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15-18 October 2017 *Napoli, Italy*
Europe/Rome timezone

A Monte Carlo perspective on small beam radiation therapy

Jan Seuntjens
Medical Physics Unit
McGill University
Canada



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Kamen Paskalev (2002)
Hugo Bouchard (2004)
Laurent Tantot (2007)
Justin Sutherland (2009)
Eunah Chung (2011)
Pavlos Papaconstadopoulos (2013)
Lalageh Mirzakhania (2017)



at CMA, Napoli, Oct. 16, 2017



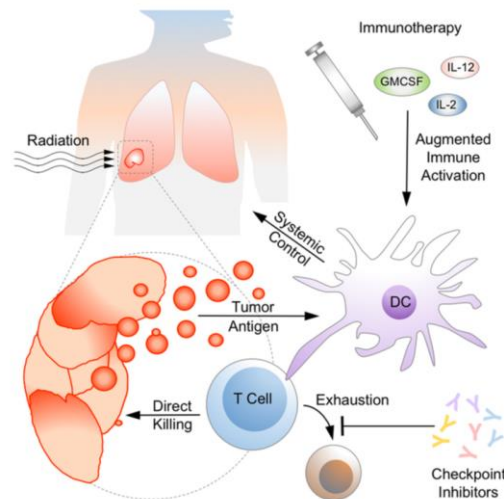
Outline

- Rationale and reminder of seminal milestone
- Small field characteristics
- Detectors and small fields
 - LCPE
 - Response decomposition
 - Detector density
 - Calibration of small fields (G-Knife, sub-LCPE fields)
- Beam model commissioning
- TPS algorithms & small fields
- Why do we care?

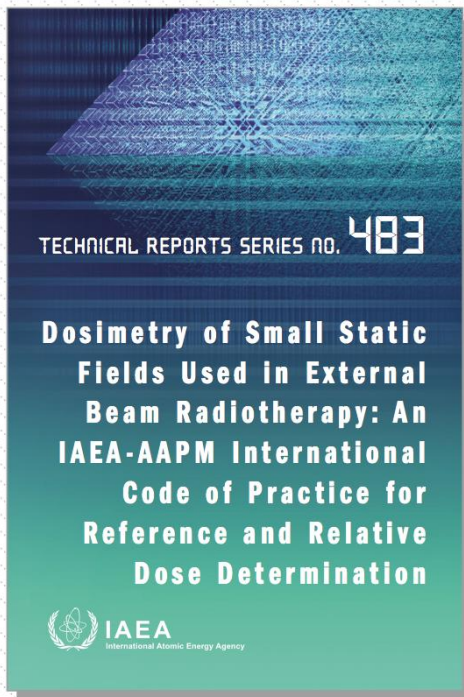
Small beam radiation therapy (SBRT)

- Biology of high dose / fraction : **BED > 100 Gy**
- Synergy of SBRT and immunotherapy
 - Melanoma
 - Renal tumours
 - Sarcomas
- Reporting of SBRT

Translational Lung Cancer Research, Vol 6, No 2 April 2017



Two important reports



Dosimetry of Small Static Fields Used in External Beam Radiotherapy

An IAEA-AAPM International Code of Practice for Reference and Relative Dose Determination

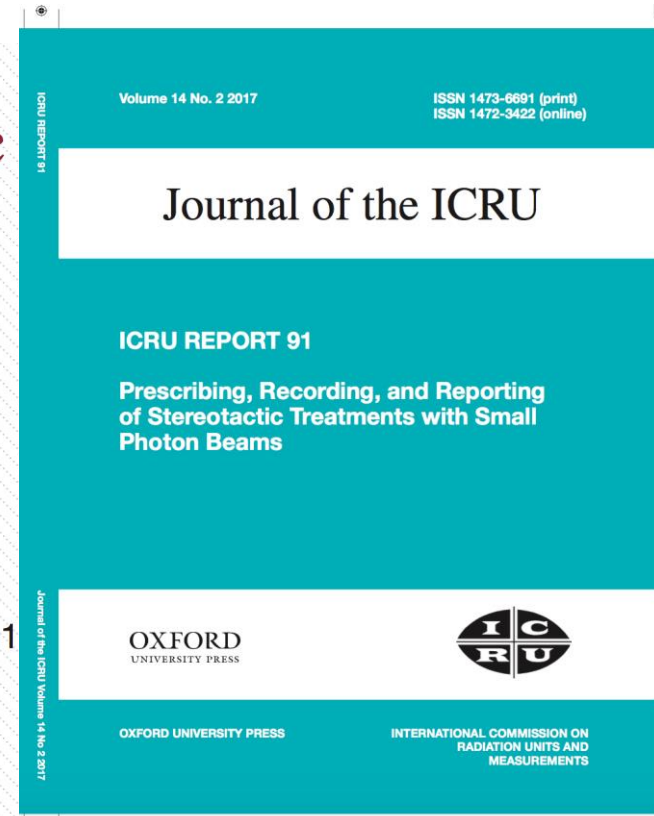
Technical Reports Series No. 483

STI/DOC/010/483 • ISBN 978-92-0-105916-1

Price: €52.00

Language: English

Forthcoming: 2016



IAEA-TRS 483: Which problems does it solve?

- Characteristics that lead to dosimetric issues of two kinds:
 - Reference dose calibration
 - Reference fields are not 10 x 10 cm², SSD/SAD is not 100 cm, etc; they are called “machine-specific reference fields” (*msr*)
 - Flattening filter-free beams, beam quality specification
 - Output factors
 - Small fields
 - Detector correction factors
- Problem that was put on the backburner: calibration of composite fields

2 A new formalism for reference dosimetry of small and nonstandard fields

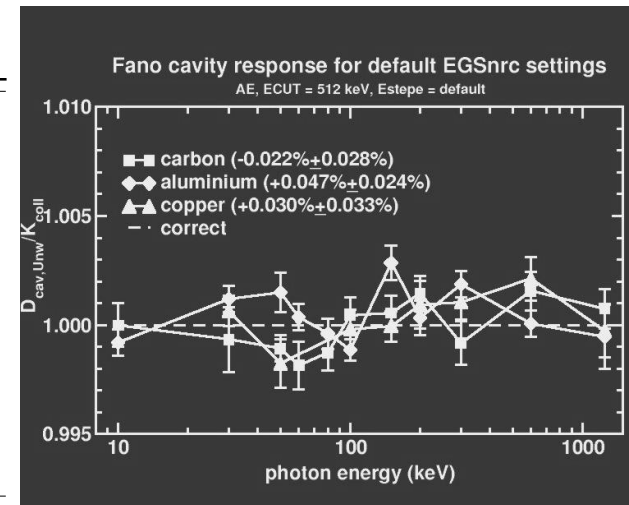
3	R. Alfonso
4	<i>International Atomic Energy Agency, Vienna, Austria and Instituto Nacional de Oncologia y Radiobiologia,</i>
5	<i>La Habana, Cuba</i>
6	P. Andreo
7	<i>International Atomic Energy Agency, Vienna, Austria and University of Stockholm-Karolinska Institutet,</i>
8	<i>Stockholm, Sweden</i>
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17	T. R. Mackie
18	<i>Department of Medical Physics, University of Wisconsin, Madison, Wisconsin</i>
19	H. Palmans ^{a)}
20	<i>National Physical Laboratory, Teddington, United Kingdom and Slovensky Metrologicky Ustav, Bratislava,</i>
21	<i>Slovakia</i>
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23	<i>Royal Marsden NHS Foundation Trust, Sutton, United Kingdom</i>
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28	S. Vatnitsky
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31	published xx xx xxxx)

The “Alfonso” paper

Reminder - Seminal enabling work

Ion chamber simulation at ^{60}Co : resolution of EGS4/PRESTA artifacts

Artifact	Aluminium	Carbon	Aluminium	Carbon
ESTEPE step control→	20%	20%	1%	1%
electron step	-9.0%	-5.0%	-1.4%	-0.7%
BCA	+3.4%	+2.6%	+1.5%	+0.9%
energy loss	+0.3%	+0.5%	+0.0%	+0.0%
discrete interactions	+0.7%	+0.7%	+0.7%	+0.7%
Totals	-4.6%	-1.2%	+0.8%	+0.9%



Application to kV and MV beams (Seuntjens et al 2001)

EGSnrc: Kawrakow, 2000

Penelope: Sempau & Andreo 2006

GEANT4: Poon et al 2003; Elles & Maire 2006

Small fields in stereotactic nonmalignant treatments

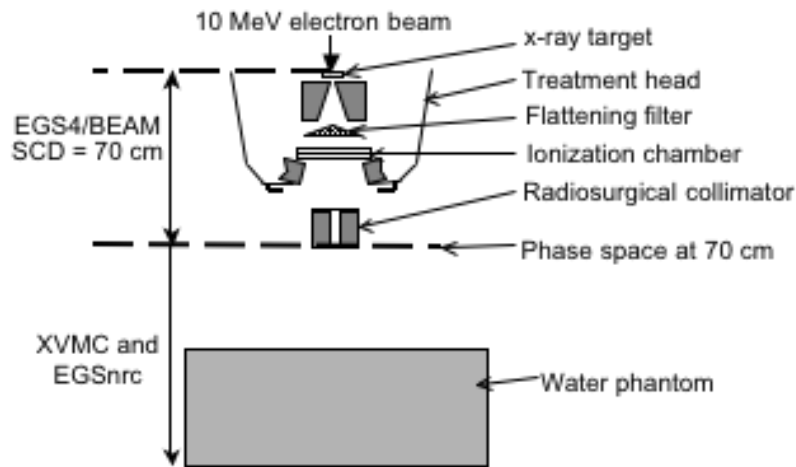


Figure 1: Beam entry trace (baseball seam) on the patient's skull for the dynamic radiosurgery technique.

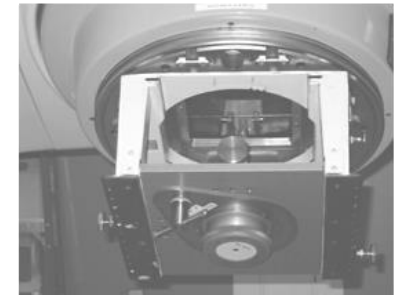
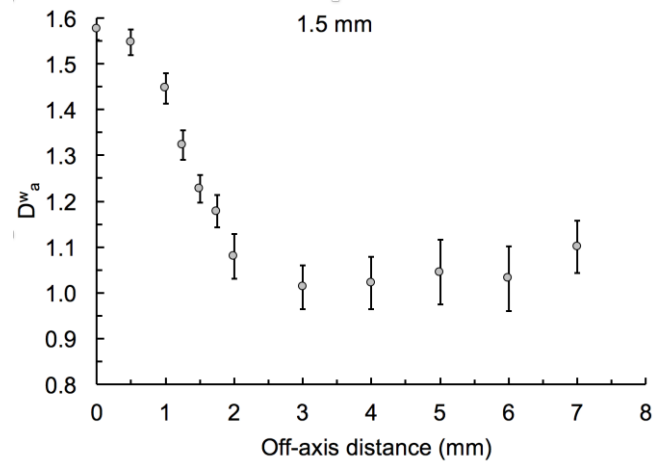
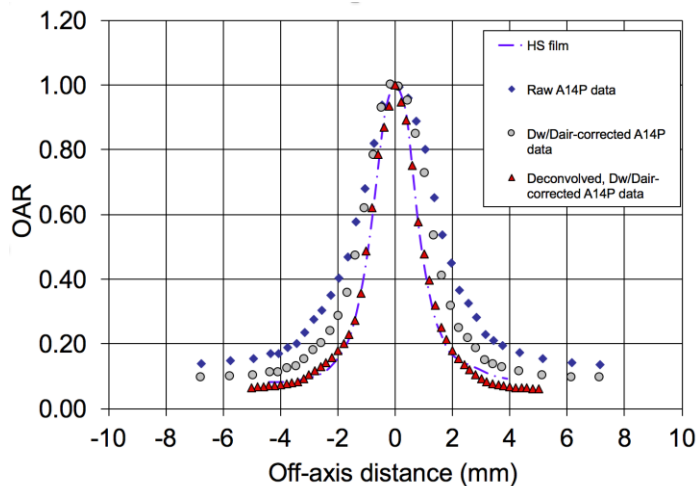


Figure 2: Radiosurgical collimator holder with a small field collimator, attached to the Clinac-18 treatment head.

Figure 4: Schematic drawing of the geometry used for Monte Carlo simulations. The treatment head along with the collimator is simulated with BEAM/EGS4. Source-collimator distance (SCD) is 70 cm. The calculations in phantoms are performed with: (i) the EGSnrc code at source-surface distance (SSD) 100 cm for phase space validation and (ii) with the XVMC code at source-axis distance 100 cm for dose distribution calculations.

McGill circa 2000 (presented at the 2001 McGill Workshop 10 days after 9/11)

Large dosimetric discrepancies!



Back in 2001 – first McGill Workshop!

Data: Paskalev et al, 2001, 2002

DOSRZ run on a A14P simplified model

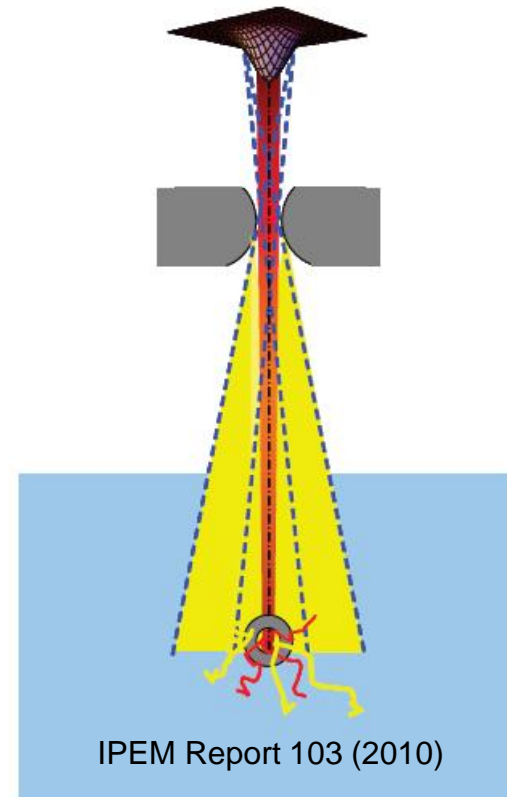
Modeling of electric field distribution was necessary!

Separate deconvolution!

Small photon field conditions

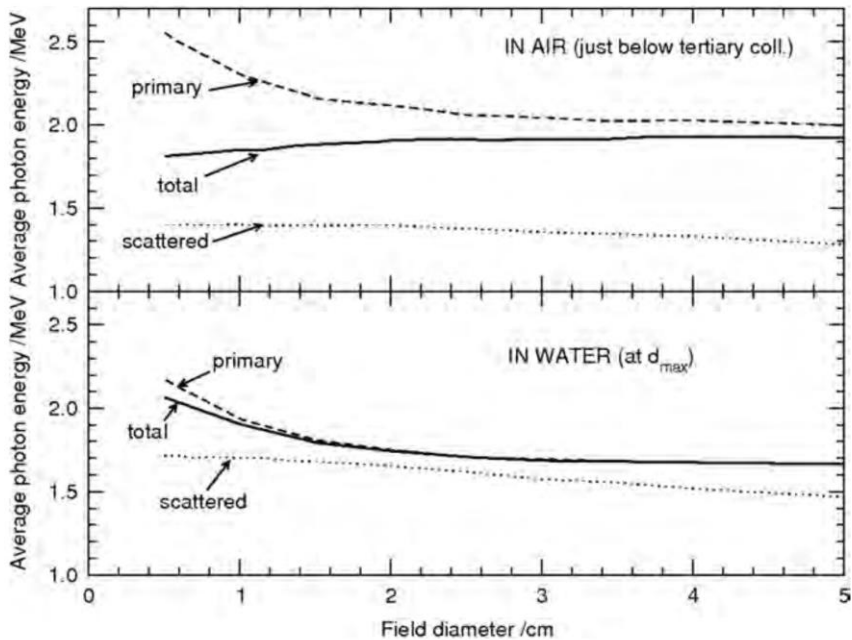
IAEA TRS 483 – ICRU 91

- Beam-related small-field conditions
 - the existence of lateral charged particle disequilibrium
 - change in photon fluence spectrum -> beam quality
 - partial geometrical shielding of the primary photon source as seen from the point of measurement
- Detector-related small-field condition
 - detector size compared to field size

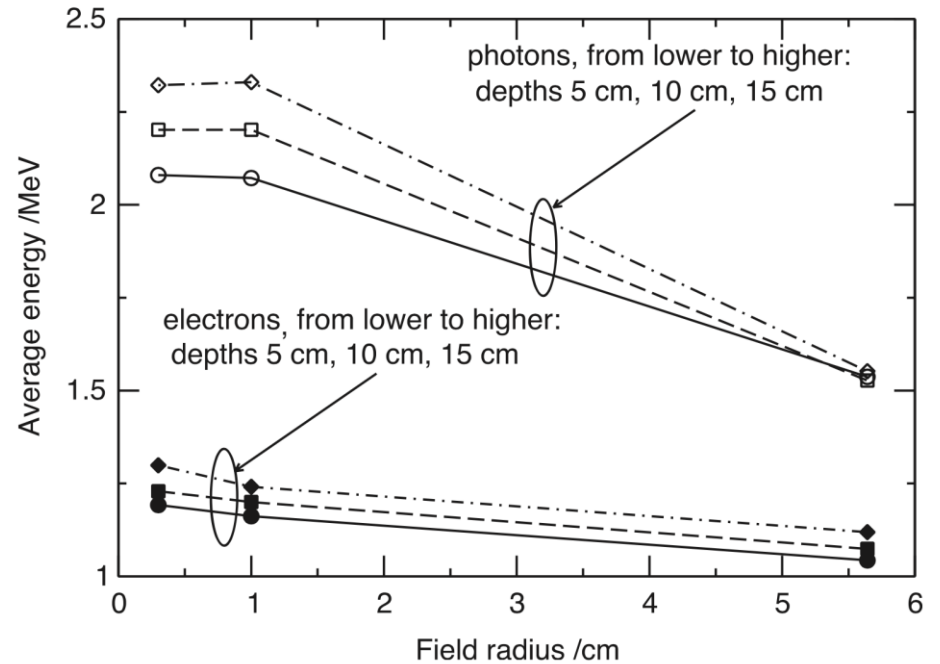


Small beams

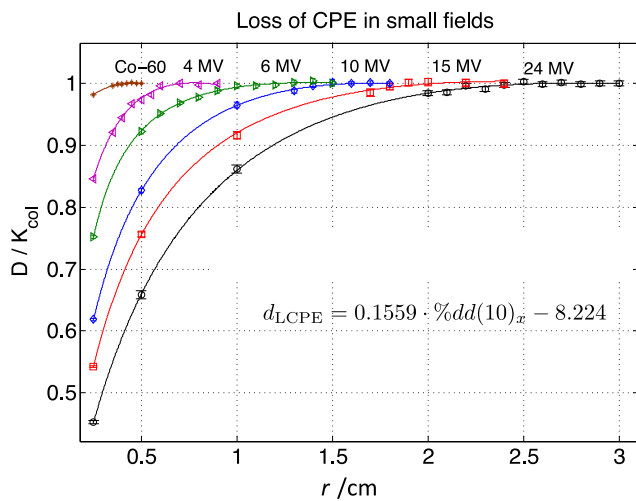
Data from Verhaegen et al 1998



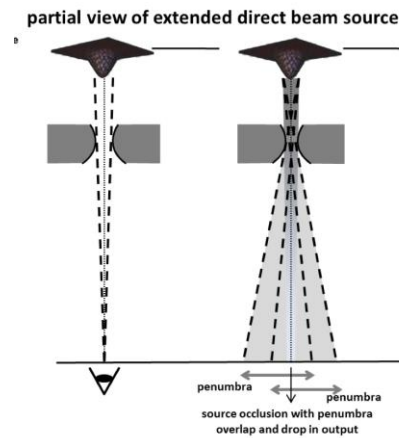
Data from Sanchez-Doblado, et al 2003



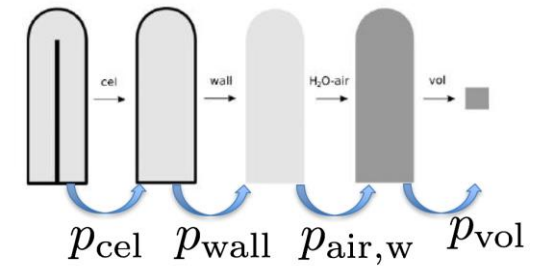
Textbook characterization of small beams



Radiation disequilibrium

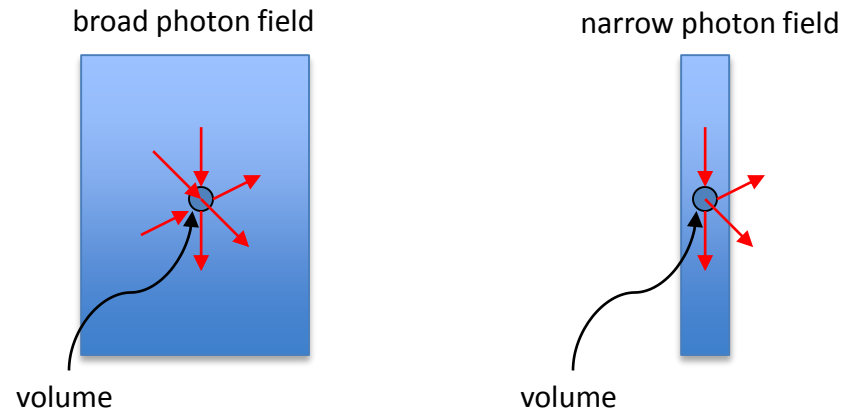


Source occlusion



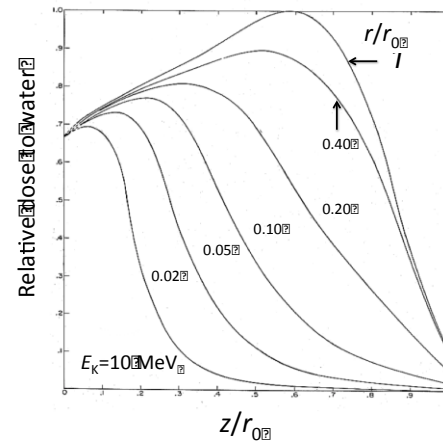
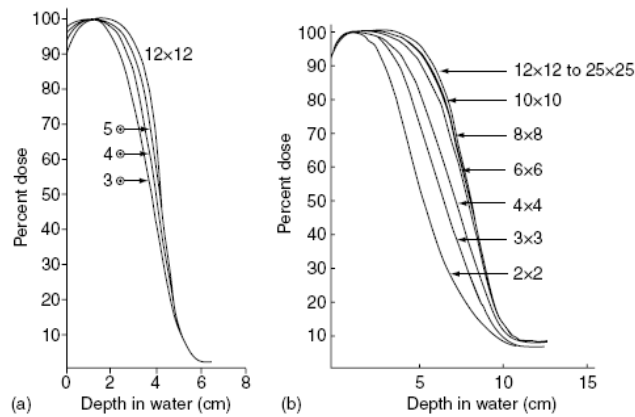
Detector correction factors

Lateral charged particle loss



A small field can be defined as a field with size smaller than the “lateral range” of charged particles

Lateral charged particle loss

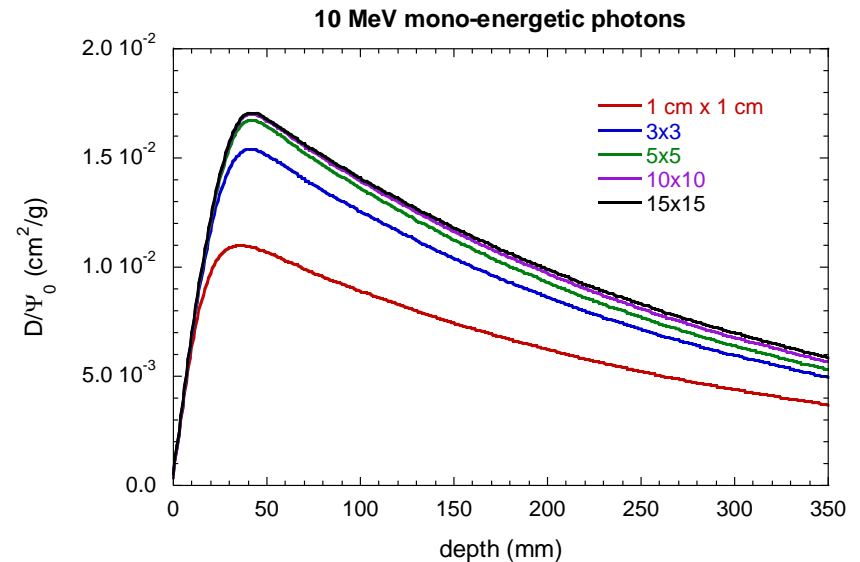
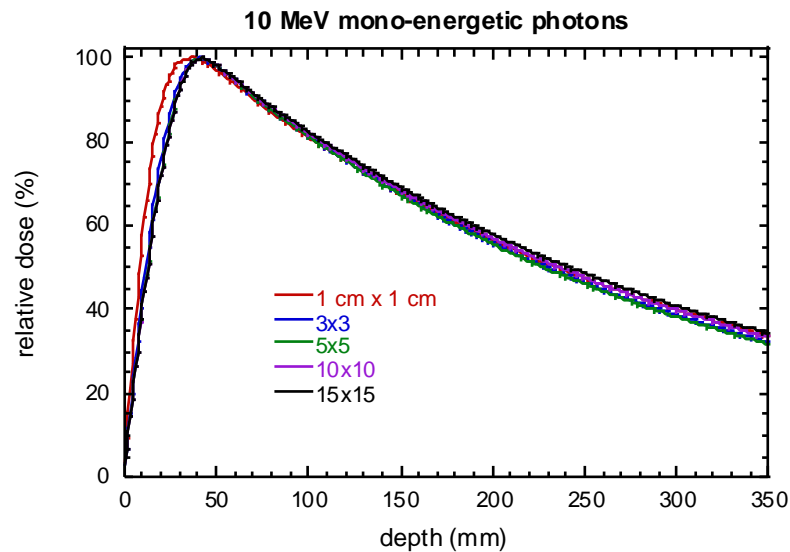


Berger and Seltzer (1982)

An electron beam can be considered "wide" when its PDD is independent of the size of the field.
 The transition to non-equilibrium conditions occurs at $r \approx r_0$ the CSDA range

Slide courtesy: P. Andreo

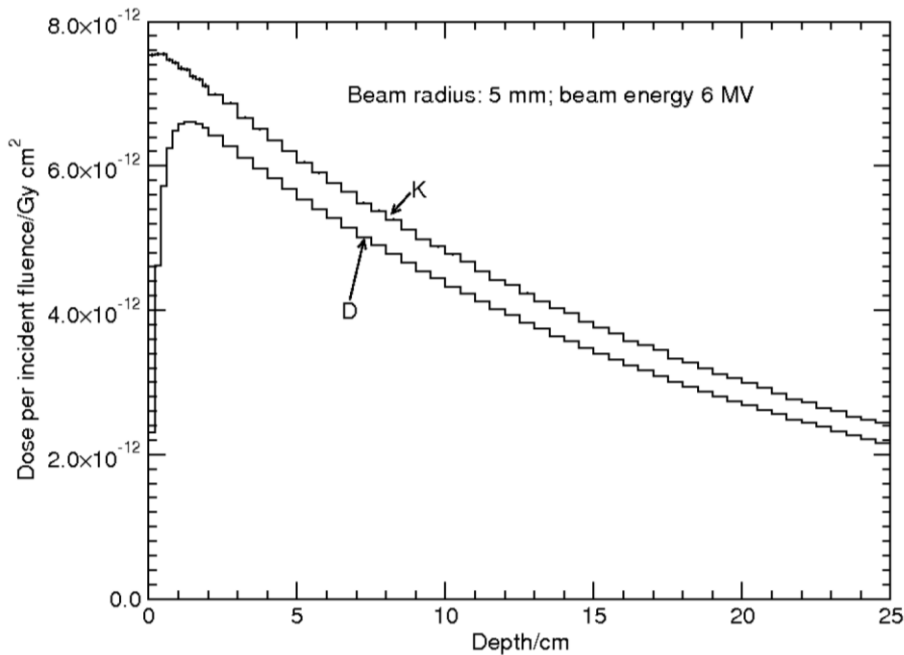
Lateral charged particle loss



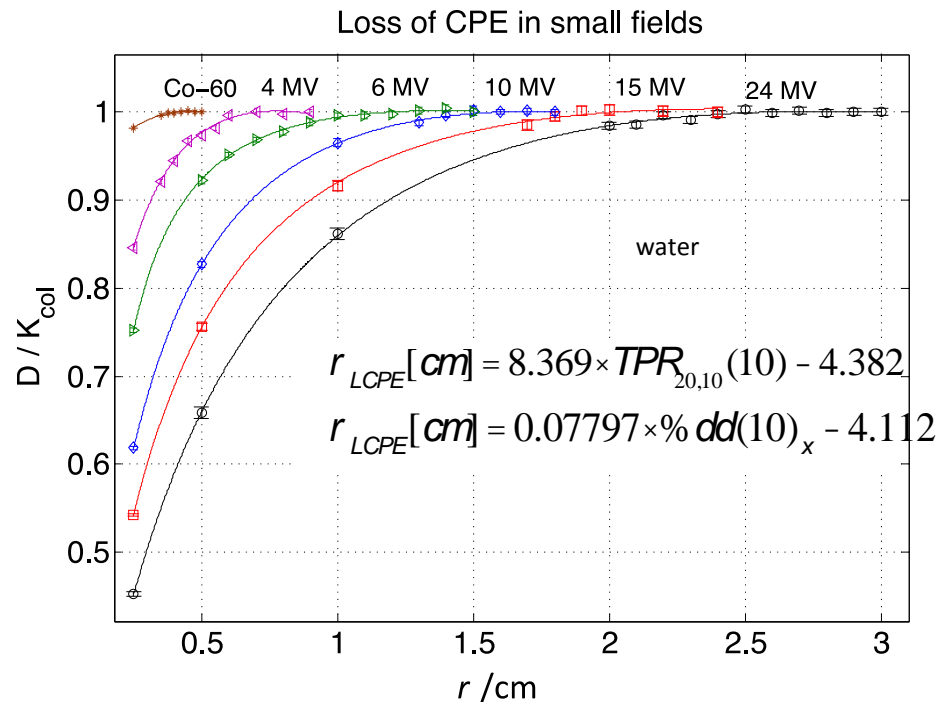
In photon beams the transition from TCPE to non-equilibrium as a function of field size is less abrupt.

Slide courtesy: P. Andreo

Lateral charged particle loss



In small fields there is **no depth** at which $D > K_{col}$

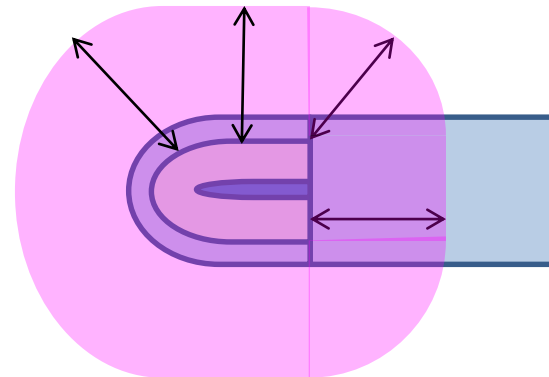


msr field versus small field

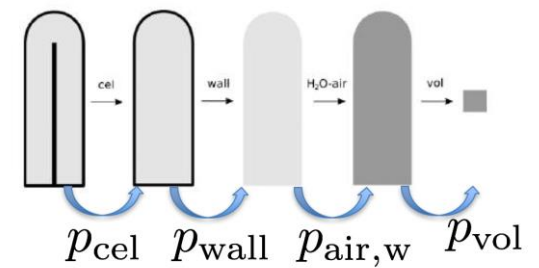
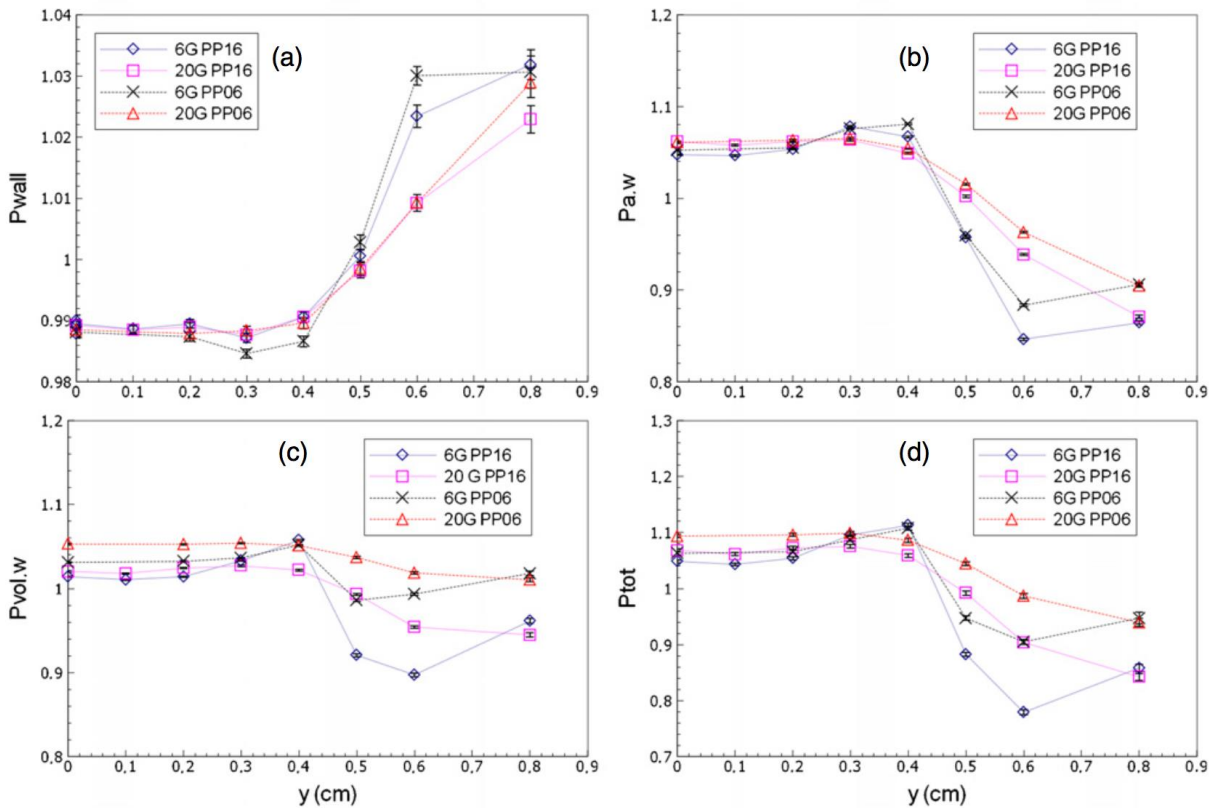
- *msr*: Largest possible reference field less than or equal to 10 x 10 cm² that can be realized on a machine and that is used for calibration $f_{msr} \geq 2r_{LCPE} + \Delta$
- Small field: one of the edges of the detector is less than a lateral charged particle equilibrium range (r_{LCPE}) away from the edge of the field

$$r_{LCPE} [cm] = 8.369 \times TPR_{20,10}(10) - 4.382$$

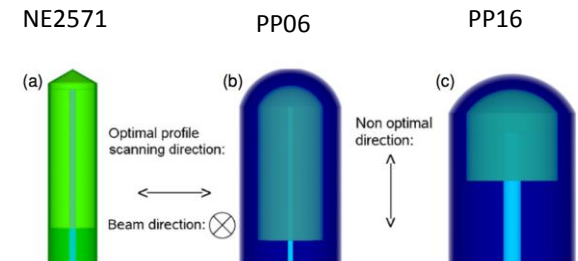
$$r_{LCPE} [cm] = 0.07797 \times \% dd(10)_x - 4.112$$



Detector response

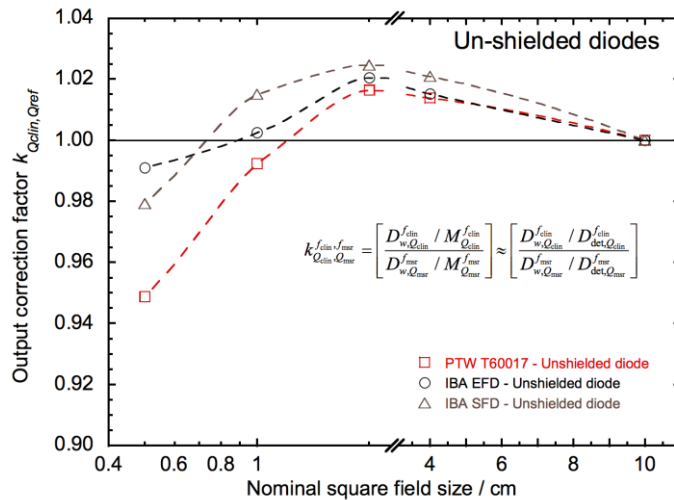
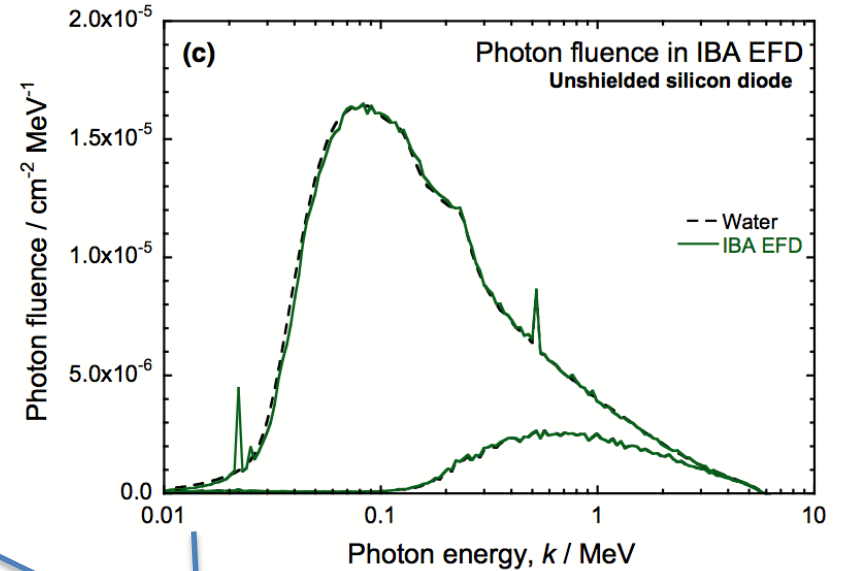
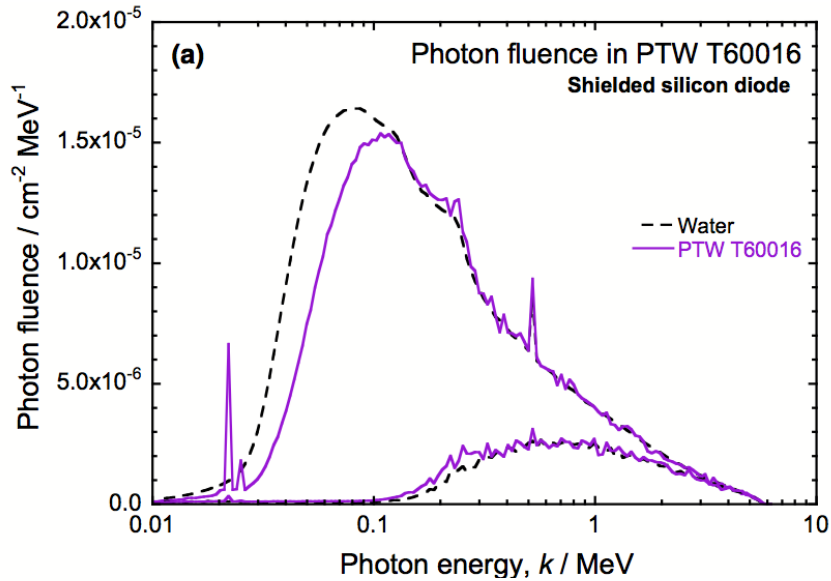


PP16 = 31016
PP06 = 31006



Crop et al 2009

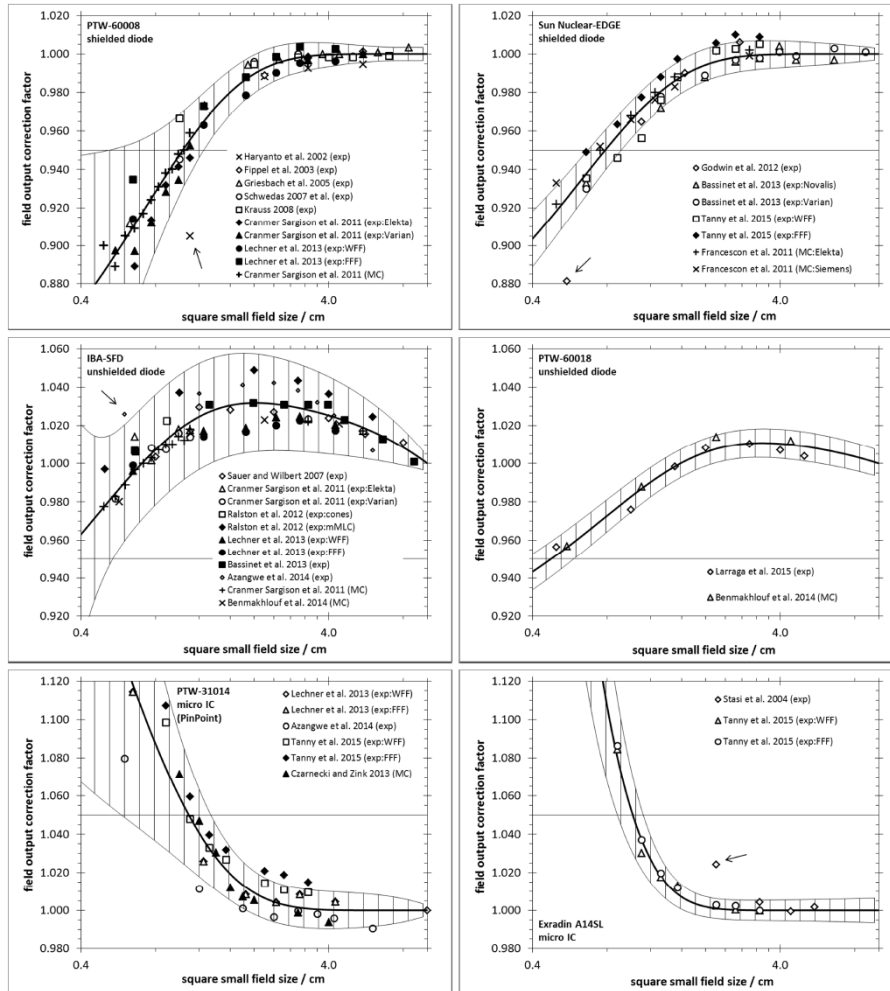
Spectra inside detectors & response



Benmakhlouf and Andreo, 2017

Benmakhlouf and Andreo, 2013

TRS 483 Small field output correction factors



Field size specification using FWHM inplane and crossplane!

ICRU Report 91 follows the TRS 483 recommendations for the measurement of **output factors** for small fields

Remarks:

1. Uncertainties are $k=2$
2. Corrections > 5% are not recommended

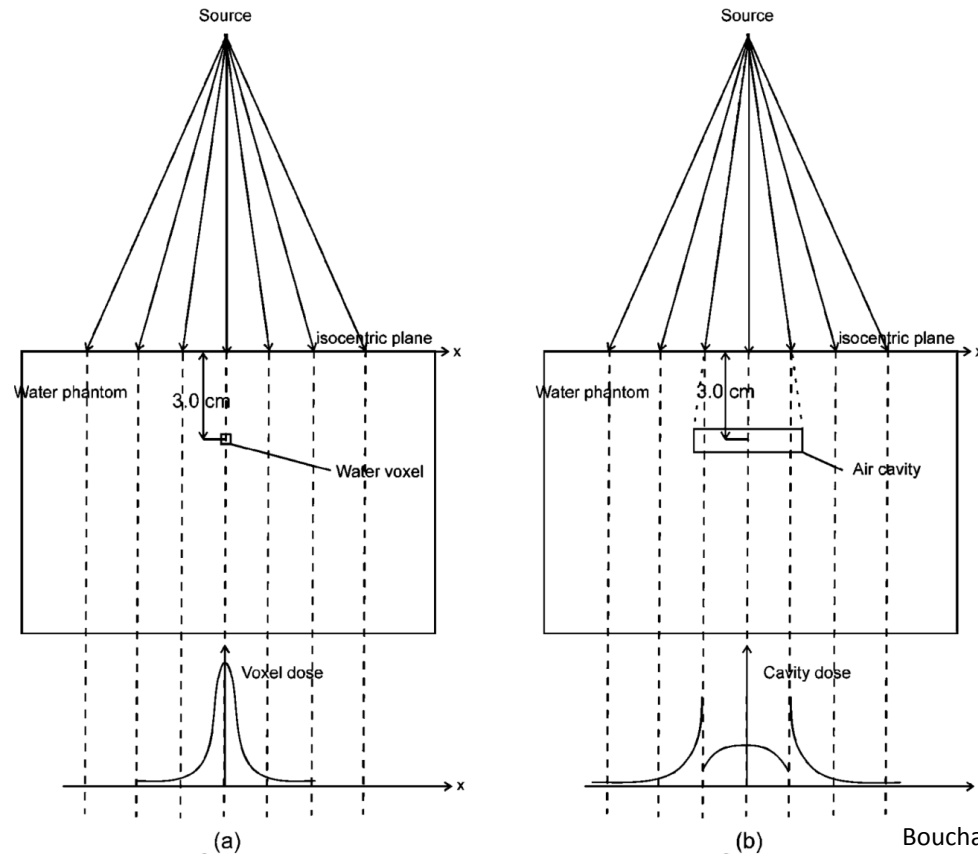
Questions post TRS-483 small field report

- More data is needed (phantoms, GammaKnife)
- Do we still need a calibration solution for modulated fields?
- Intermediate field calibration for machines that do not fulfill *msr* calibration conditions.

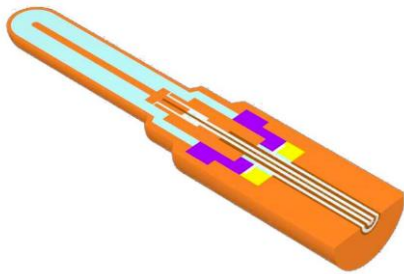
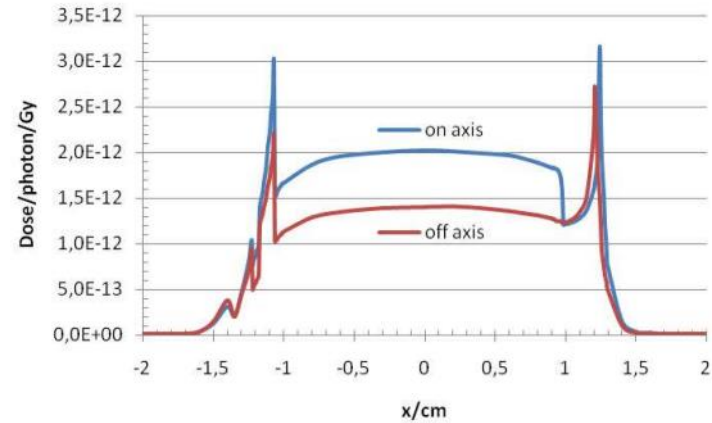
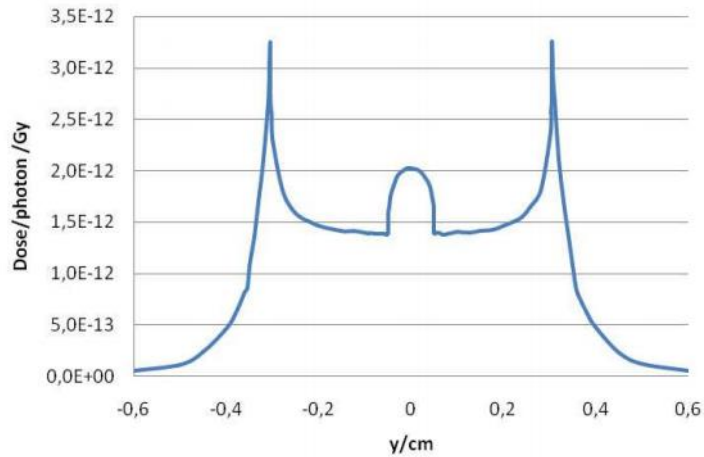
Related question

- Do we need alternative techniques to determine relative output?
- Do we need alternative techniques to calibration “sub-*msr*” fields?

Insights gained using MC: Decomposing the detector response



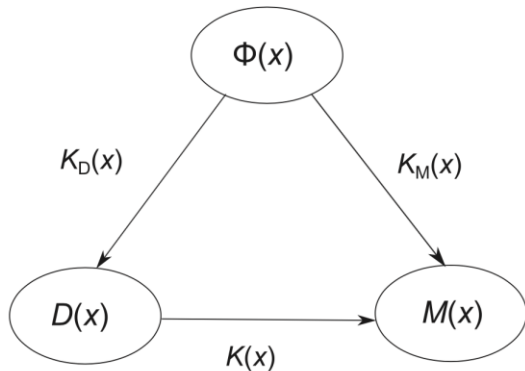
Decomposing detector response



The “batman” mask

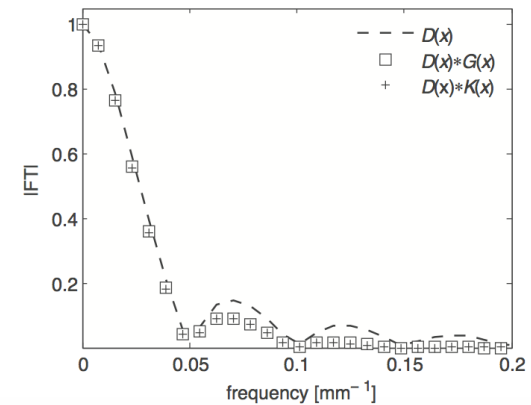
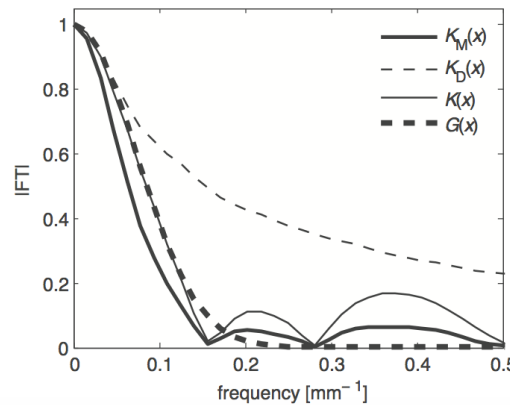
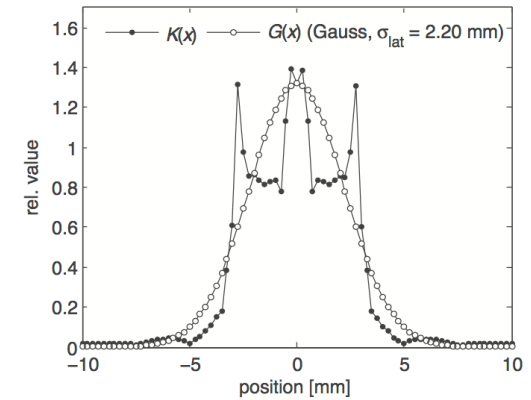
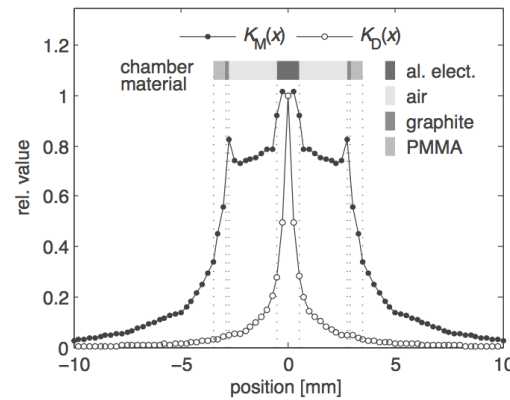
Tantot and Seuntjens, 2008

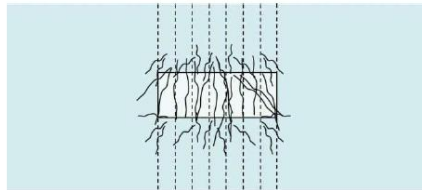
Decomposing the detector response



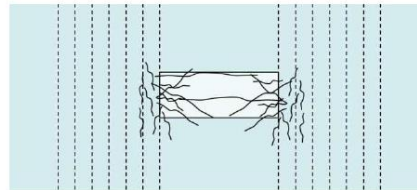
Looe et al, 2012

Gaussian kernels are a first order approximation

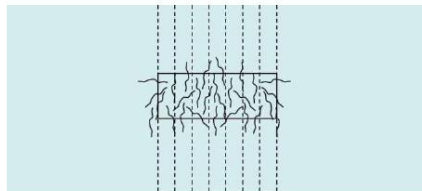


D_{in} D_{out} 

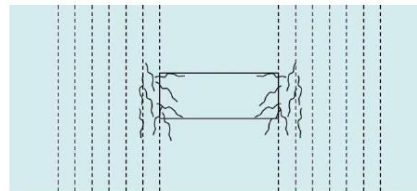
(a)



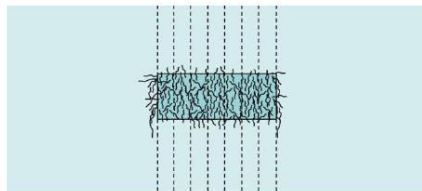
(b)



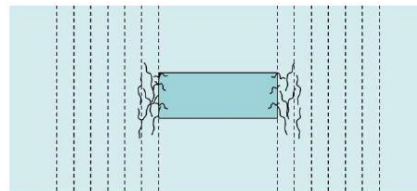
(c)



(d)



(e)



(f)

Water vapour

Liquid water

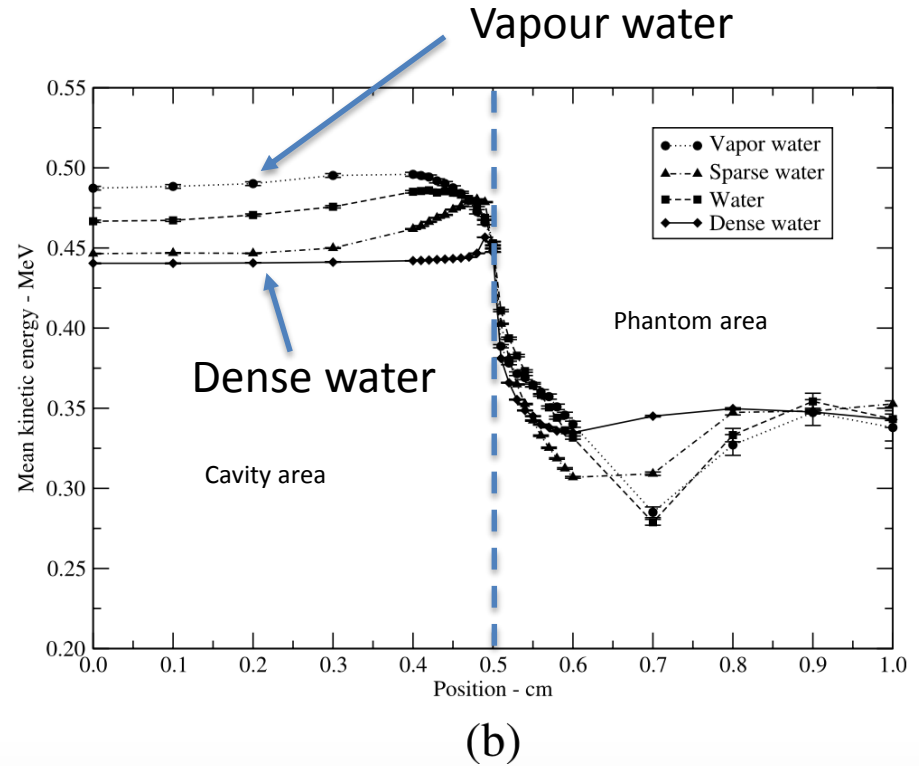
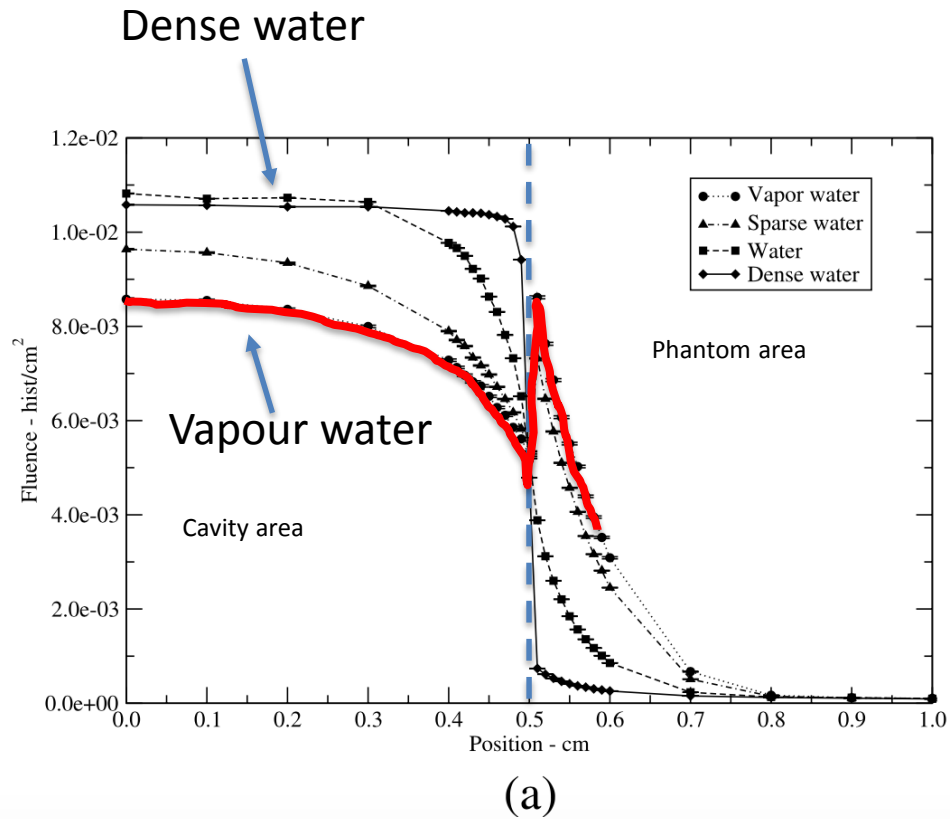
Dense water

$$D_{cav}^{small\ field} = w_1 D_{in} + w_2 D_{out}$$

$$= (w_1 - w_2) D_{in}(\rho) + w_2 D_{Fano}$$

Bouchard et al 2015AB

Batman and Fano



Fluence function and mean kinetic energy in a 5 mm radius cavity filled with different densities under Fano conditions

$E=1.25$ MeV

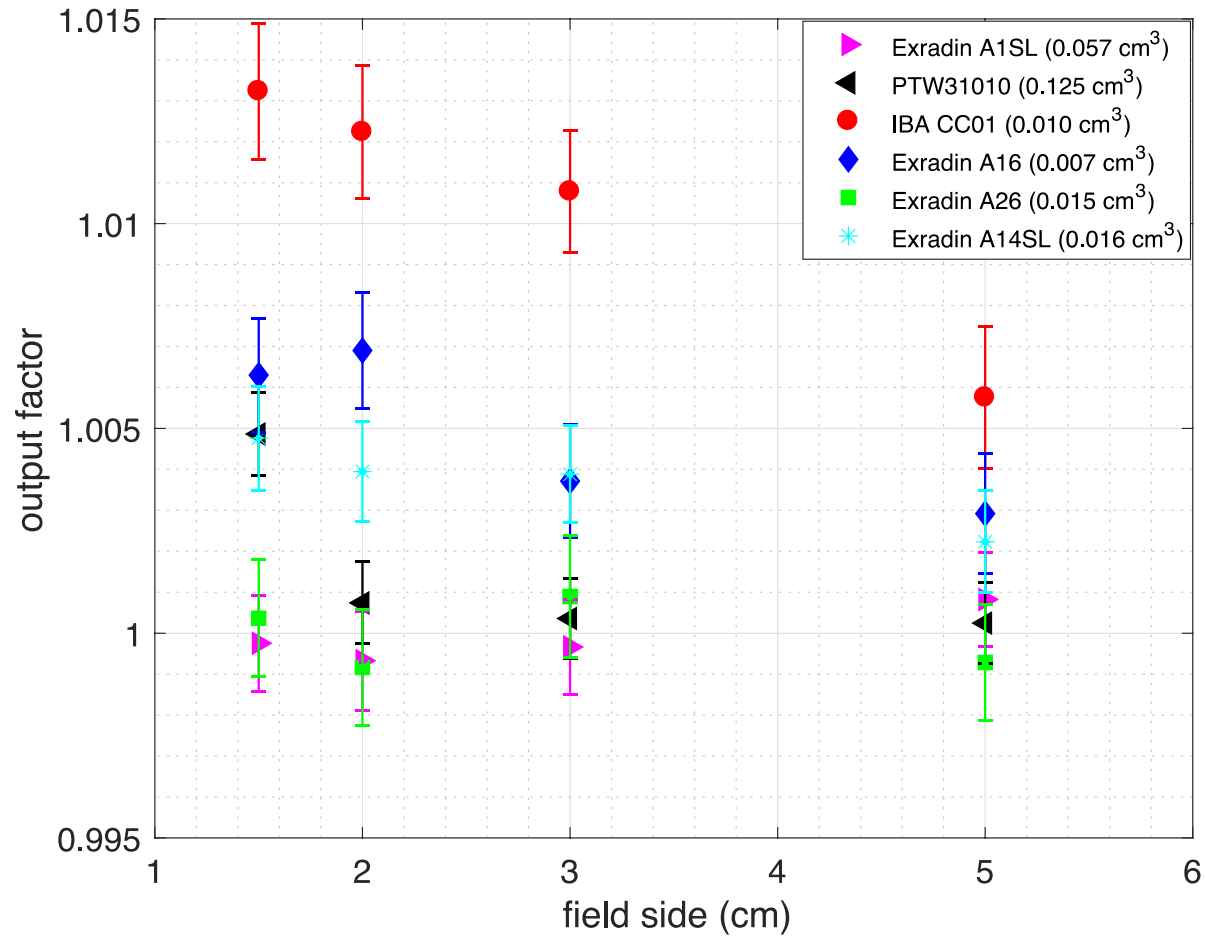
Bouchard et al 2015AB

Field sizes between *msr* and small

- The LCPE criterion is violated for field sizes below

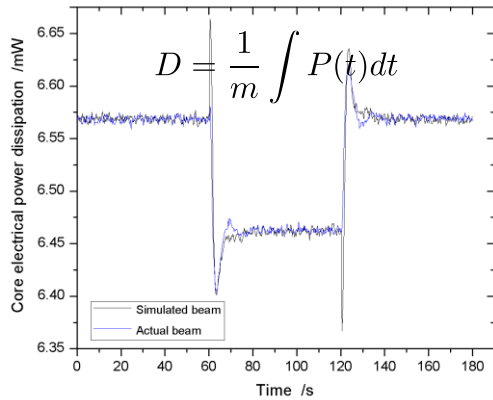
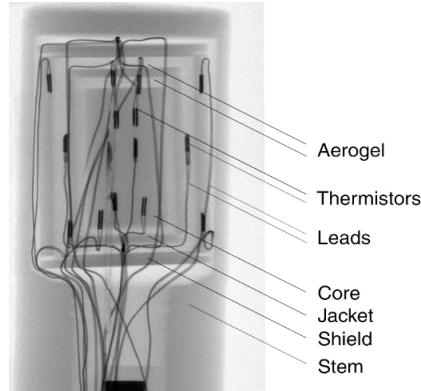
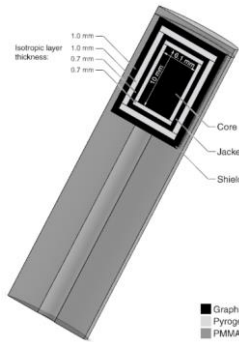
$$2r_{\text{LCPE}} + \Delta$$

- For 6 MV and reference class chambers this limits the smallest *msr* field to be larger than ~ 4 cm
- New upcoming radiation equipment may/will not have calibration fields this large
- To what extent can we live with correction factors that start to contain some more significant perturbation effects?

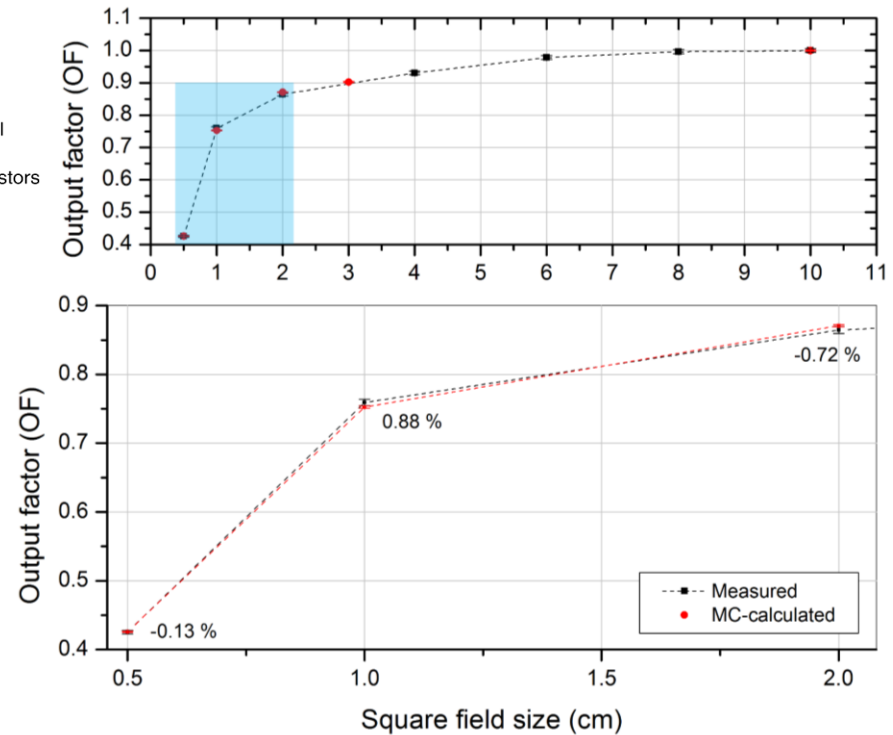


Preliminary Mirzakhania et al, 2017

More advantageous reference detector?

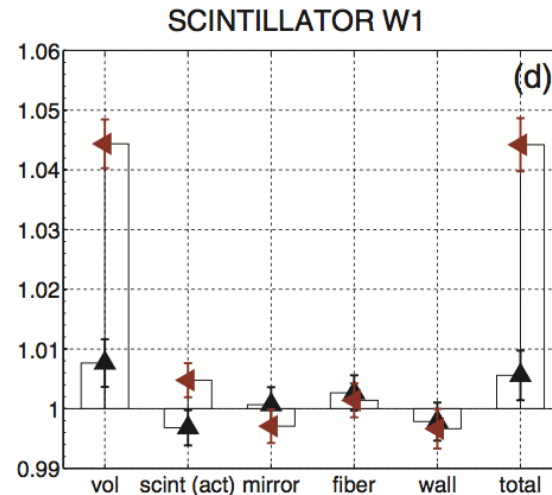
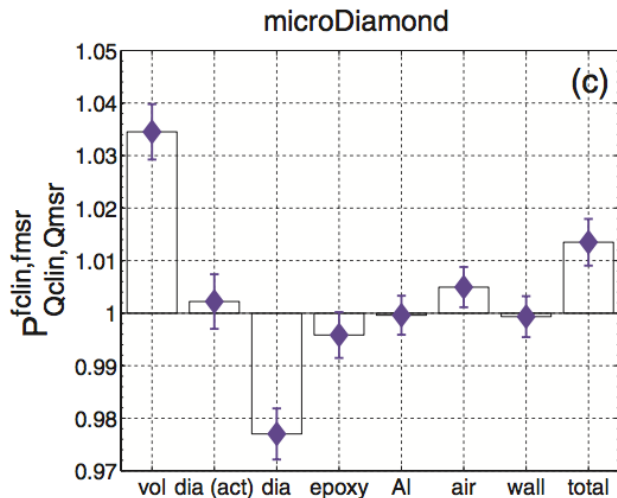
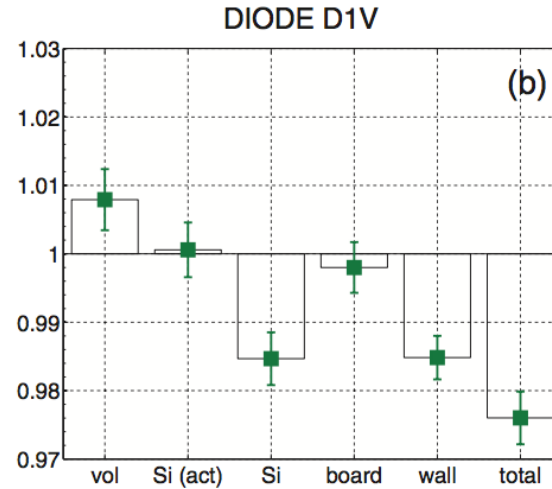
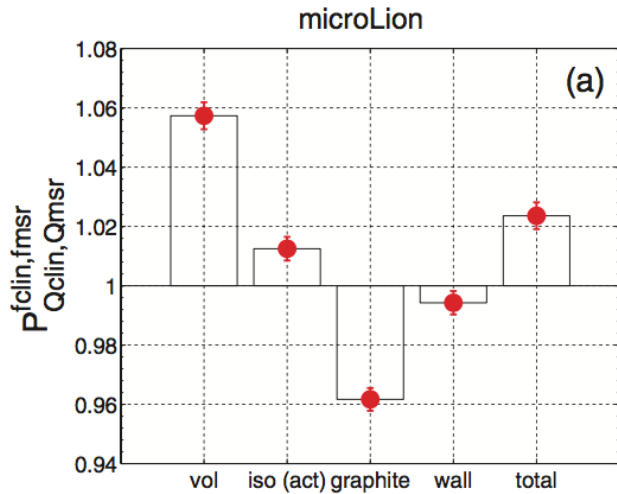


self-calibrate & self-check



J. Renaud et al, 2017

Playing with compensated detector designs



Other authors:
Underwood et al
and others

Papaconstadopoulos
et al, 2014

GammaKnife calibration



ABS parallel



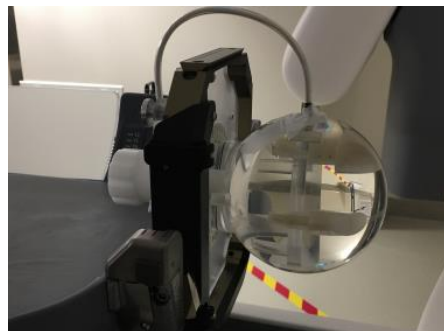
ABS perpendicular



ABS 45°



Solid Water



Lucy 0°



Lucy 270°

GammaKnife *msr* correction factors

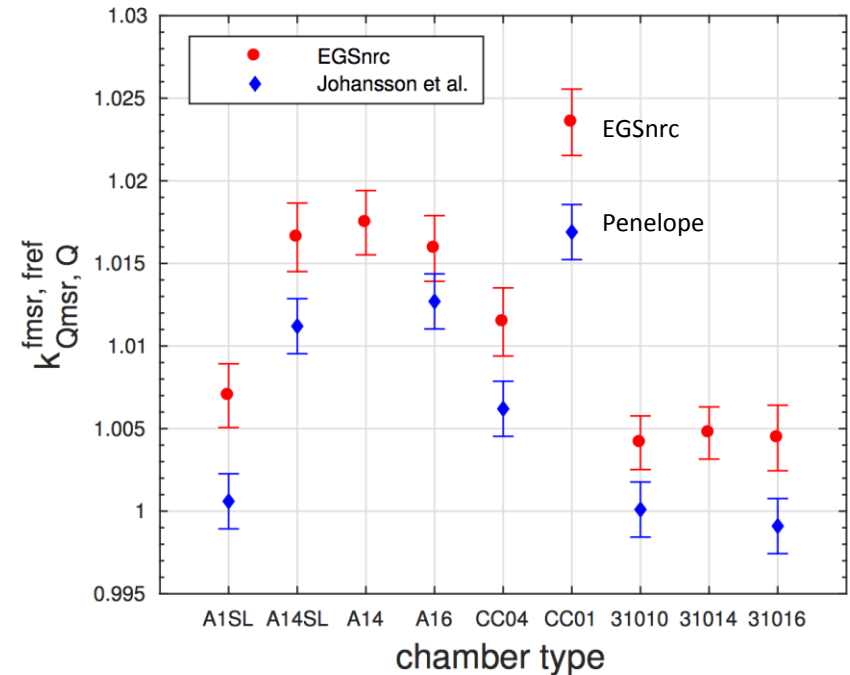
TABLE I. Chambers specifications used in this study.

chamber type	sensitive volume (cm ³)	cavity length (mm)	cavity radius (mm)	electrode material
Exradin A1SL ^a	0.057	5.7	2.1	C-552
Exradin A14SL ^a	0.016	2.1	2.1	C-552
Exradin A14 ^a	0.016	2.0	2.0	C-552
Exradin A16 ^a	0.007	1.7	1.2	C-552
IBA CC04 ^a	0.040	3.6	2.0	C-552
IBA CC01 ^b	0.010	3.6	1.0	Steel
PTW 31010 ^b	0.125	6.5	2.8	Aluminium
PTW 31014 ^b	0.015	5.0	1.0	Aluminium
PTW 31016 ^b	0.016	2.9	1.45	Aluminium

^a Chambers modeled in both codes: EGSnrc and PENELOPE

^b Chambers modeled in EGSnrc

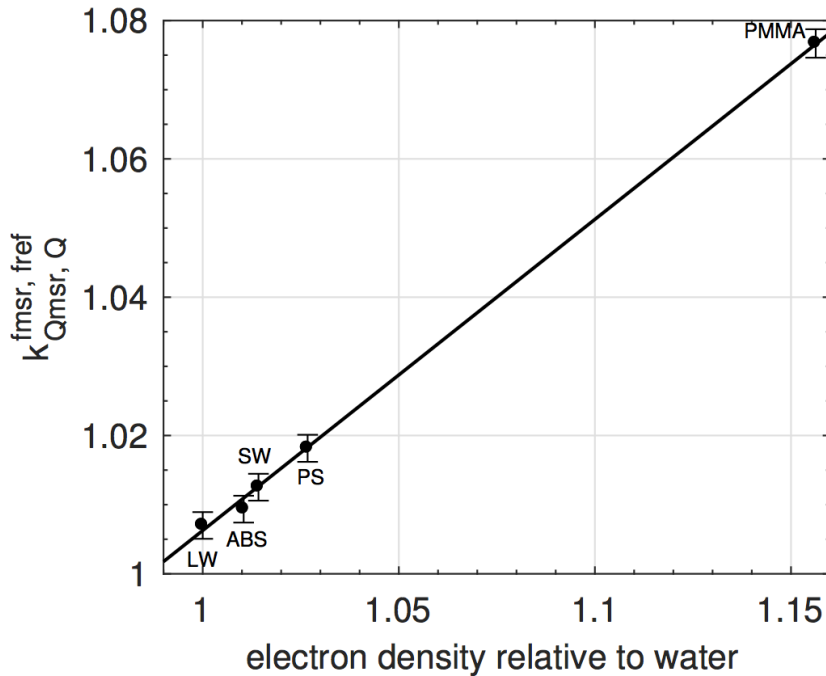
$r_{LCPe} \sim 4$ mm, for a 16 mm field we are close to *msr* limit for the largest chambers.



$$D_{W, Q_{msr}}^{f_{msr}} = M_{Q_{msr}}^{f_{msr}} \cdot N_{D, W, Q_0}^{f_{ref}} \cdot K_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}}$$

Mirzakhania et al, 2017

Phantoms of different plastics

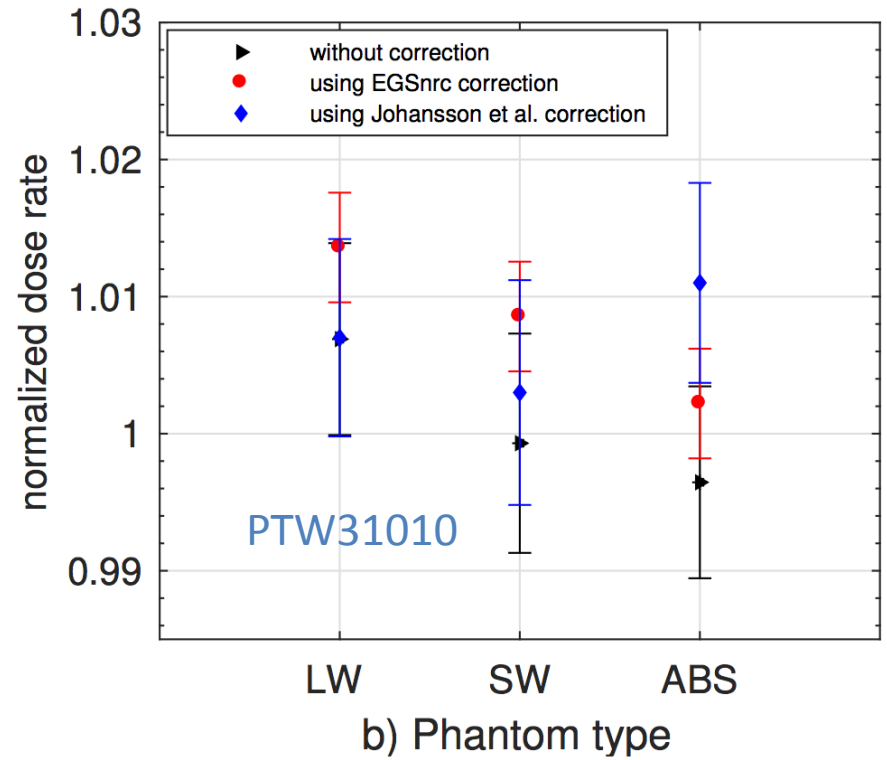
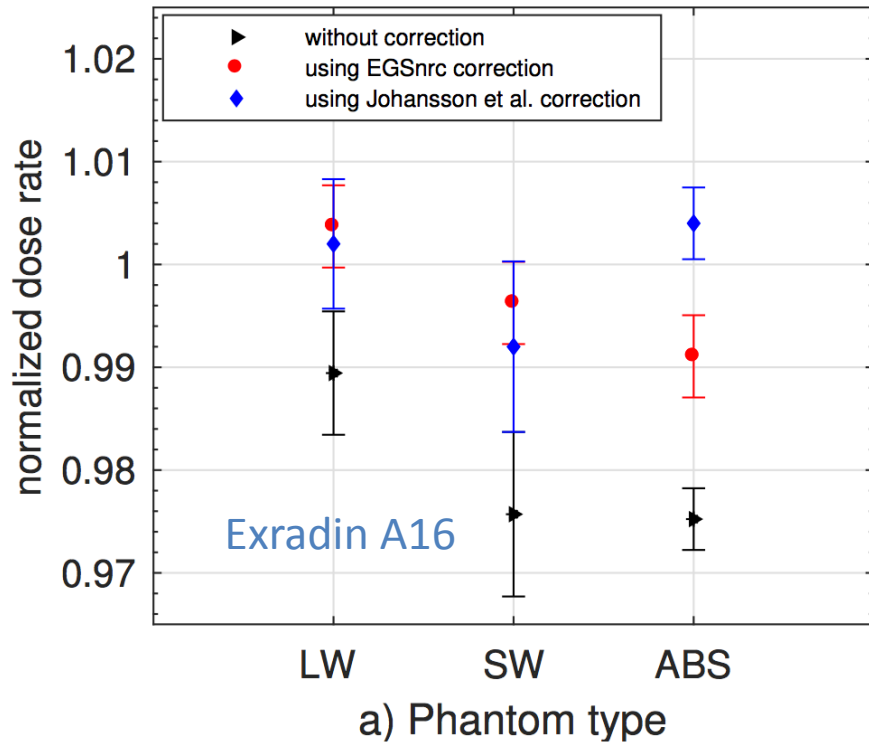


chamber type	parallel		perpendicular	
	$k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}}$	a	$k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}}$	a
Exradin A1SL	1.0070	0.4499	1.0070	0.4234
Exradin A14SL	1.0166	0.4487	1.0105	0.4299
Exradin A14	1.0175	0.4599	1.0148	0.4259
Exradin A16	1.0159	0.4248	1.0174	0.4352
IBA CC04	1.0115	0.4316	1.0112	0.4263
IBA CC01	1.0235	0.4445	1.0354	0.4329
PTW 31010	0.9976	0.4082	1.0307	0.4421
PTW 31014	0.9986	0.4280	1.0169	0.4327
PTW 31016	1.0044	0.4340	1.0072	0.4387

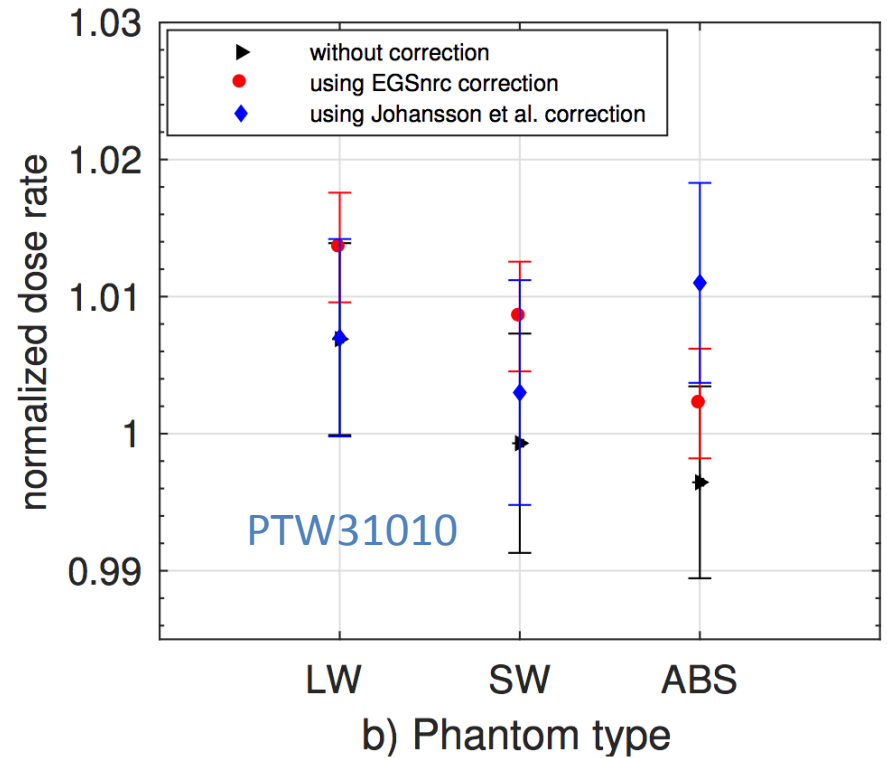
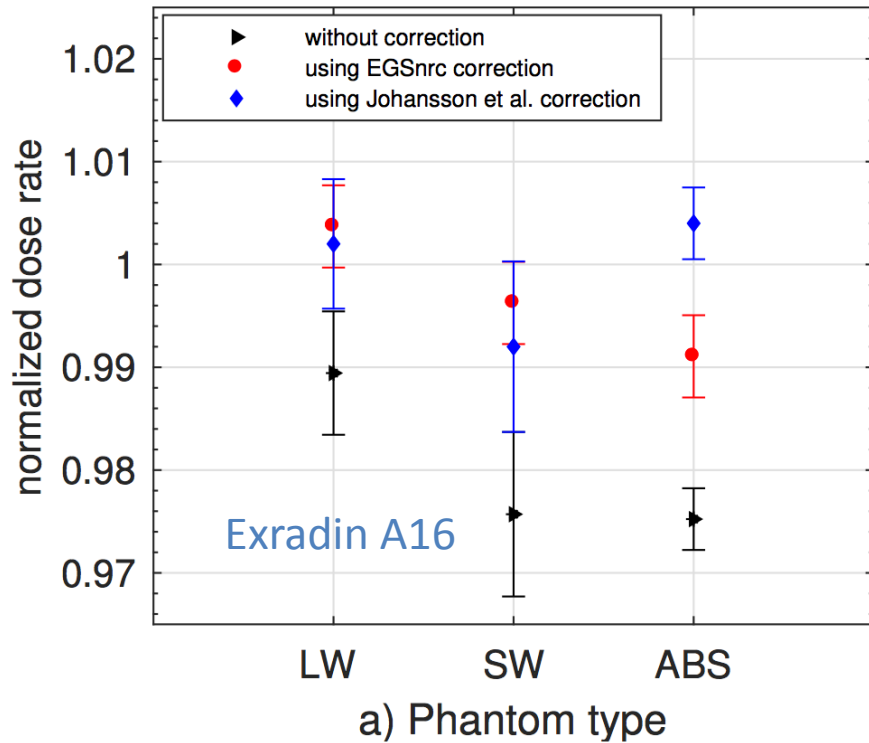
$$k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}} = (\rho_e^{rel} - 1)a_{ch} + k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}}$$

Single global fit to all phantom e-
density dependence,
 $b=0.4285 \pm 2.5\%$

$$k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}} = k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}} k_{Q_{msr}}^{w, plastic} = k_{Q_{msr}, Q_0}^{f_{msr}, f_{ref}} [(\rho_e^{rel} - 1)b + 1]$$

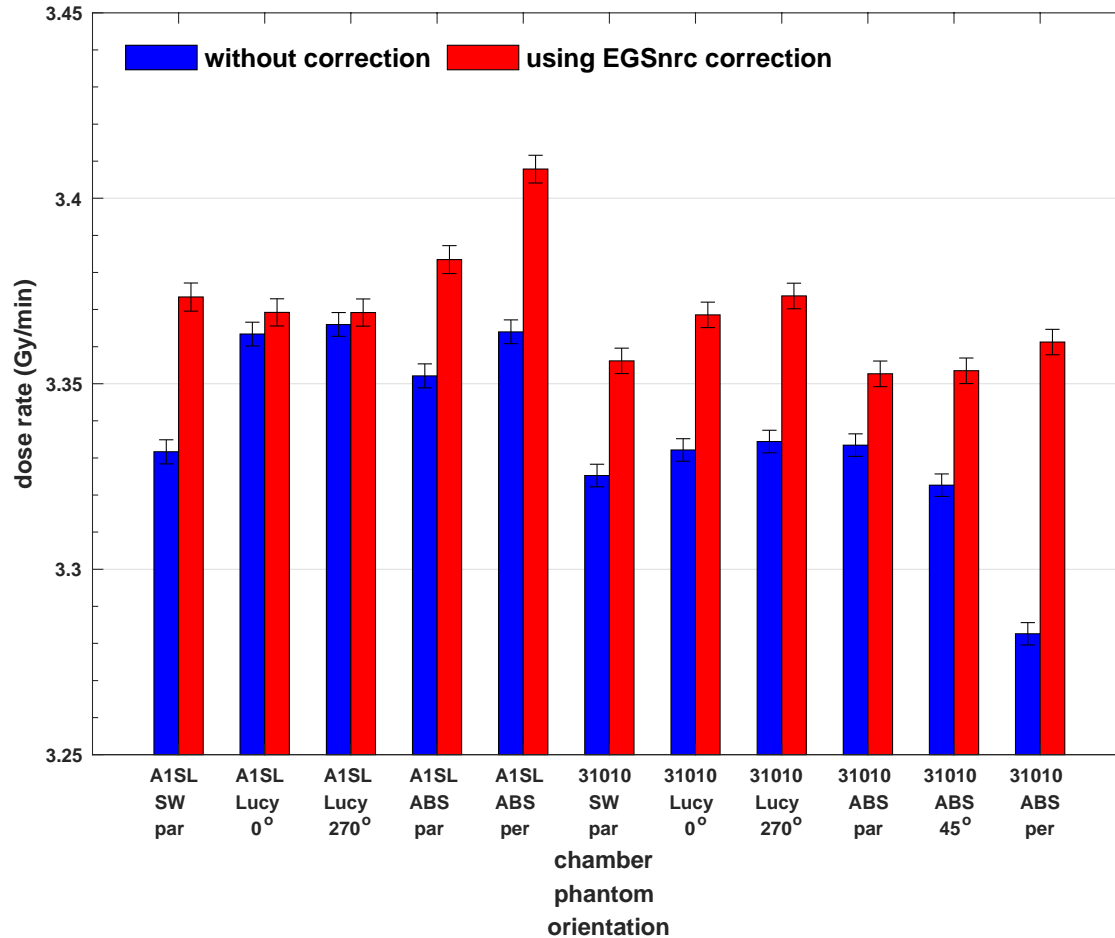


Consistency of intercomparison improves from 1.29% to 0.59%



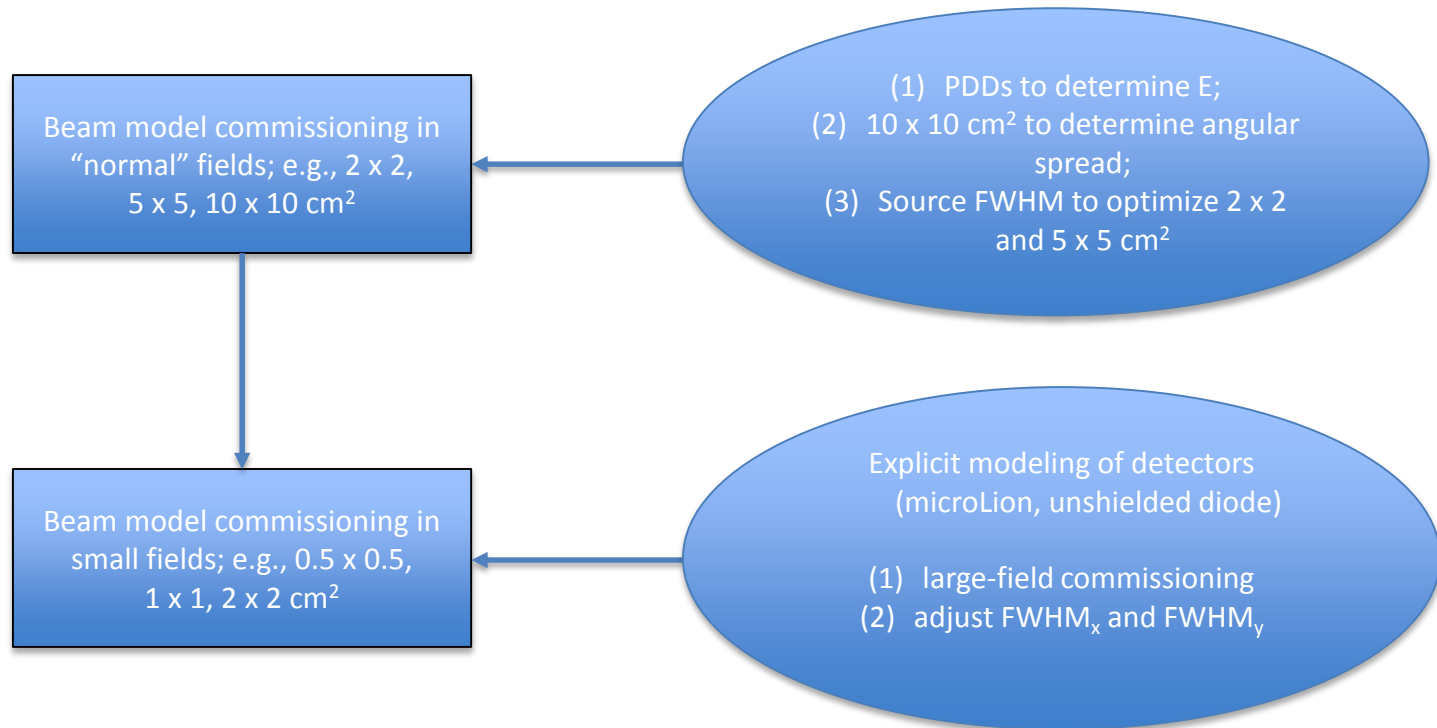
Consistency of intercomparison improves from 1.29% to 0.59%

GammaKnife correction factors



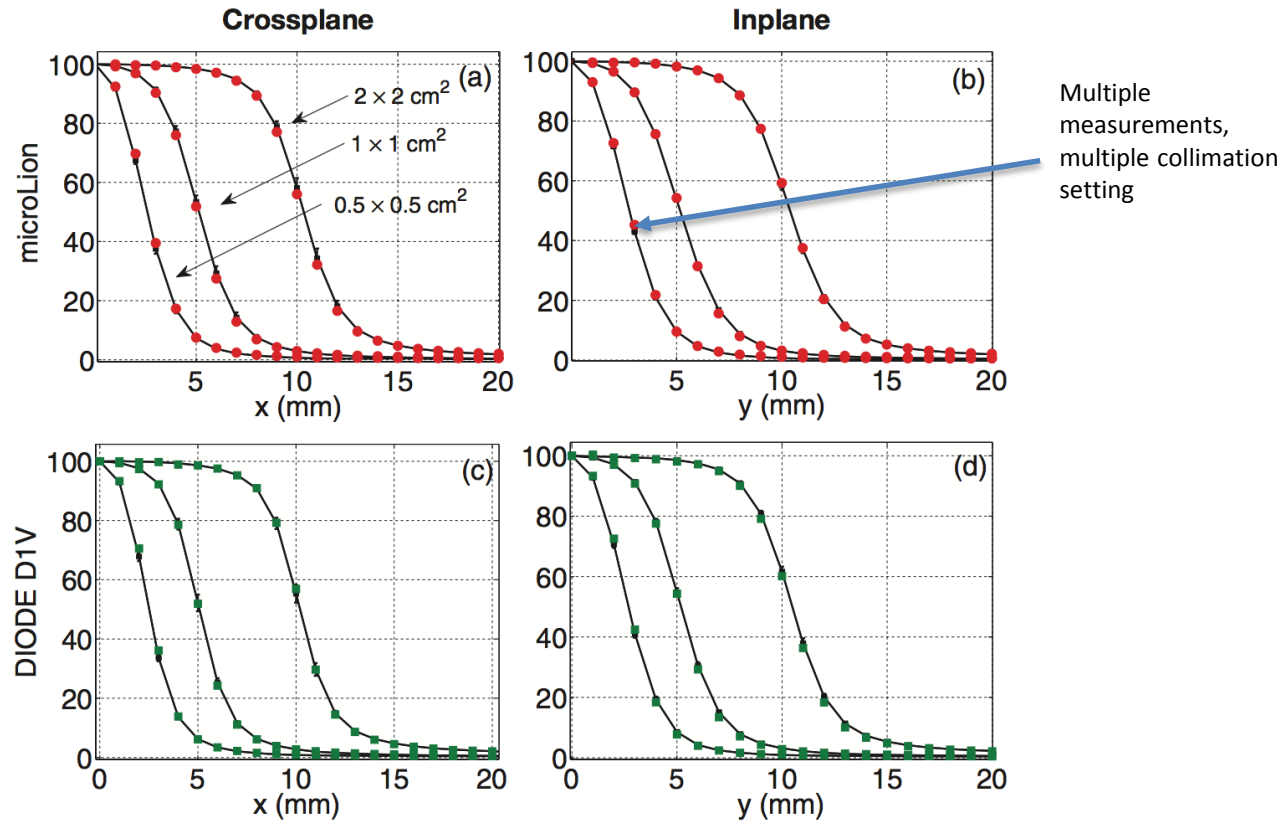
Measurements at the Sunnybrook ICON system

MC beam model commissioning small fields



There is a strong coupling between detector used and optimized MC model parameters

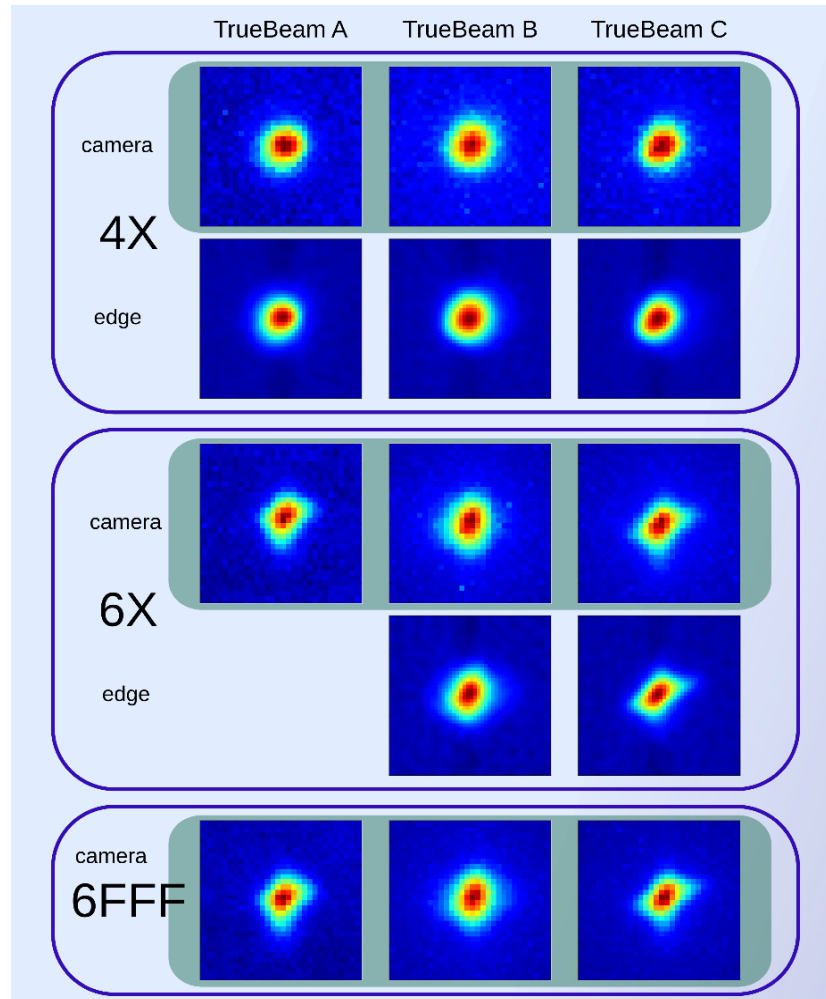
Beam model commissioning small fields



Papaconstadopoulos et al 2015

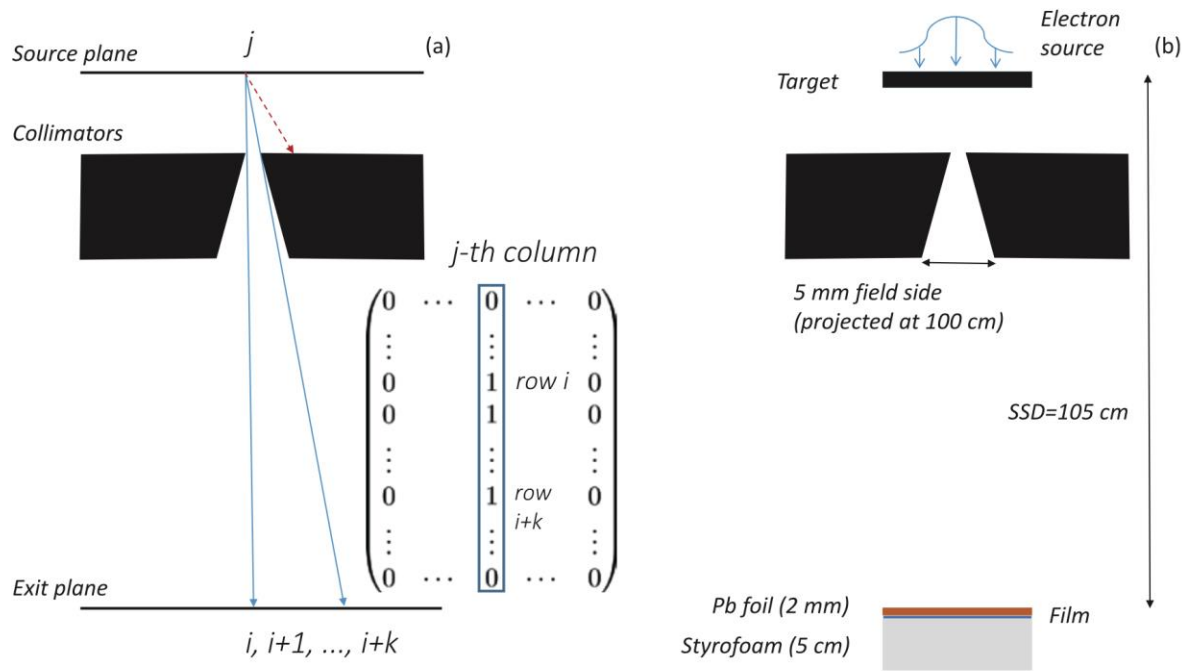
Beam models suitable for SRT planning algorithms are accelerator spot size dependent

Variability in source intensity distribution. Spot sizes range between 2.5 mm and 4.6 mm and the typical spot size is also not perfectly circular



Sawkey et al, 2012

Linac source size and occlusion

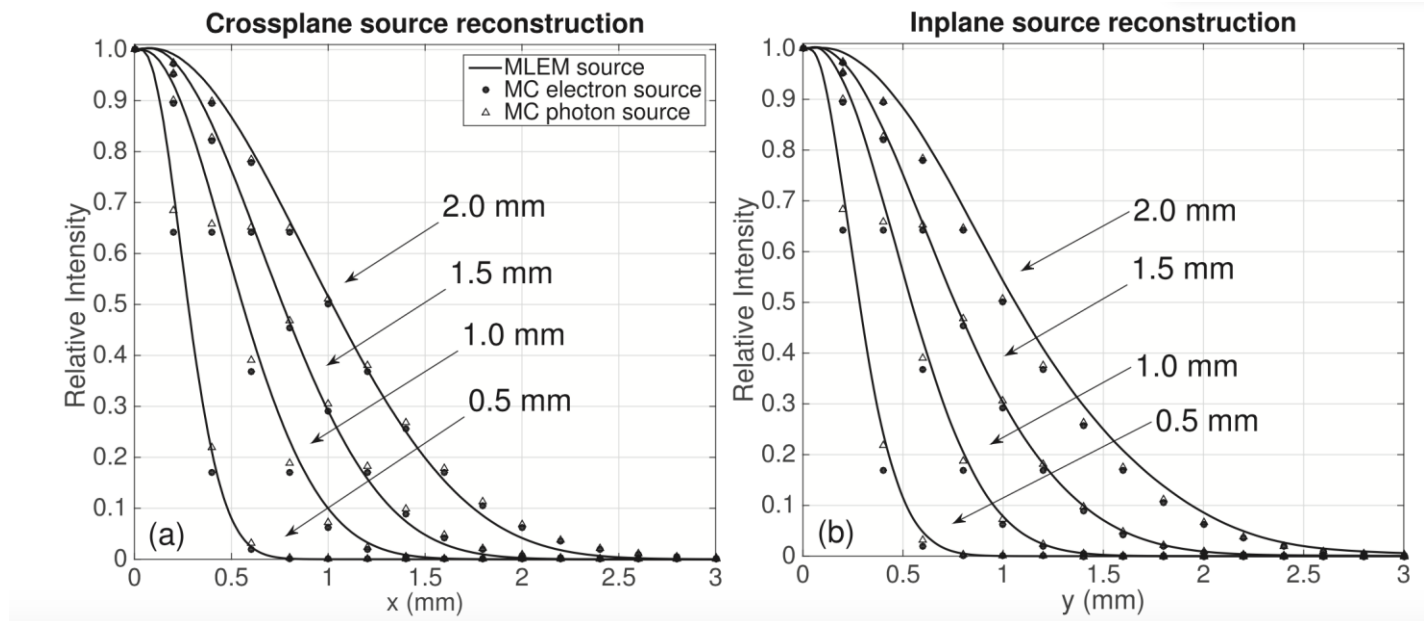


maximum-likelihood expectation-maximization algorithm

$$\varphi^{\text{new}} = \frac{\varphi}{\mathbf{A}^T \mathbf{1}} \mathbf{A}^T \frac{\mathbf{m}}{\mathbf{A} \varphi},$$

Papaconstadopoulos et al 2016

Internal consistency- MLEM vs. MC



Papaconstadopoulos et al 2016

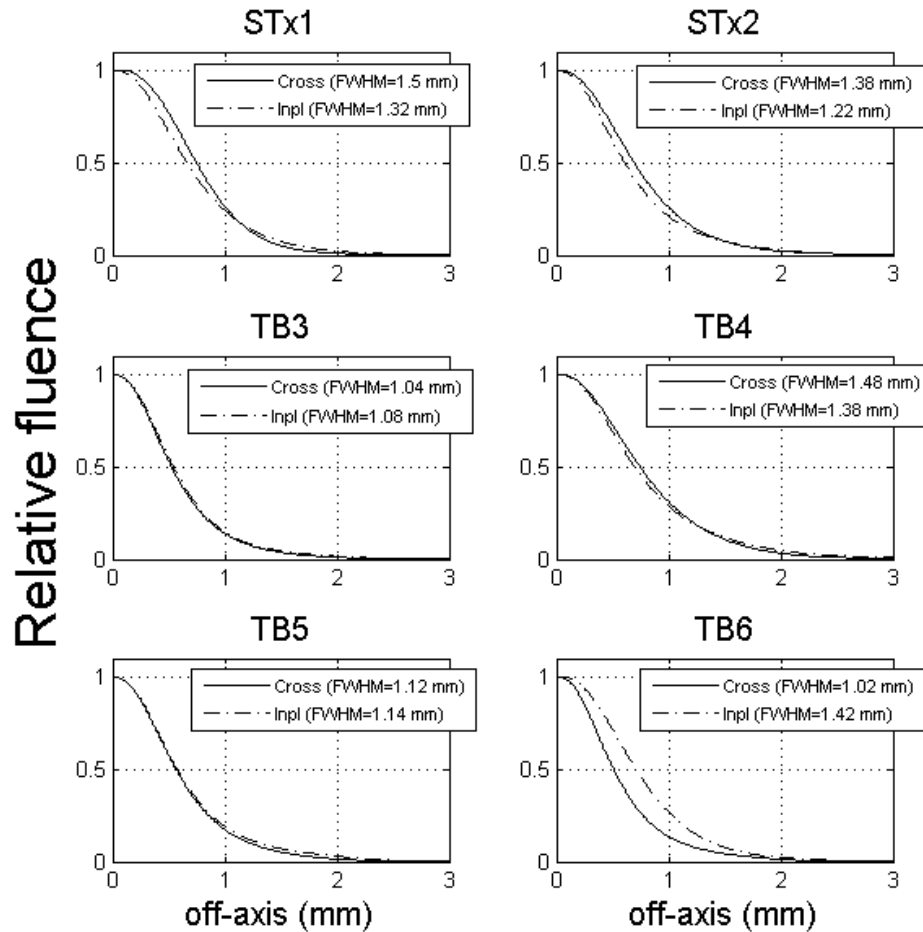
MC versus MLEM on Novalis Tx

Table 3. FWHM and FWTM values of the reconstructed source using the film profile measurements, the MC electron source (incident on the target) and the photon source (at 0.2mm depth in the target) as determined during model commissioning.

		FWHM _x (mm)	FWTM _x (mm)	FWHM _y (mm)	FWTM _y (mm)
Detailed MC commissioning	e^- source	1.25	2.26	1.10	2.00
	γ source	1.28	2.32	1.13	2.04
	rec source	1.22	2.29	1.21	2.32
	σ_{total} (rec source)	0.12	0.27	0.11	0.20
MLEM	σ_{jaw} (rec source)	0.11	0.21	0.10	0.15
	σ_{exp} (rec source)	0.03	0.12	0.04	0.11
	σ_{psf} (rec source)	0.04	0.16	0.01	0.07

Note: The total and component uncertainties of the reconstruction are presented at the 1σ level.

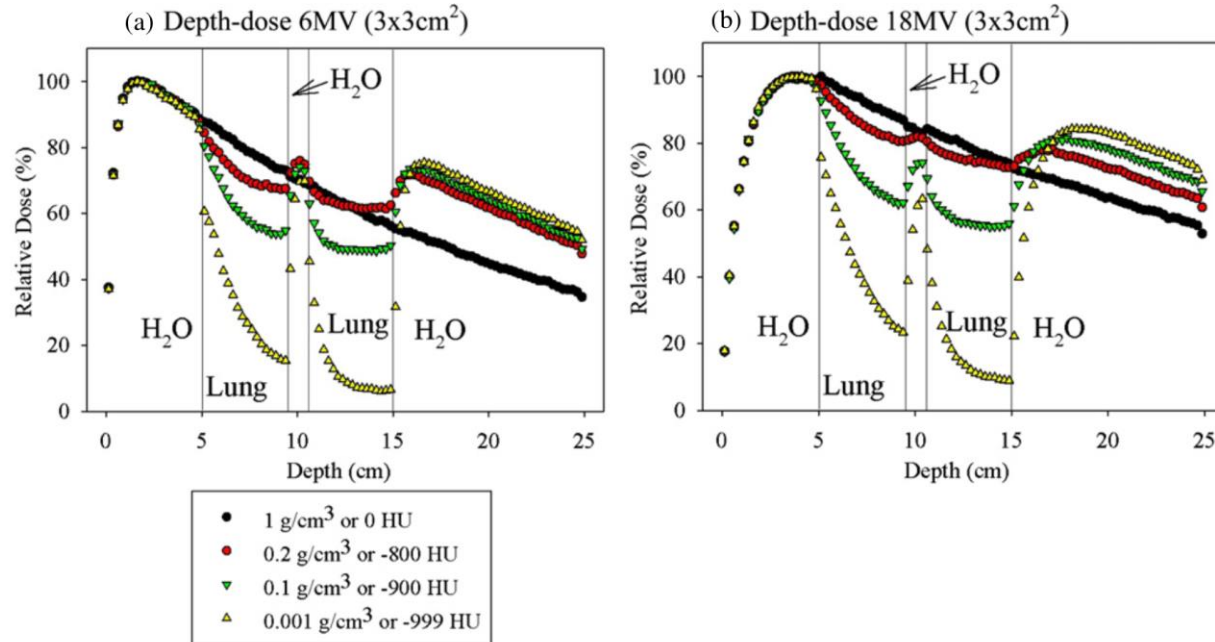
Linac source size variation



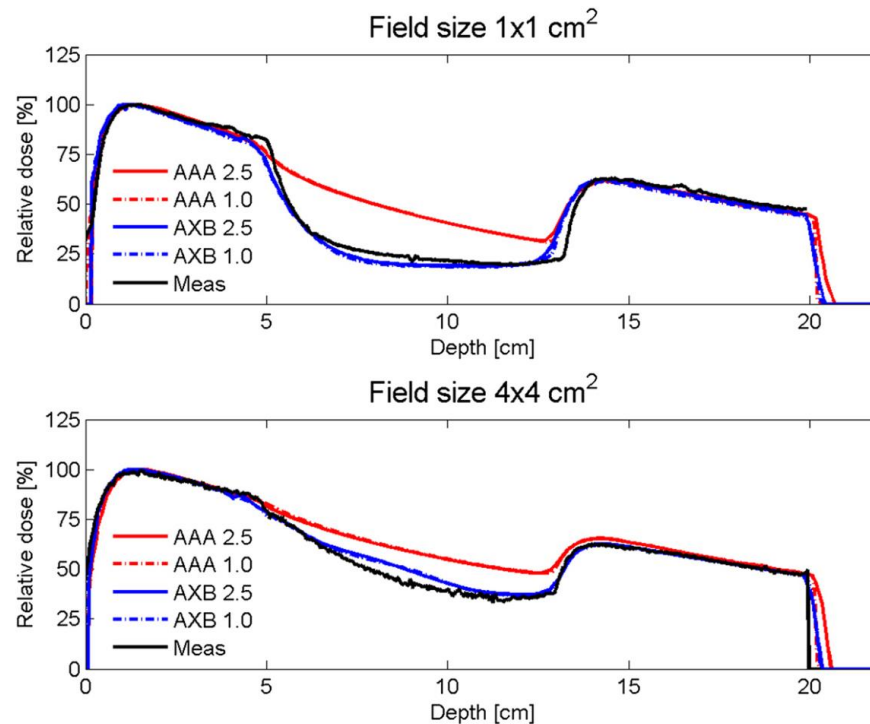
- Source size measurements with simple methods
- Measurement-less small-field output factor prediction
- Variations from accelerator to accelerator

Treatment Planning Algorithms – small fields

- Factor based
 - Successfully used in cranial SRS
- Model based
 - Beam model
 - coupled angular - energy distribution of a representative set of particles in the beam (photons and contamination particles)
 - Source parameters - TPS parameterizes the source size – impact on dose calculation accuracy
 - Collimation system - Backup collimation, alignment of different collimation systems
 - Patient model
 - Type a (or category 1)
 - equivalent path-length scaling for inhomogeneity corrections
 - Type b (or category 2)
 - changes in lateral electron transport are considered in some fashion
 - **Advanced type-b: MC or deterministic transport algorithms**

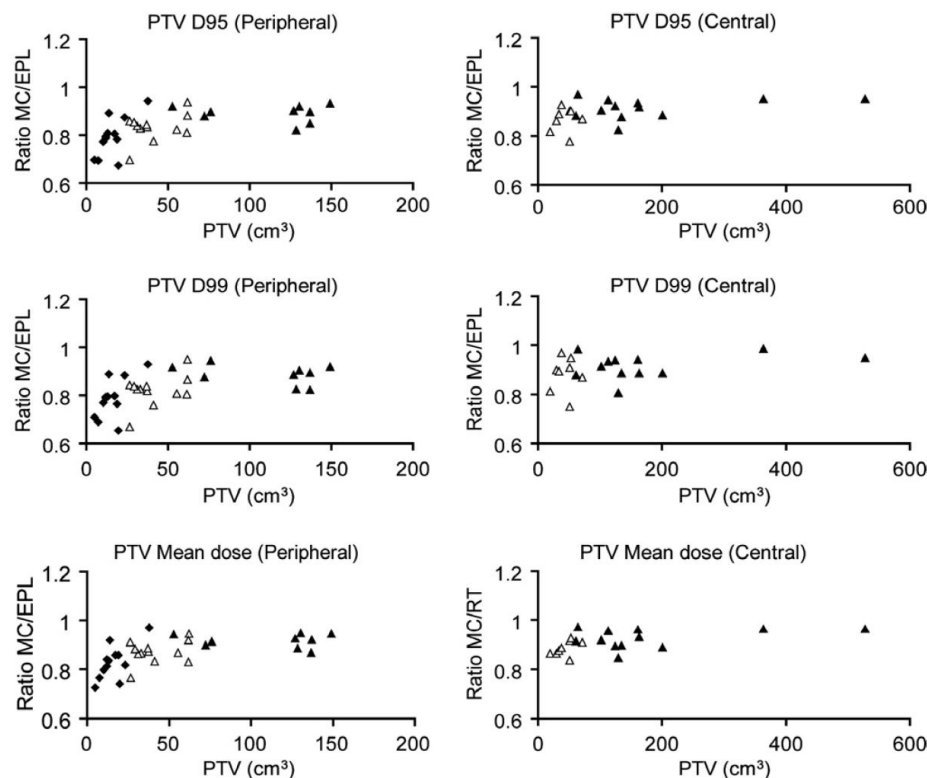


Monte Carlo-calculated central-axis depth-dose profiles for a lung slab phantom geometry irradiated by a 6 MV and a 18 MV beam (3 x 3 cm² field size) with a 1 × 1 × 1 cm³ tumour embedded in the lung, with decreasing lung slab density. [Disher, et al., 2012](#)

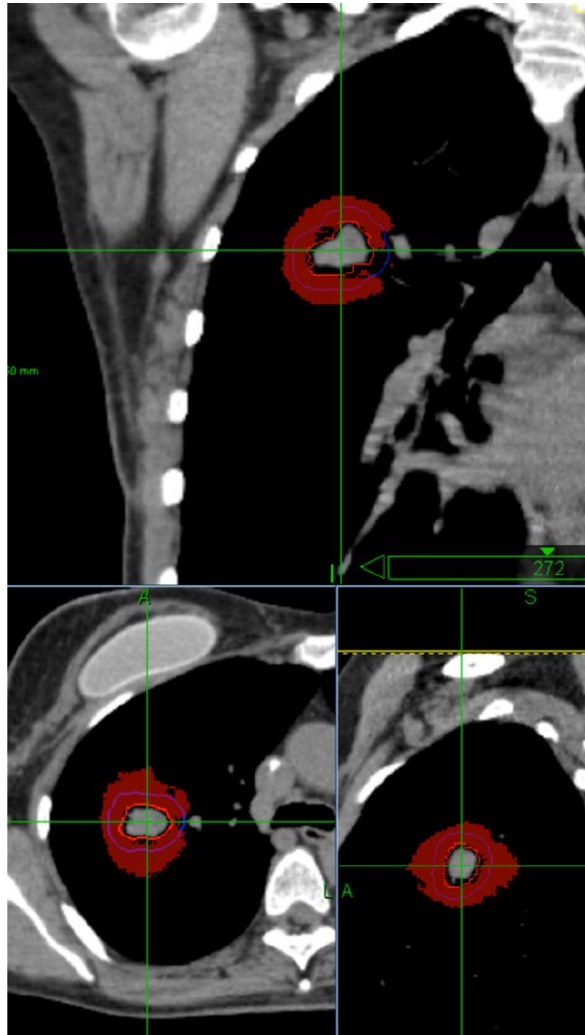


Comparison of category 2 algorithms AAA and Acuros XB (AXB, Varian) calculated with measured percentage depth doses for field sizes of 1 x 1 cm² and 4 x 4 cm². The phantom consists of foam, with a low-density $\rho = 0.03 \text{ g cm}^{-3}$ and a thickness of 8 cm sandwiched between two layers of polystyrene with a density of $\rho = 1.05 \text{ g cm}^{-3}$. [Kroon, et al., 2013](#)

Considerations for Clinical Prescription Using Category 2 Dose Calculation Algorithms in Small Fields



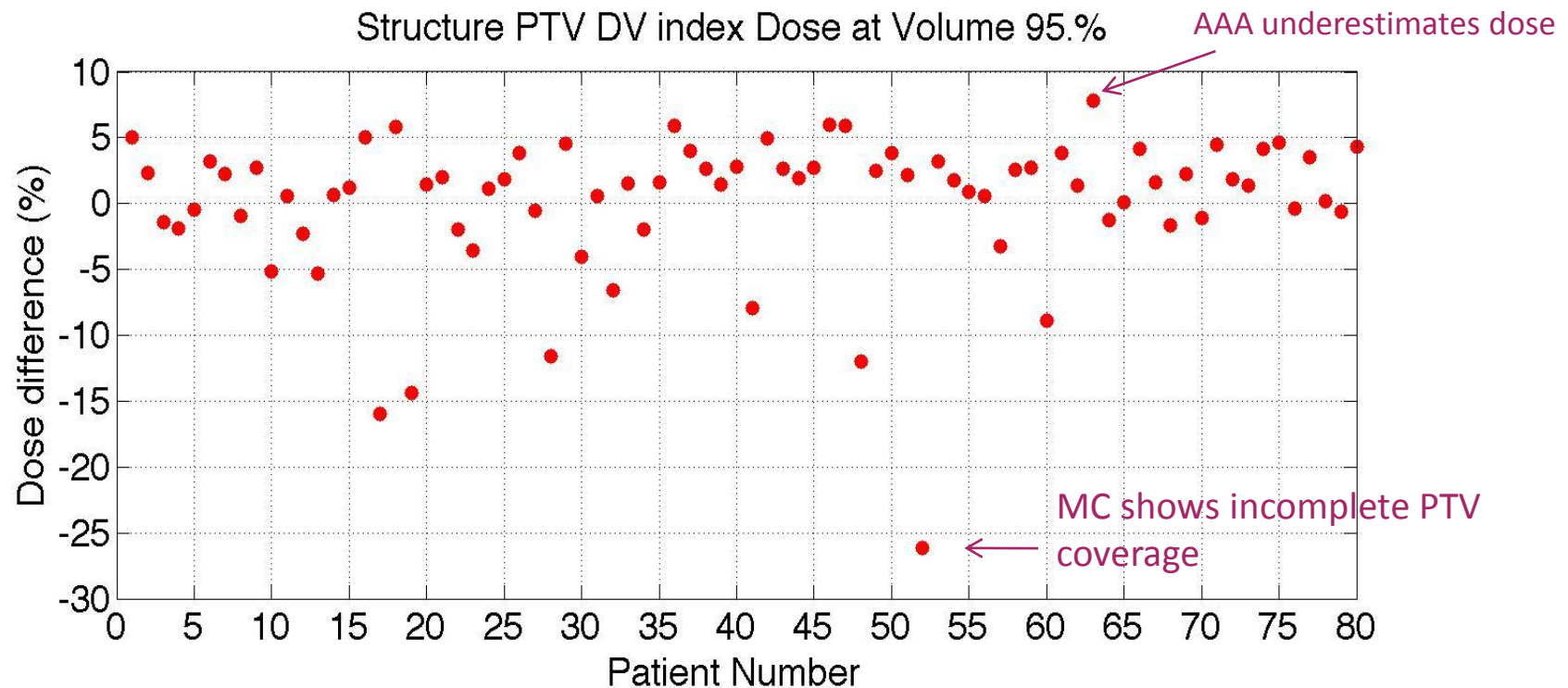
Ratio of MC and EPL calculated PTV $D_{95\%}$, $D_{99\%}$ and mean dose for peripheral and central pulmonary tumors. Bold diamonds represent tumors <3 cm, open triangles represent tumors of 3–5 cm and bold triangles represent tumors >5 cm. Data is for the CyberKnife 6 MV beam. [van der Voort van Zyp, et al., 2010](#)).



Region of dose difference exceeding 15 Gy outside the GTV, between equivalent path length correction (EPL) and Monte Carlo for CyberKnife (6 MV) treatments of a tumor with size 3.6 cm^3 . Dose prescribed 60 Gy. ([Lacornerie, et al., 2014](#))

--> **ICRU Report 91** mandates the use of advanced type b model-based dose calculation algorithms (Monte Carlo, etc)

Large scale lung SBRT dose calculations



- Positive results indicate the dose is underestimated by AAA
- Negative results indicate the PTV coverage is overestimated by AAA
- Range: +8% to -26%

E. Soisson et al 2012

Why do we care?

- 217 primary stage I non-small cell lung cancer (NSCLC) treated using SBRT between 2011 and 2015
- 37 pts developed distant metastases; median follow-up time 24 months
- 2 institutions

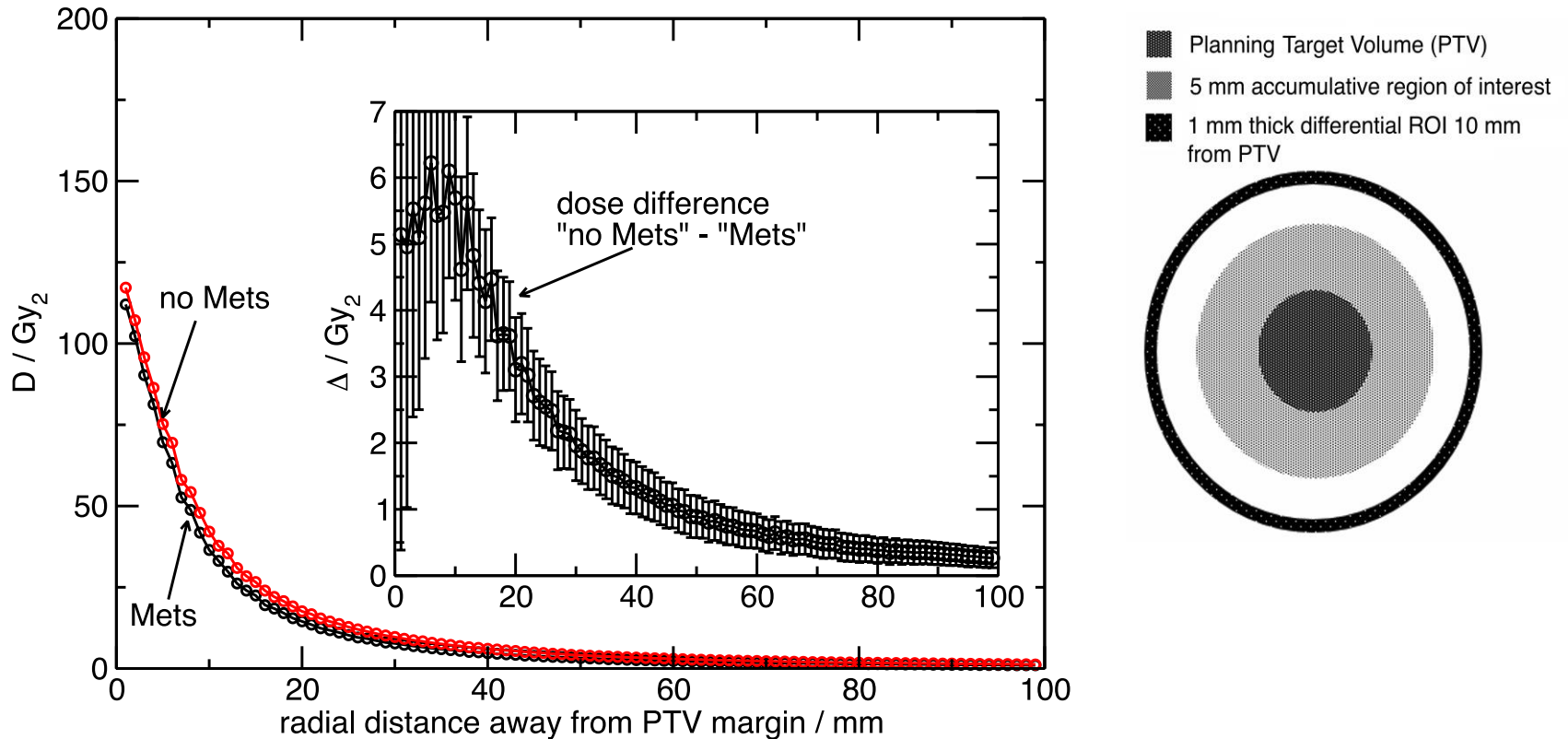
Radify (M.A. Renaud) →



AAA versus MC

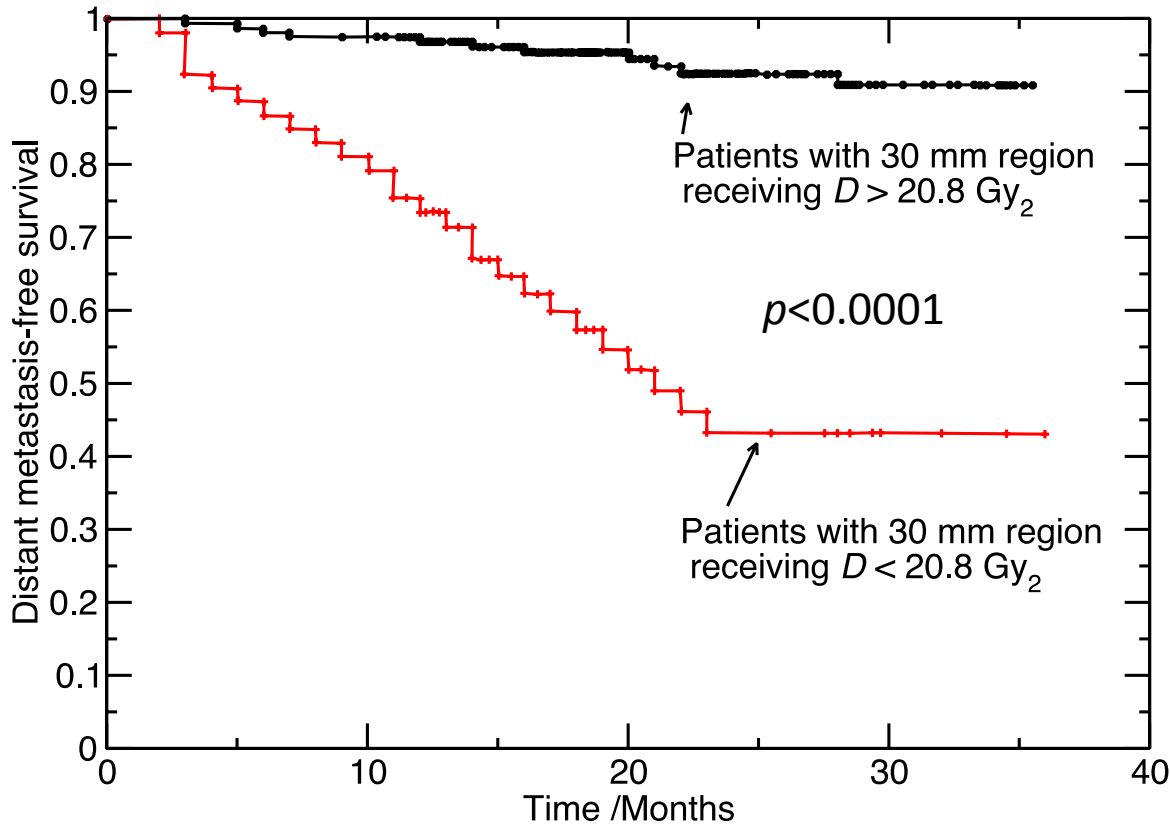
Poster session #56 (Boustead et al, 2017)

Dose difference → different outcome in terms of distant metastasis probability

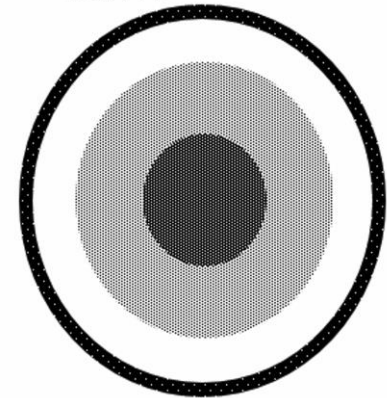


A. Boustead et al; preliminary

Same data: Distant metastasis-free survival



- Planning Target Volume (PTV)
- 5 mm accumulative region of interest
- 1 mm thick differential ROI 10 mm from PTV



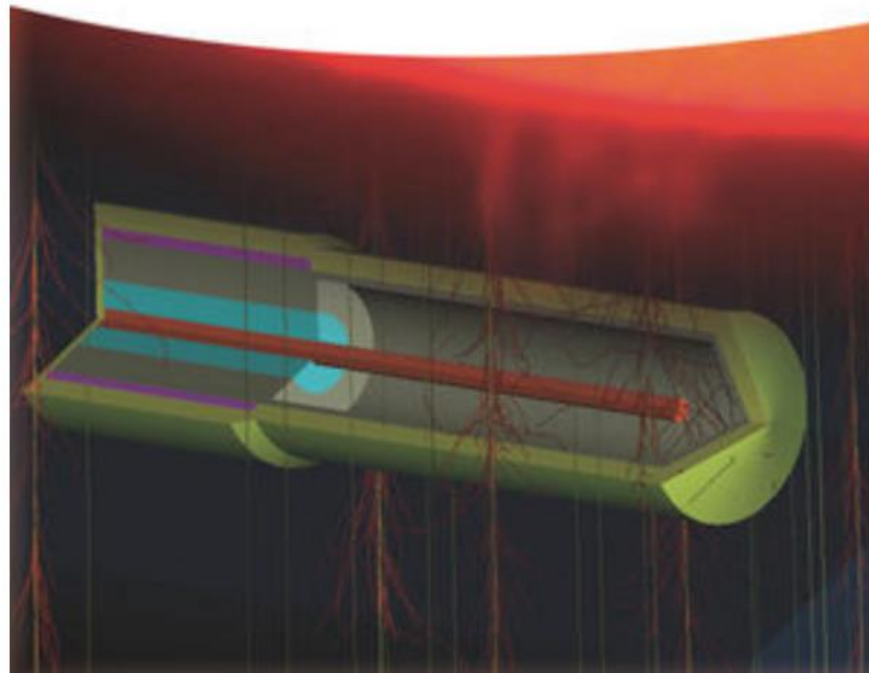
A. Boustead et al; preliminary

Conclusions

- Small photon beams are tricky
- Successful SRT hinges on accurate small field dosimetry
- In the past two decades our understanding and formalization of small field dosimetry has significantly improved
 - Calibration
 - Detectors and correction factors
 - Dose calculation algorithms
- Monte Carlo techniques have played and continue to play a core role in our understanding of radiation dosimetry of these fields

Pedro Andreo, David T. Burns, Alan E. Nahum,
Jan Seuntjens, and Frank H. Attix

Fundamentals of Ionizing Radiation Dosimetry



FIORD is here!
The new Attix book

Thank you!

