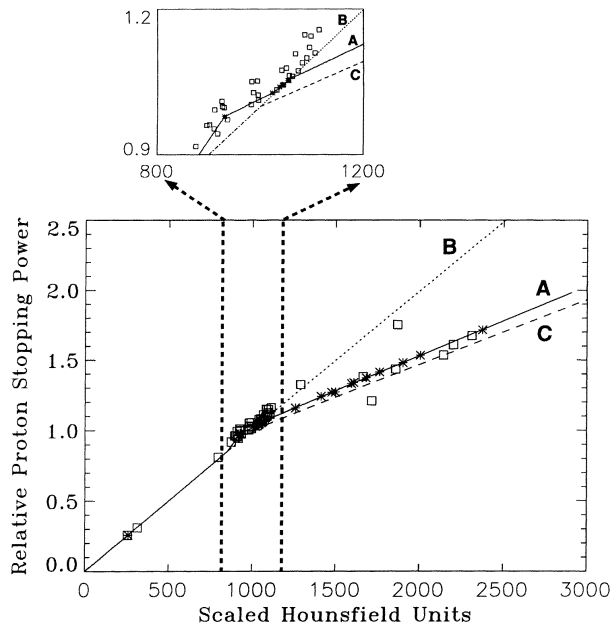
Grant Number 098285

# Why do we need proton CT?



# Why do we need proton CT?

## Stoichiometric calibration



U. Schneider, Phys. Med. Biol. **41**  
(1996) 111–124

IOP PUBLISHING

Phys. Med. Biol. **57** (2012) R99–R117

PHYSICS IN MEDICINE AND BIOLOGY

doi:10.1088/0031-9155/57/11/R99

## TOPICAL REVIEW

# Range uncertainties in proton therapy and the role of Monte Carlo simulations

Harald Paganetti

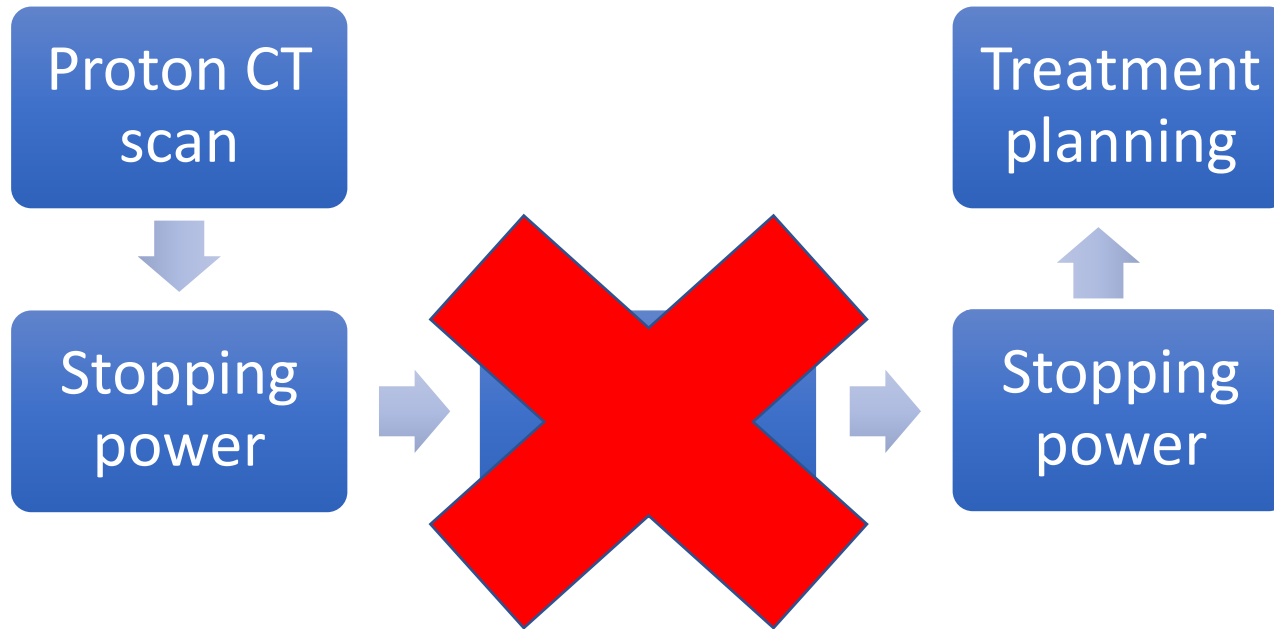
Source of range uncertainty in the patient	Range uncertainty without Monte Carlo	Range uncertainty with Monte Carlo
<b>Independent of dose calculation</b>		
Measurement uncertainty in water for commissioning	$\pm 0.3$ mm	$\pm 0.3$ mm
Compensator design	$\pm 0.2$ mm	$\pm 0.2$ mm
Beam reproducibility	$\pm 0.2$ mm	$\pm 0.2$ mm
Patient setup	$\pm 0.7$ mm	$\pm 0.7$ mm
<b>Dose calculation</b>		
Biology (always positive) ^	$\pm \sim 0.8\%$	$\pm \sim 0.8\%$
CT imaging and calibration	$\pm 0.5\%$ <sup>a</sup>	$\pm 0.5\%$ <sup>a</sup>
CT conversion to tissue (excluding I-values)	$\pm 0.5\%$ <sup>b</sup>	$\pm 0.2\%$ <sup>g</sup>
CT grid size	$\pm 0.3\%$ <sup>c</sup>	$\pm 0.3\%$ <sup>c</sup>
Mean excitation energy (I-values) in tissues	$\pm 1.5\%$ <sup>d</sup>	$\pm 1.5\%$ <sup>d</sup>
Range degradation; complex inhomogeneities	$-0.7\%$ <sup>e</sup>	$\pm 0.1\%$
Range degradation; local lateral inhomogeneities *	$\pm 2.5\%$ <sup>f</sup>	$\pm 0.1\%$
Total (excluding *, ^)	2.7% + 1.2 mm	2.4% + 1.2 mm
Total (excluding ^)	4.6% + 1.2 mm	2.4% + 1.2 mm

4

Uncertainty in proton stopping power leads to uncertainty in where protons stop



# Why do we need proton CT?

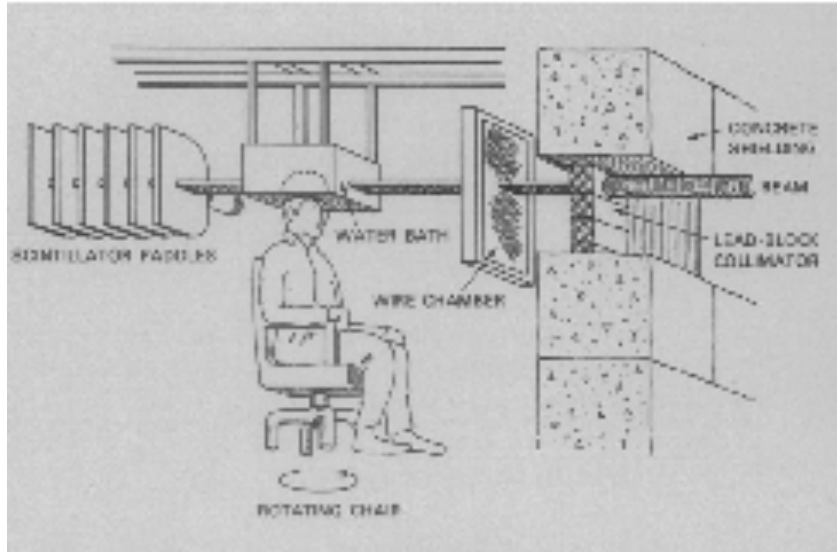


Aim to reduce stopping power uncertainty to 1%

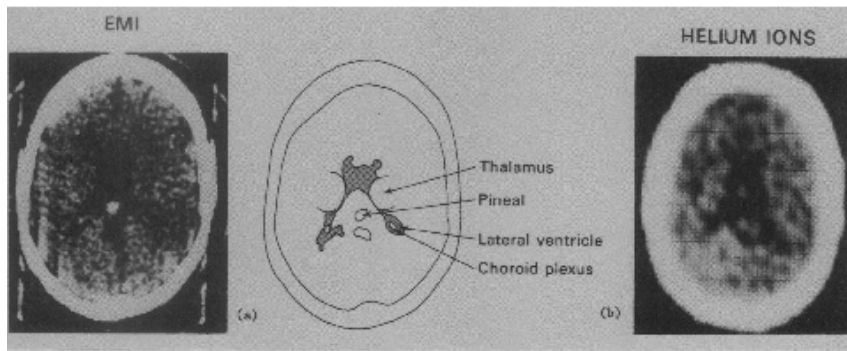
# HADRON COMPUTED TOMOGRAPHY

A 40 years old idea

The first alpha scanner ever trialled on humans



The next application of the solution [for Computed Tomography]... concerns the recent use of the peak in the Bragg curve for the ionisation caused by protons, to produce small regions of high ionisation in tissue. The radiotherapist is confronted with the problem of determining the energy of the incident protons necessary to produce the high ionisation at just the right place, and this requires knowing the variable specific ionisation of the tissue through which the protons must pass.



A. M. Cormack. Representation of a Function by Its Line Integrals, with Some Radiological Applications. *Journal of Applied Physics*, 34(9), 1963.

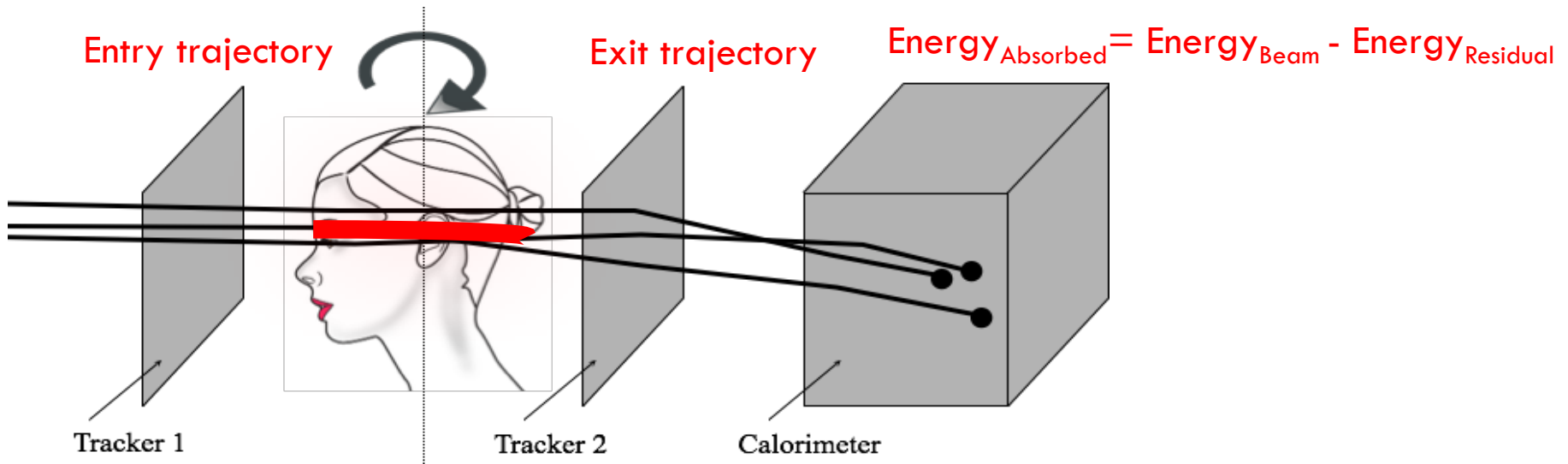
K.M. Crowe et al.. Axial Scanning with 900 MeV Alpha Particles. *Nuclear Science, IEEE Transactions on*, 22(3):1752–1754, June 1975.

# Basics of proton CT



Entry trajectory  
Exit trajectory  
Energy<sub>Absorbed</sub>

} Repeat millions of times!



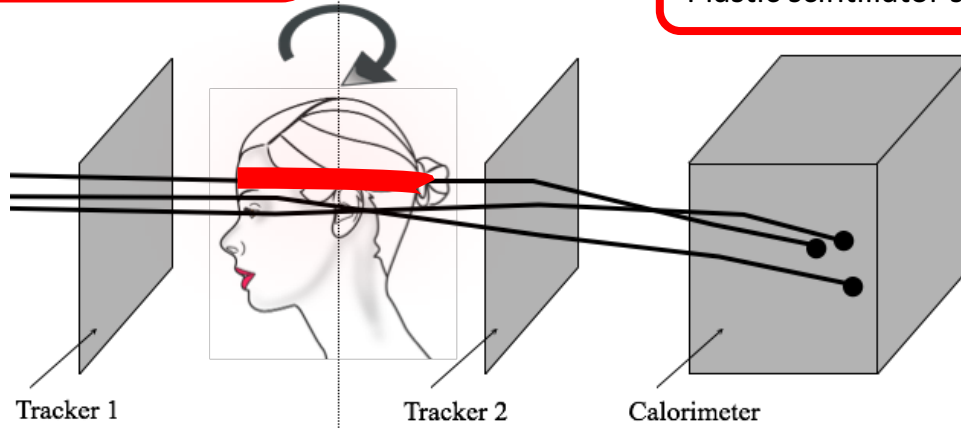
G. Poludniowski et al., Br J Radiol 2015; 88: 20150134.

Category	Parameter	Value
Proton beam	Energy	$\geq 200$ MeV (head)
		$\geq 250$ MeV (body)
	Flux <sup>a</sup>	$\geq 3000$ protons $cm^{-2} s^{-2}$
Imaging dose	Maximum absorbed dose <sup>b</sup>	$< 20$ mGy
Image quality	Spatial resolution, $\sigma$	$\approx 1$ mm
	Relative stopping-power accuracy	$< 1\%$
Time	Data acquisition time	$< 10$ min
	Reconstruction time	$< 10$ min

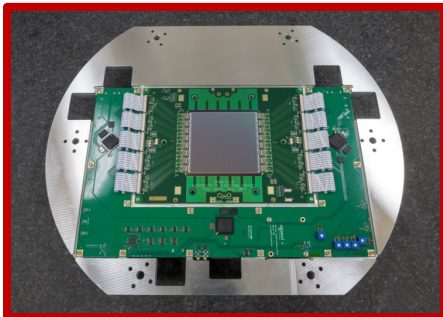
# Basics of proton CT

Multi-wire Proportional Chambers  
Double-Sided Strip Detectors  
Scintillating Fibres Hodoscopes  
Gas Electron Multiplier detectors

Scintillator-based calorimeter  
Plastic scintillator stack

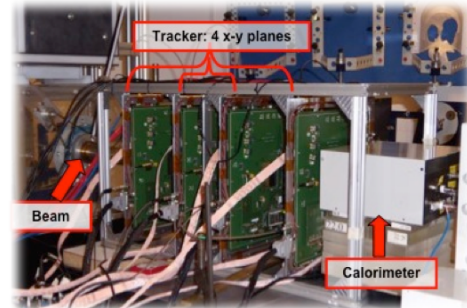


## PRaVDA

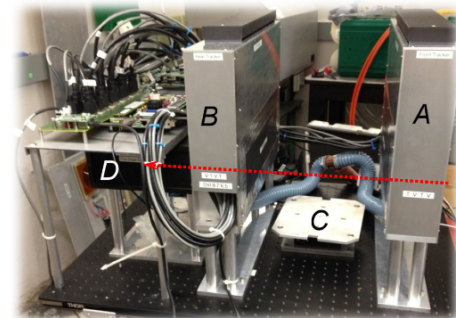


The first solid-state energy-range detector for proton CT.

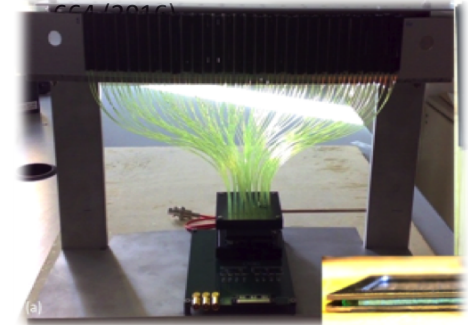
Unlike calorimeters, position sensitive detectors allow for multiple proton tracks to be detected in a single readout cycle potentially reducing CT scan times.



V. Sipala et al., POS (RD11) 013, 2011



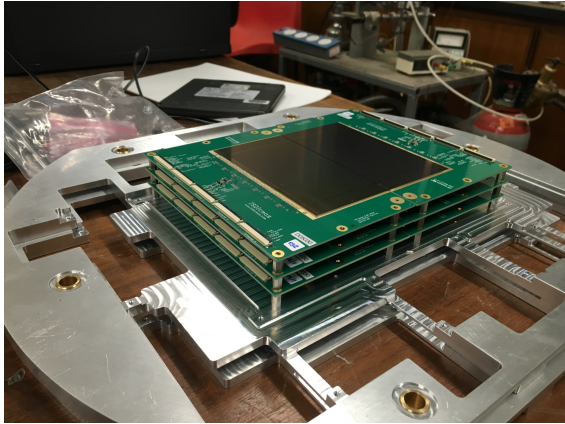
V.A. Bashkurov et al., Med. Phys. **43**, 2016



D. Lo Presti et al., J. Inst. **9**, C06012, 2014

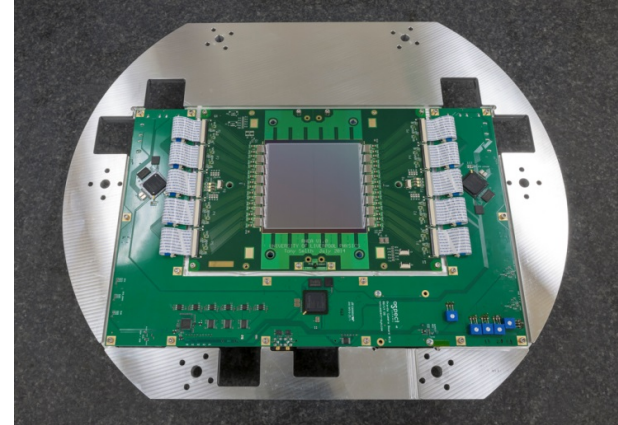
# Detector technology

## CMOS Active Pixel Sensors



- 2D-positional detectors
- Analog readout
- kHz readout (high occupancy per R/O cycle)
- Moderately radiation tolerant
- Mosaic tiling of edge-less sensors to cover larger areas
- High material budget

## Silicon Strip Sensors



- 1D-positional detectors
- Binary readout (in our implementation)
- MHz readout (low occupancy per R/O cycle)
- Radiation tolerant to LHC doses
- Dead areas when tiling to larger areas
- Low material budget



# The PRaVDA proton CT system

## 2 tracking units

- 4 sets of 3 layers of Silicon Strip Detectors (SSD)
- Crossed at 60°

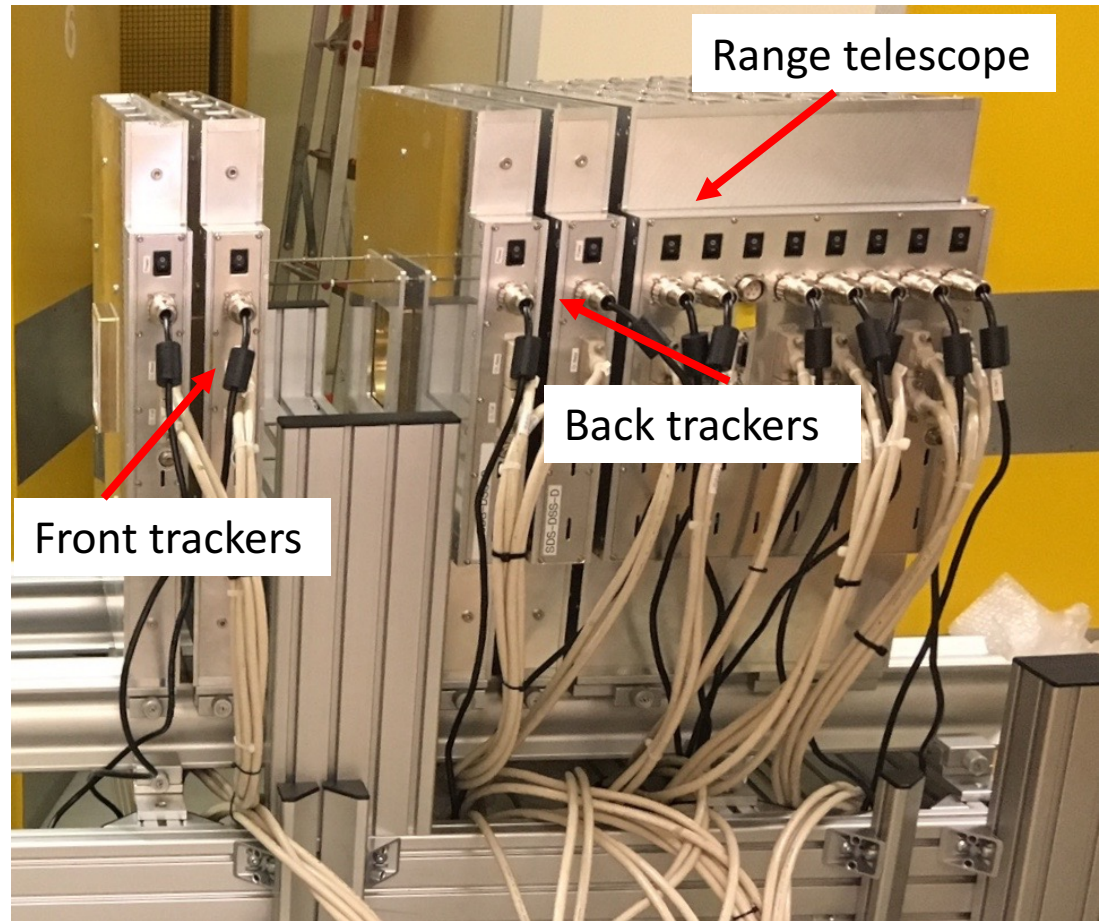
## Range Telescope

- 21 layers (SSD)
- 1D tracking

Readout frequency = 26 MHz

Max hit rate =  $2 \times 10^8$  hits/second  
(uniform field)

Total data throughput = 66 Gb/s



# Why do we need a Monte Carlo simulation?

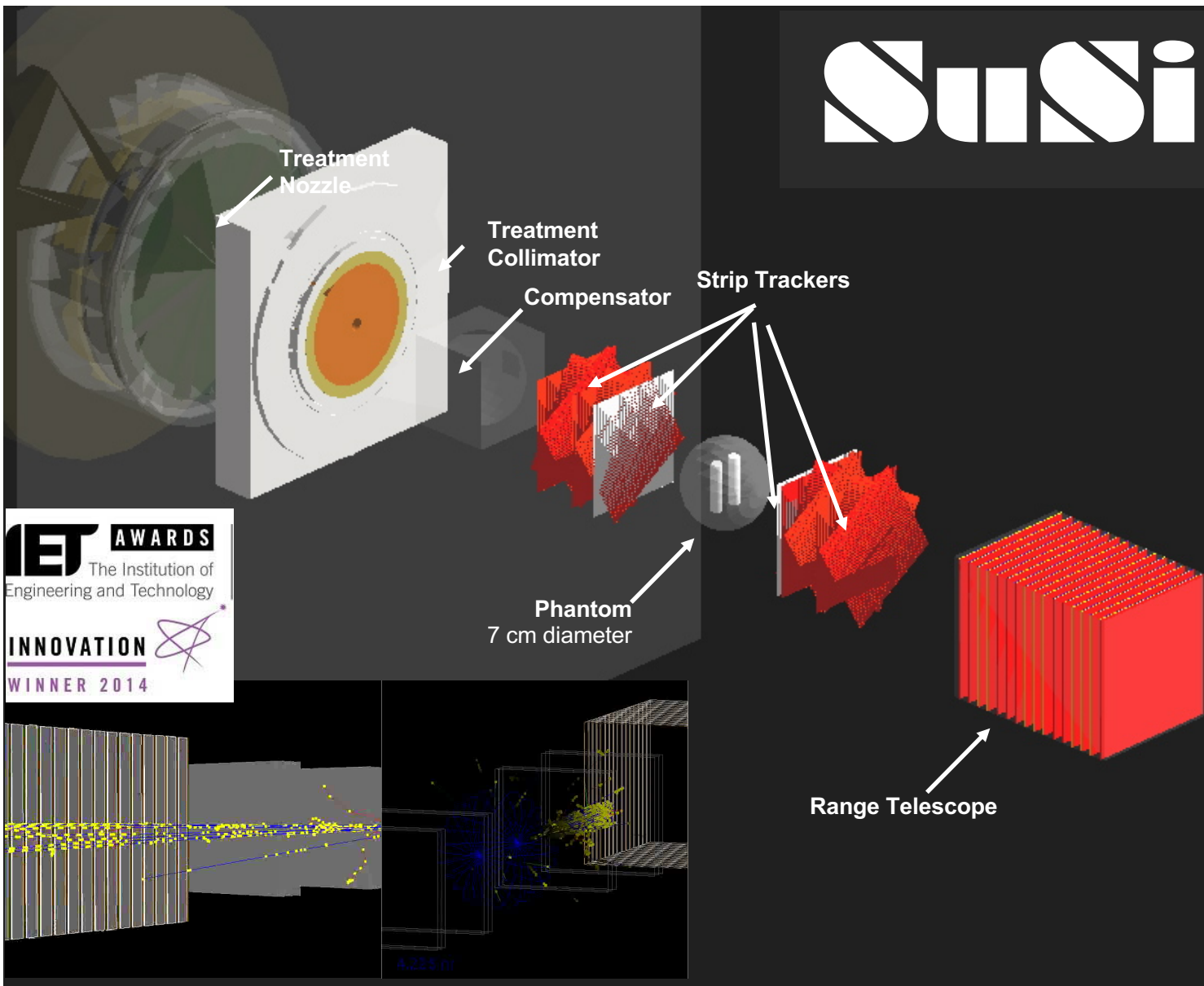
## Detector design

- 2 different technologies
- SSD derived from HEP
- CMOS sensors derived from medical imaging

## Radiation tolerance and shielding

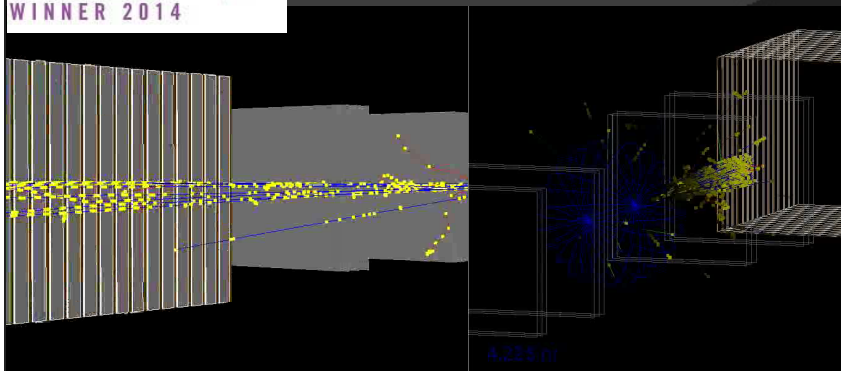
## Tracking algorithms (trackers and range telescope)

## CT reconstruction algorithms



Geant 4.10.1  
Standard opt\_3  
QGSP\_BIC

University of  
Birmingham BlueBEAR  
HPC cluster and GridPP

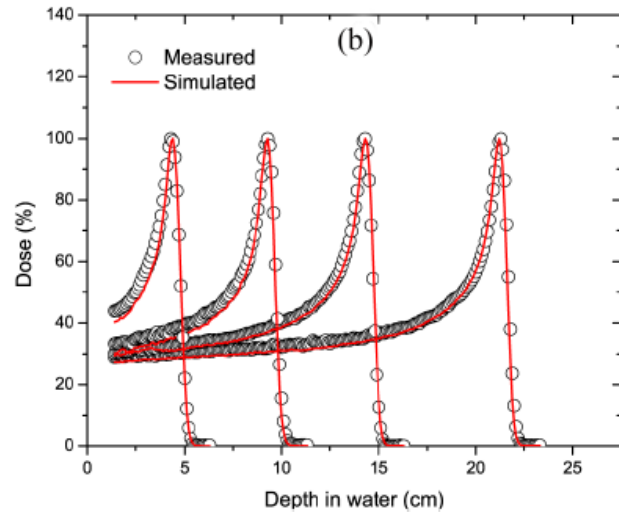


More details on beam line models in **Tony Price's** talk yesterday: "Code sharing of MC beam models for advanced radiotherapy" (ID: 201) and poster "A validated model of the University of Birmingham Medical Beamline (ID:248)

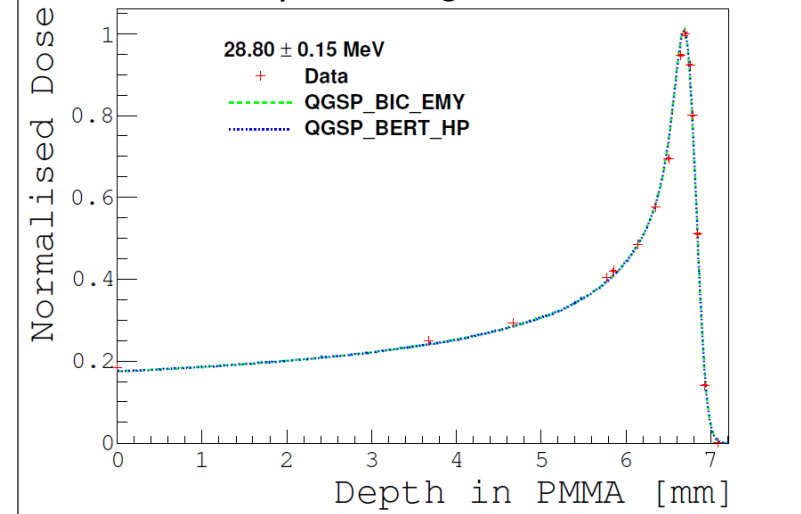


# SuSi – validation results

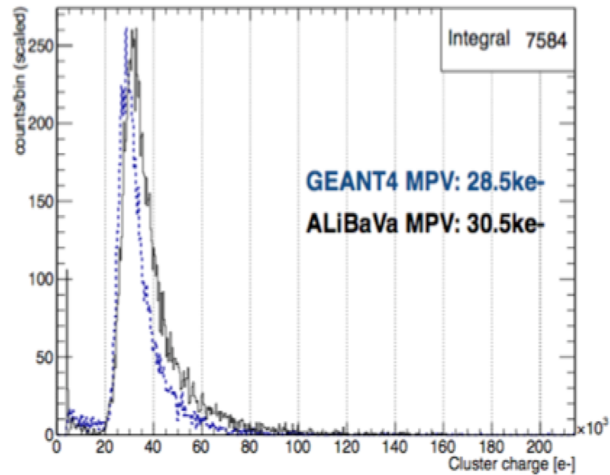
iThemba beamline



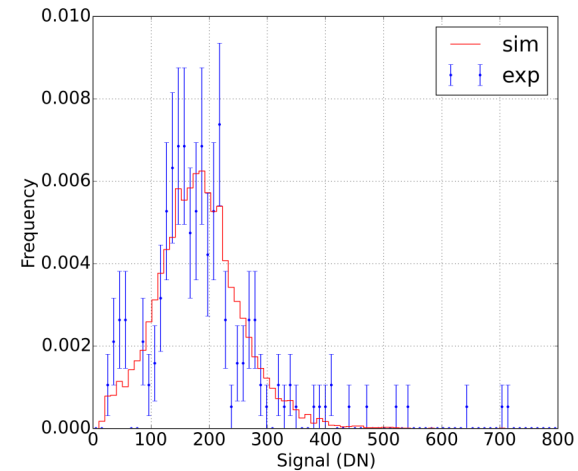
University of Birmingham beamline



SDS – 191 MeV p



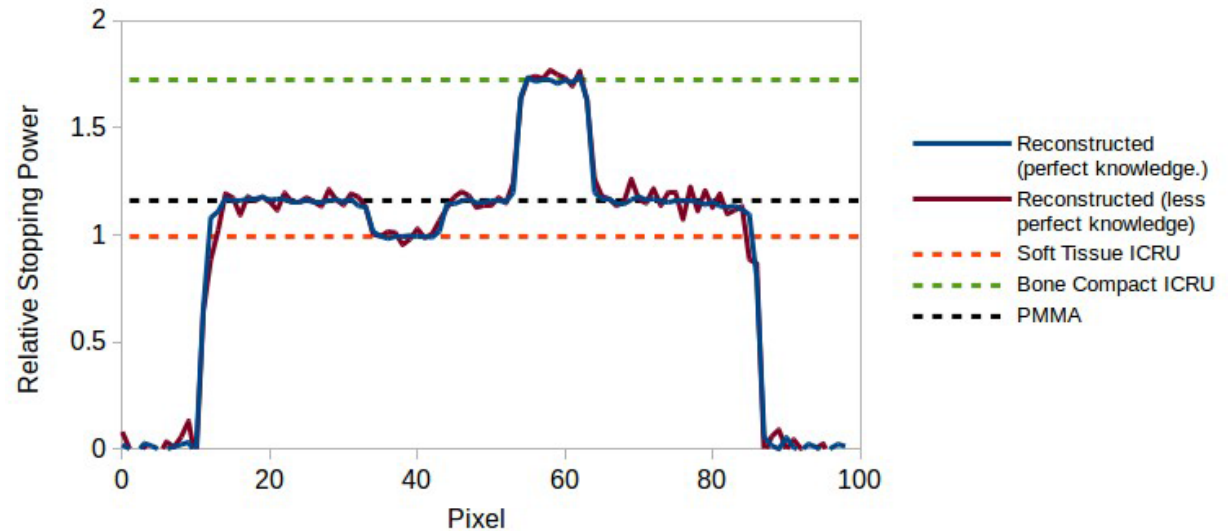
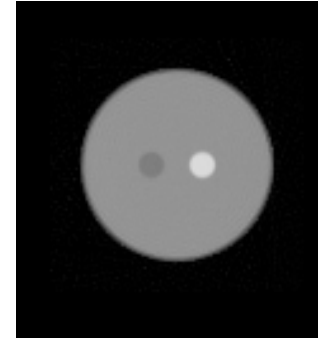
CMOS sensor – 29 MeV p



# Proton CT reconstruction algorithm

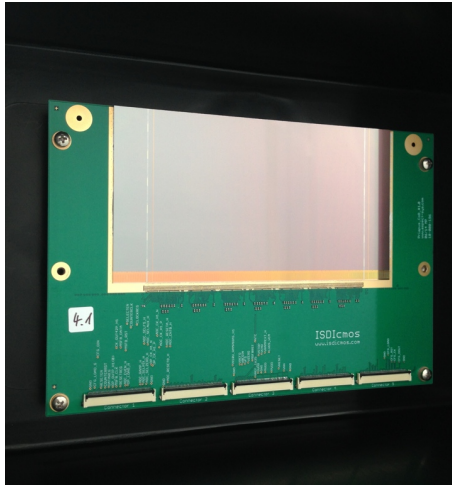
Novel algorithm for CT reconstruction:  
Back projection-then-filtering

Stopping power uncertainty <0.2%



Poludniowski, G., Allinson, N.M. and Evans, P.M., 2014. Proton computed tomography reconstruction using a backprojection-then-filtering approach. *Physics in medicine and biology*, 59(24), p.7905.

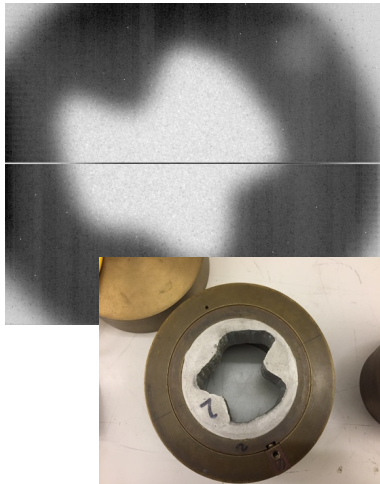
# The PRaVDA CMOS imager



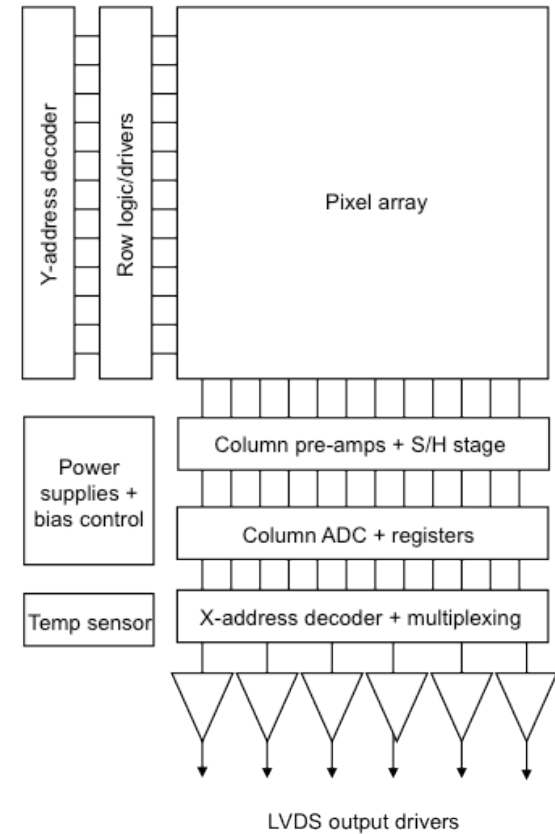
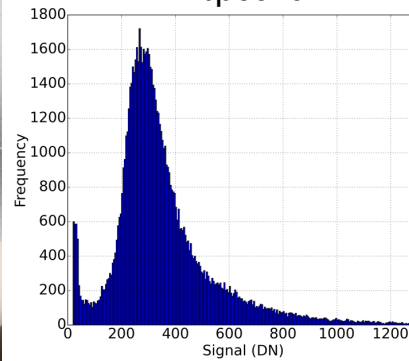
Design specifications:

- 0.35  $\mu\text{m}$  technology
- 5 cm  $\times$  10 cm imaging area
- 3-side buttable
- 194  $\mu\text{m}$  pixel
- 150 e<sup>-</sup> noise floor
- 1 kHz frame rate (11 bits)

Patient collimator



29 MeV protons signal spectrum



Further readings:

M. Esposito et al, J. Inst 2015; 10 (06), C06001

T. Price et al., J. Inst. 2015; 10 (05), P05013

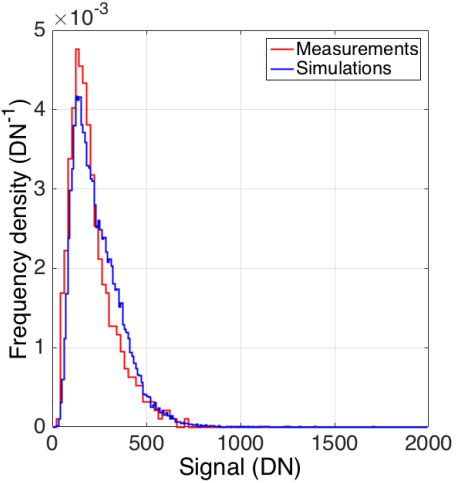
G. Poludniowski et al., Phys. Med. Biol. **59**  
(2014) 2569–2581

# Charge sharing model

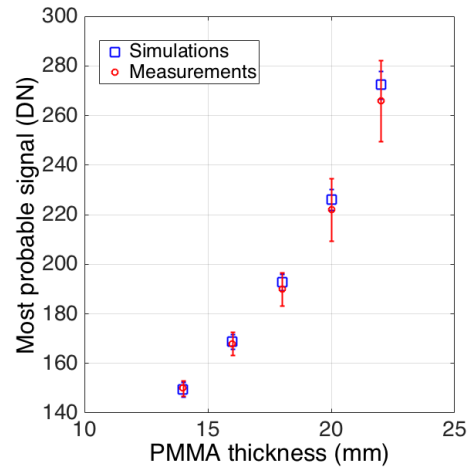


Geant4-based simulations of charge collection in CMOS Active Pixel Sensors, M. Esposito et al., Jinst 12 P03028, 2017

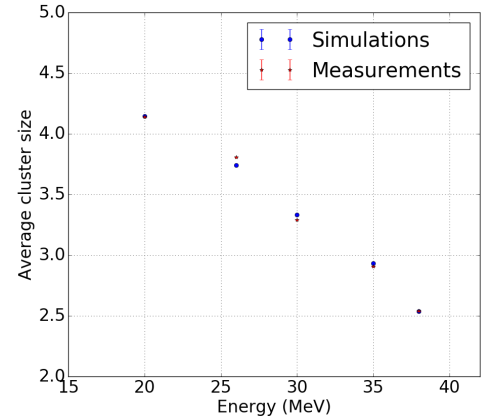
## Validation results



Signal spectrum(38 MeV p)

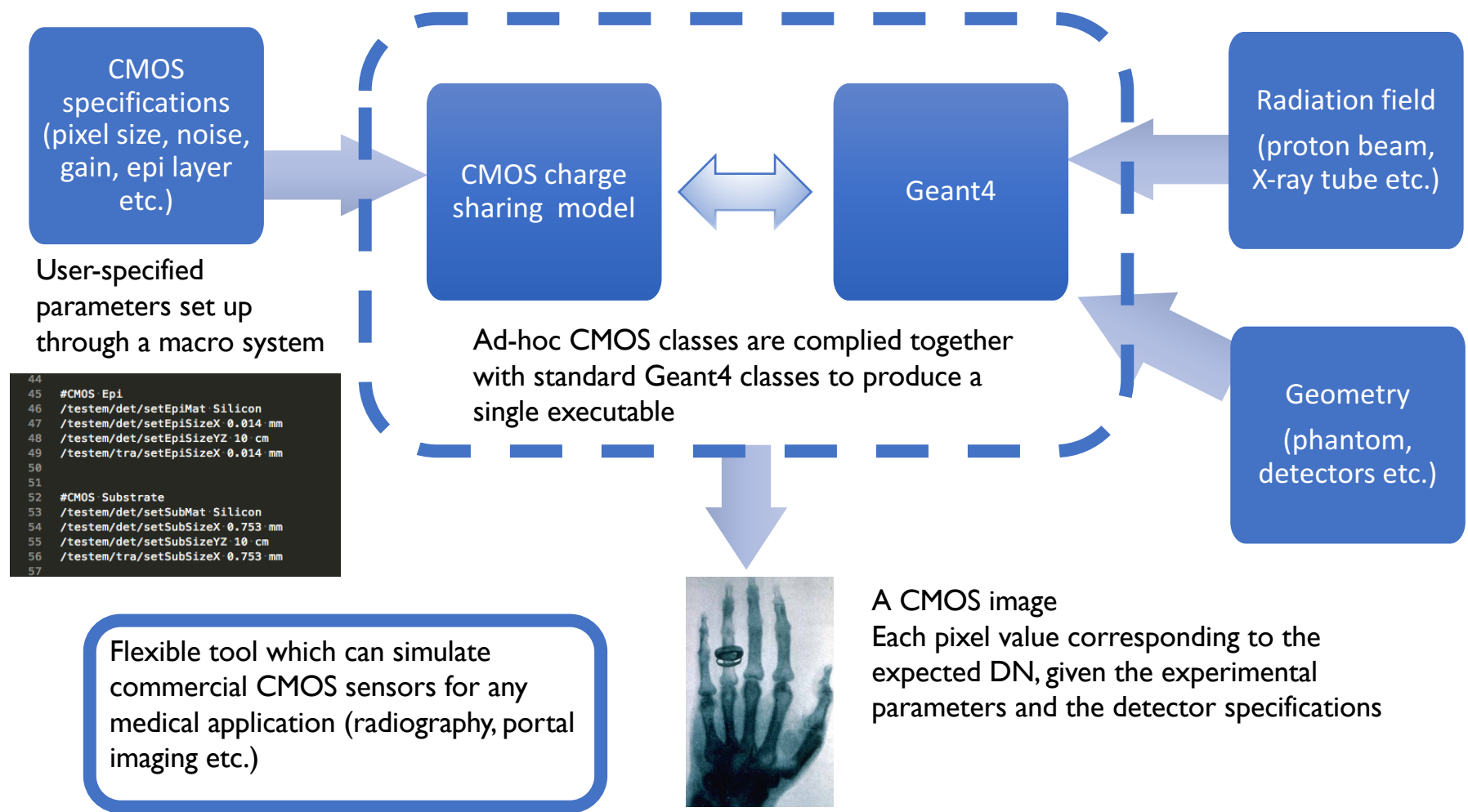


Most probable signal (20-38 MeV p)

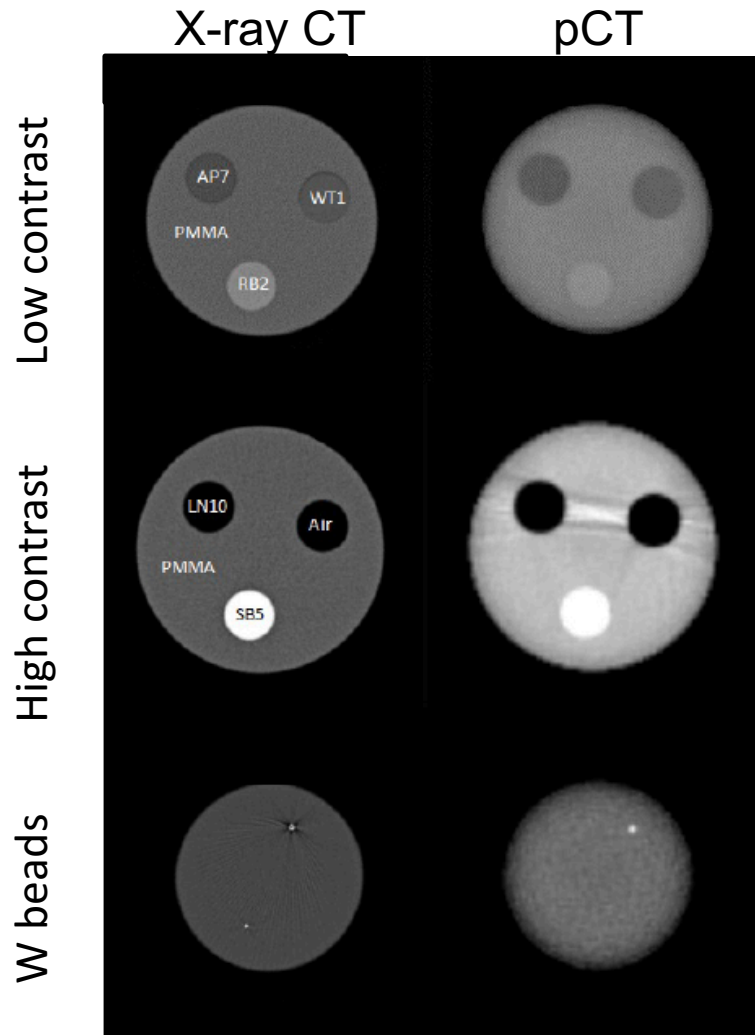
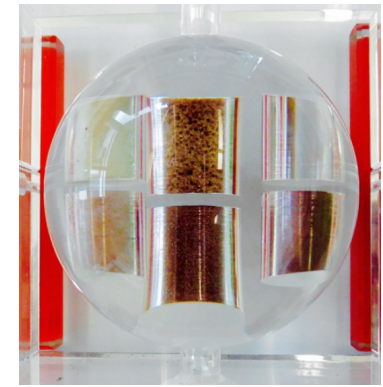


Average cluster size (20-38 MeV p)

# Integration into the Geant4 toolkit



# PRaVDA proton CT



Material	Density [g/cm <sup>3</sup> ]	Expected RSP	pCT RSP	Percent error
PMMA	~1.16	1.15	1.15	0.0
AP7	0.92	0.95	0.94	-0.7
WT1	1.00	1.00	0.98	-1.6
RB2	1.40	1.21	1.22	1.2
SB5	1.84	1.63	1.62	-0.4
LN10	0.25-0.35	0.25	0.29*	-
AIR	0.00	0.00	0.09*	-

\*The image slices containing the LN10 insert and air cavity manifest streak artefacts that compromise quantitative accuracy. For that reason, percentages error is not shown for these two materials.

# Conclusions

- PRaVDA has developed 2 solid-state technologies for proton CT
  - Design heavily relied on MC simulations
  - Simulation of charge sharing in CMOS Active Pixel Sensors
  - Model of a multi-step process from e/h pair generation to digitalisation
  - Flexible tool to be integrated into Geant4
  - Developed for proton CT but can be seamlessly extend to:
    - Different commercial CMOS sensors (just setting sensor specs)
    - Different radiation field/geometry
    - E.g. radiography, mammography, portal imaging, fluoroscopy etc.
  - Happy to share the code for different applications/experiments
- [mesposito@lincoln.ac.uk](mailto:mesposito@lincoln.ac.uk)
- On our first proton CT stopping power uncertainty equal or lower of 1.6% - preliminary analysis



# Acknowledgments

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Michael Koeberle

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