## Radiotherapy in Hungary: present status and future needs



#### **Tibor Major, PhD**

National Institute of Oncology Radiotherapy Department Budapest, Hungary





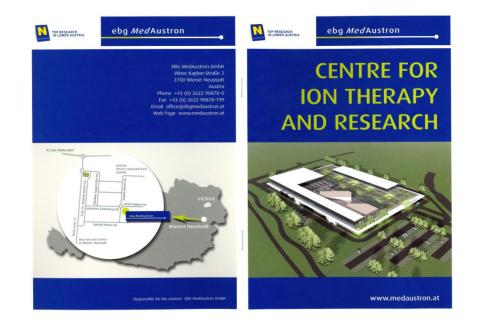
Academia Europaea Section Workshops, Bergen, 10 September, 2012

## Outline

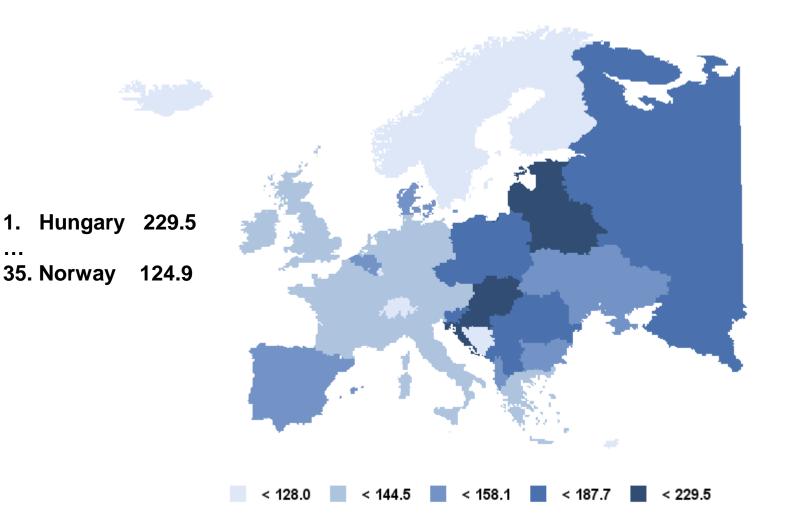
Cancer statistics and radiotherapy in Hungary

The role of particle therapy in cancer management

MedAustron – RegIonCo project

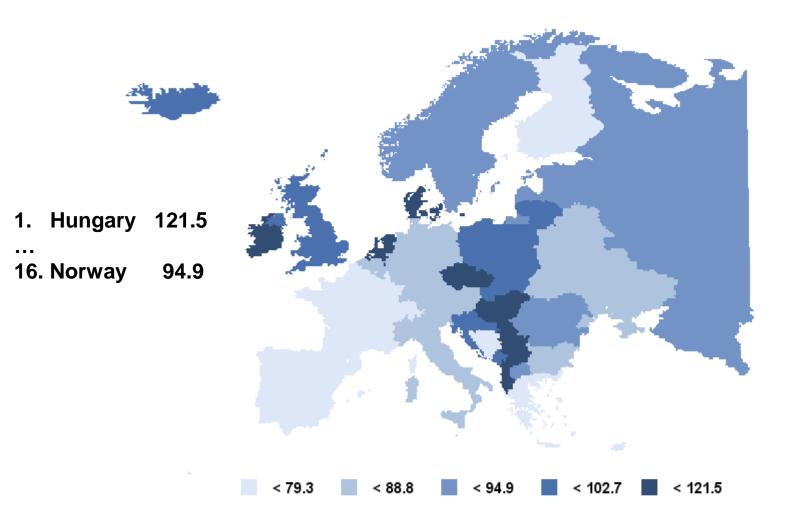


## Estimated age-standardised mortality rate for males per 100 000 (all cancers excl. non-melanoma skin cancer)



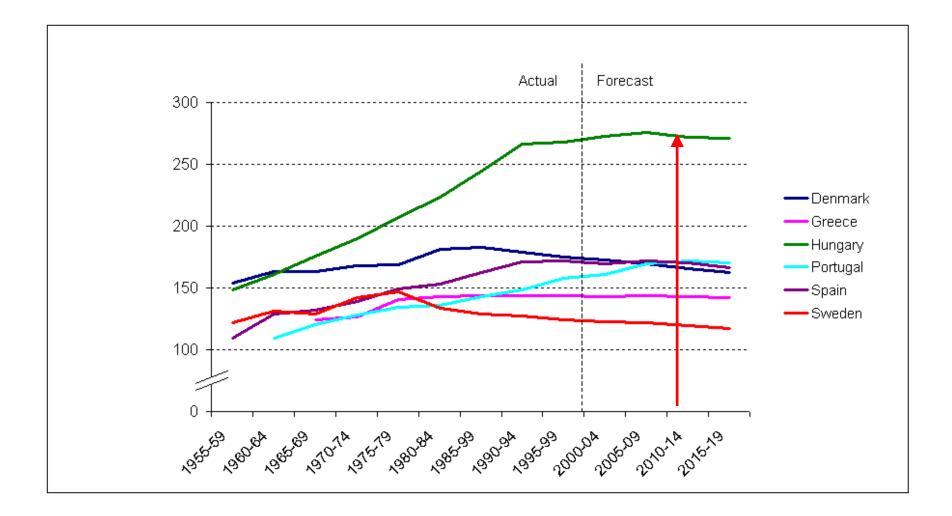
GLOBOCAN 2008 (IARC) - 2.9.2012

## Estimated age-standardised mortality rate for males per 100 000 (all cancers excl. non-melanoma skin cancer)

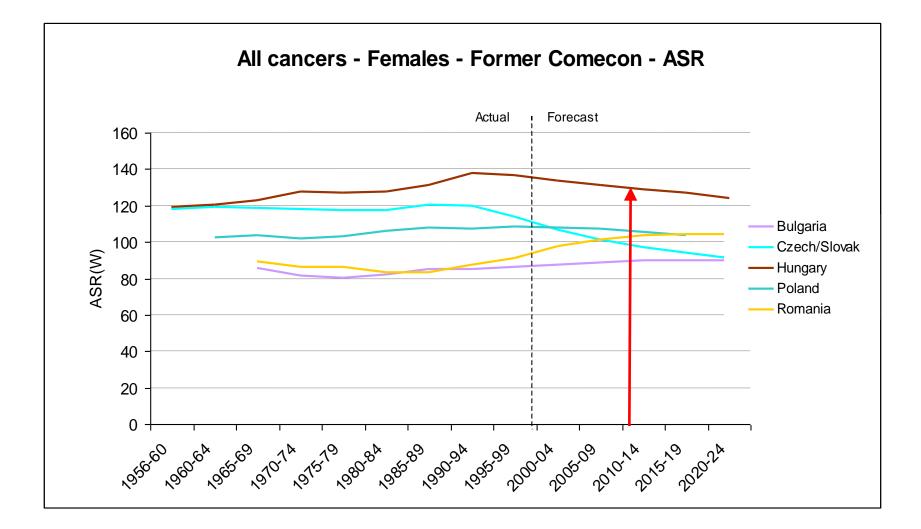


GLOBOCAN 2008 (IARC) - 2.9.2012

# Cancer mortality for male population 1955-2019



# Cancer mortality for female population 1955-2019

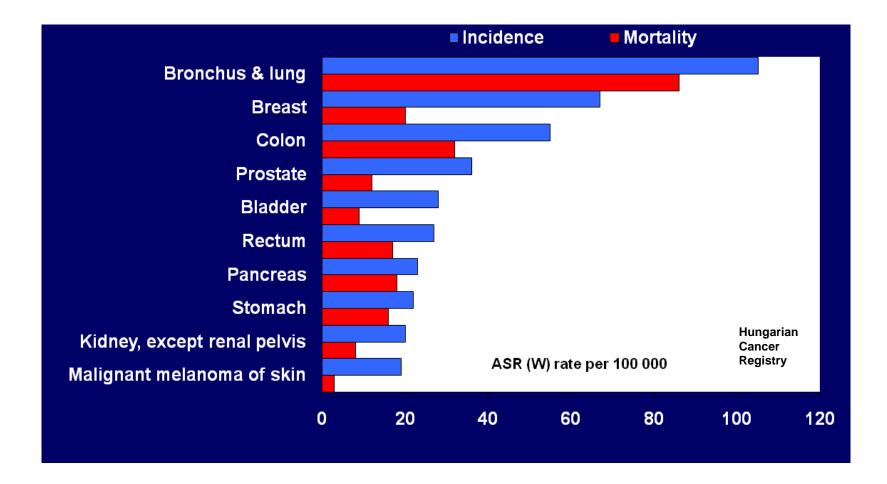


## **Cancer patient characteristics in Hungary in 2010**

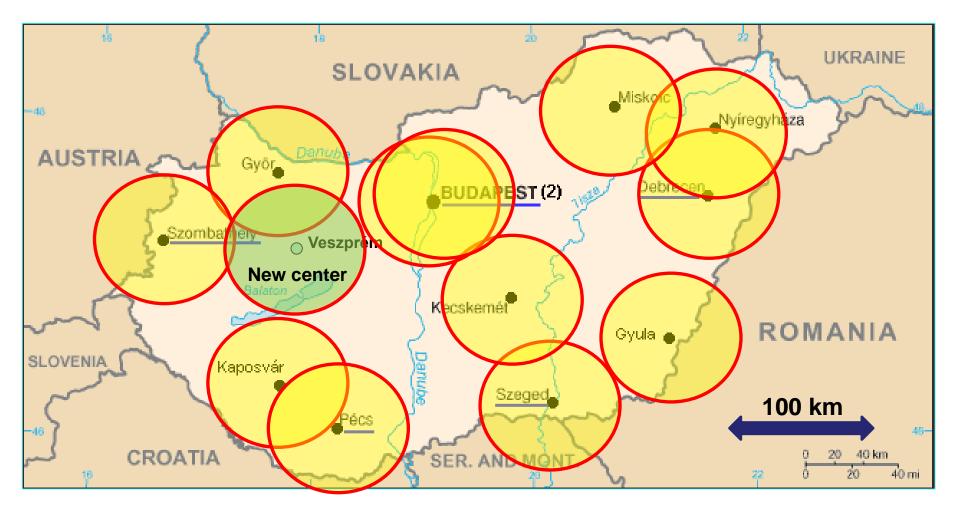
Diagnoses ICD-10	Incidence
1. Skin (non-melanoma) C44	10 957
2. Lung & trachea C33-34	10 706
3. Colo-rectal C18-21	9 632
4. Breast C50	7 543
5. Head & neck C11-14, C30-32	5 177
6. Lymphoma & leukemia C81-96	3 877
Altogether	69 262

Patients treated by radiotherapy: 23 119 (33.4 %)

## Top 10 cancers in Hungary (2010)



## Radiotherapy centers in Hungary (n=12)

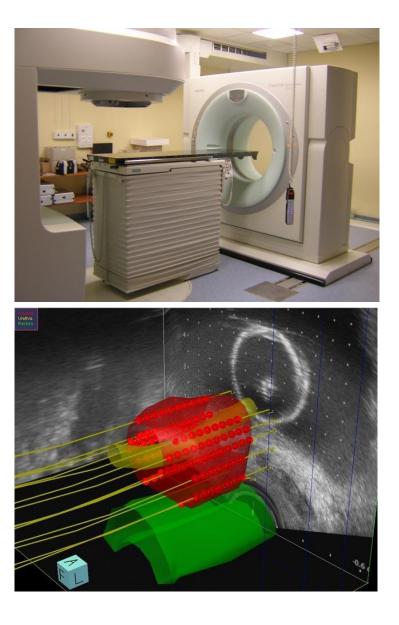


**Regional centers: 5** New center: 1

## **Radiotherapy infrastructure and staff in Hungary**

Linear accelerators (X, e)	26
Cobalt units (Co-60)	9
HDR afterloading (Ir-192)	11
Prostate seed BT (I-125)	1

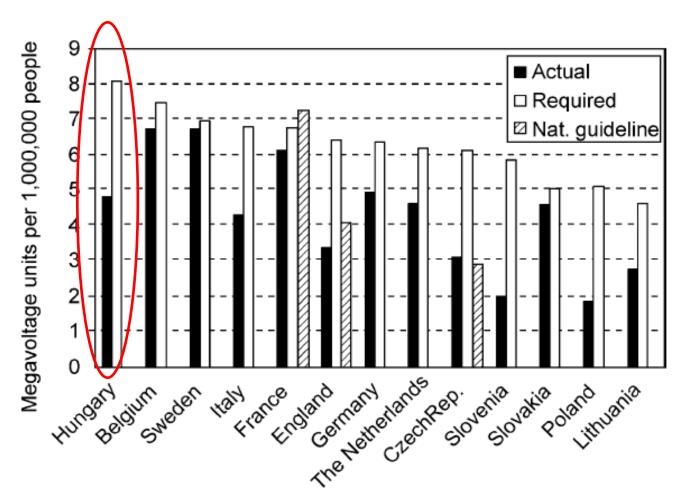
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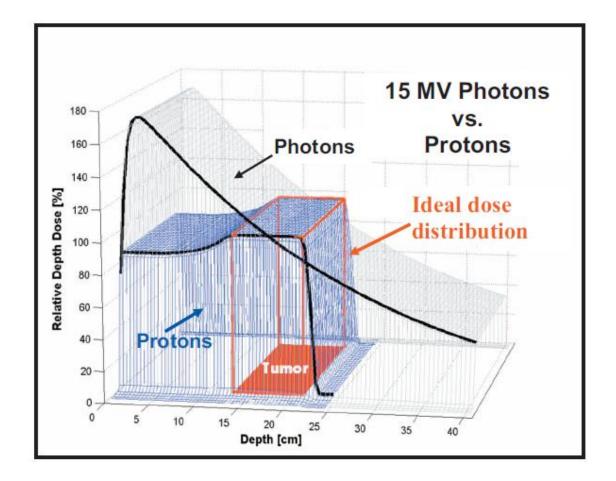
Towards evidence-based guidelines for radiotherapy infrastructure and staffing needs in Europe: the ESTRO QUARTS project

Søren M. Bentzen<sup>a,\*</sup>, Germaine Heeren<sup>b</sup>, Brian Cottier<sup>c</sup>, Ben Slotman<sup>d</sup>, Bengt Glimelius<sup>e, f</sup>, Yolande Lievens<sup>g</sup>, Walter van den Bogaert<sup>g</sup>

Radiother Oncol 75:355-365 (2005)

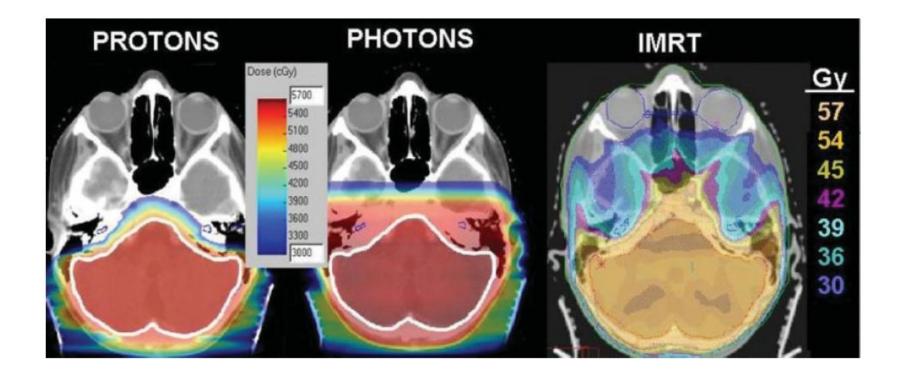


# Comparison between depth dose curves for photon and proton beams



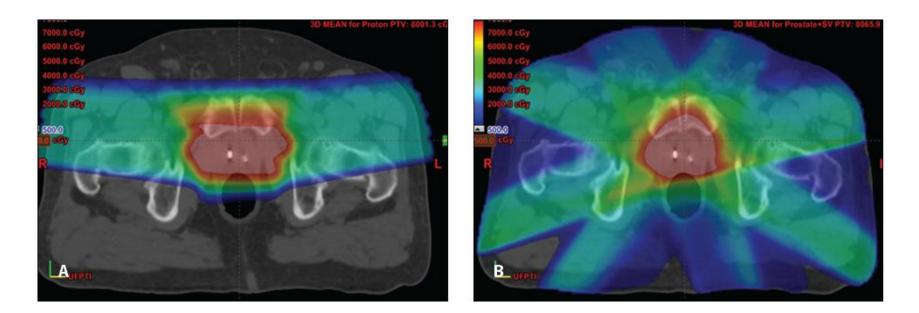
Smith AR, Med. Phys. 36:556-568. 2009.

## A comparison of dose distributions for a medulloblastoma patient using protons, photons and IMRT



Greco C, Wolden S, Cancer 109: 1227-1238. 2007.

### Irradiation of prostate with protons and photons



Proton

#### **Photon - IMRT**

Hoppe B, et al. Oncology, June, 644-662. 2011

How costly is particle therapy? Cost analysis of external beam radiotherapy with carbon-ions, protons and photons

Andrea Peeters<sup>a,e</sup>, Janneke P.C. Grutters<sup>a,d</sup>, Madelon Pijls-Johannesma<sup>a,b,\*</sup>, Stefan Reimoser<sup>c</sup>, Dirk De Ruysscher<sup>a,b</sup>, Johan L. Severens<sup>d,e</sup>, Manuela A. Joore<sup>d,e,1</sup>, Philippe Lambin<sup>a,b,1</sup>

<sup>a</sup> Maastricht Radiation Oncology (MAASTRO Clinic), The Netherlands; <sup>b</sup>GROW Research Institute, University Medical Center, Maastricht, The Netherlands; <sup>c</sup>Turner & Townsend, München, Germany; <sup>d</sup>Department of Health Organization, Policy & Economics, Maastricht University, The Netherlands; <sup>e</sup>Department of Clinical Epidemiology and Medical Technology Assessment, University Hospital Maastricht, The Netherlands

Radiother. Oncol. 95:45-53. 2010

#### Costs

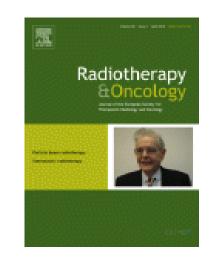
- capital
- operational

### **Three facilities**

- combined (carbon ion and proton)
- proton only
- photon

### Four tumor types

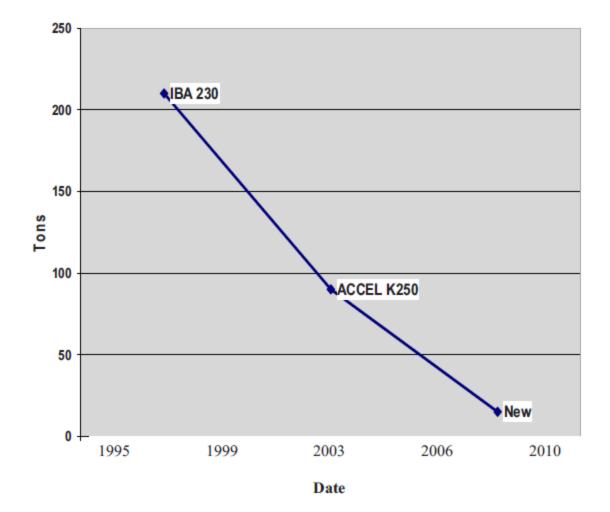
- prostate
- lung
- head & neck
- skull-base chordoma



## **Base-case analysis: operational model and cost estimates**

Operation Particle facility				
	Combined (carbon-ion and proton)	Proton- only	facility	
Number of rooms	3 rooms <sup>a</sup>	3 rooms <sup>b</sup>	2 rooms <sup>c</sup>	
Treatment category <sup>d</sup>	Special	Special	Standard	
Working days per week	5	5	5	
Hours/day available for treatment	14	14	14	
Days of operation/year	250	250	250	
Time/fraction (average)	18 min	18 min	10 min	
Number of fractions/room	11,667	11,667	21,000	
Treatment room utilization	98%	98%	100%	
Treatment room availability	95%	98%	98%	
Total number of fractions/year	32,585	33,614	41,160	
Number of fractions/patient (average)	18	20	18	
Total number of patients/year	1810	1681	2287	
Capital costs Medical equipment and IT Particle therapy equipment	13,750,000 90,000,000	<u>11,250,000</u> 60,000,000	14,250,000 <sup>e</sup> -	
Building	34,850,000	23,680,000	9,180,000	
Total capital costs	138,600,000	94,930,000	23,430,000	
Assumed lifecycle	30 years	30 years	30 years	
Capital costs/year	4,620,000	3,164,333	781,000	
Operational costs				
Cost of operation/year	10,952,350	5,736,450	2,758,350	
Cost of renewal/year	3,697,750	2,080,200	1,562,700	
Cost of staff/year	6,366,304	6,366,304	2,599,716	
Yearly interest for financing (5% over 20 years)	11,121,623	7,617,429	1,880,084	
Operational cost/year	32,138,027	21,800,383	8,800,850	
Total costs/year	36,758,027	24,964,716	9,581,850	
Cost/fraction	1128	743	233	
Ratio to photon	4.8	3.2	-	

### Weight (size) of clinical cyclotrons vs. date



Smith AR, Med. Phys. 36:556-568. 2009

## Proton therapy is too expensive for the minimal potential improvements in outcome claimed

Robert J. Schulz, Ph.D., FAAPM Yale University, Johnson, Vermont 05656 (Tel: 802-635-7351, E-mail: schulz@pshift.com)

Alfred R. Smith, Ph.D., FAAPM MD Anderson Proton Therapy Center, Houston, Texas 77054 (Tel: 713-516-1519, E-mail: alsmith@mdanderson.org)

Colin G. Orton, Ph.D., Moderator

#### POINT/COUNTERPOINT Med. Phys. 34:1135-38, April 2007

Proton therapy (PT) is "faith-based" radiotherapy without enough evidence

Not widely available for patients who may benefit from it

PT is 2-3 times more expensive than IMRT

Most of the cancer deaths are due to metastatic disease

Dose escalation will increase local control that may improve long-term DFS PT has less side effects and about 2 times less integral dose

Cost will be reduced as more and cost effective facilities are being designed

Cost calculation should include the cure of injuries obtained by the treatment

### Current status of radiotherapy with proton and light ion beams Greco C, Wolden S. Cancer 109:1227-1238. 2007

## Evidence based

- Ocular tumours (uveal melanoma > 4 mm)
- Base of skull tumours (chordomas)

## • Some evidence

- Pediatric malignancies (e.g. retinoblastoma)
- Paraspinal tumours
- Head & neck ACC (for C-ions)
- Experimental
  - Lung tumours
  - Breast (APBI)
  - Pelvic tumours (prostate, uterine cervix, bladder, GI)
  - Sarcomas

## Particle therapy facilities in operation I.

WHO, WHERE	COUNTRY	PARTICLE	S/C*, MAX. ENERGY (MeV)	BEAM DIRECTION	START OF TREATMENT
ITEP, Moscow	Russia	р	S 250	1 horiz.	1969
St.Petersburg	Russia	р	S 1000	1 horiz.	1975
PSI, Villigen	Switzerland	p**	C 250	1 gantry, 1 horiz.	1996
Dubna	Russia	р	C 200****	horiz.	1999
Uppsala	Sweden	р	C 200	1 horiz.	1989
Clatterbridge	England	р	C 62	1 horiz.	1989
Loma Linda	CA.,USA	р	S 250	3 gantry, 1 horiz.	1990
Nice	France	р	C 65	1 horiz.	1991
Orsay	France	p*****	C 230	1 gantry,2 horiz.	1991
NRF - iThemba Labs	South Africa	р	C 200	1 horiz.	1993
IU Health PTC, Bloomington	IN.,USA	р	C 200	2 gantry, 1 horiz.	2004
UCSF	CA.,USA	р	C 60	1 horiz.	1994
HIMAC, Chiba	Japan	C-ion	S 800/u	horiz.,vertical	1994
TRIUMF, Vancouver	Canada	р	C 72	1 horiz.	1995
HZB (HMI), Berlin	Germany	р	C 72	1 horiz.	1998
NCC, Kashiwa	Japan	р	C 235	2 gantry	1998
HIBMC,Hyogo	Japan	р	S 230	1 gantry	2001
HIBMC,Hyogo	Japan	C-ion	S 320/u	horiz.,vertical	2002
PMRC(2), Tsukuba	Japan	р	S 250	gantry	2001
NPTC, MGH Boston	MA.,USA	p***	C 235	2 gantry, 1 horiz.	2001
INFN-LNS, Catania	Italy	р	C 60	1 horiz.	2002
Shizuoka Cancer Center	Japan	р	S 235	3 gantry, 1 horiz.	2003
STPTC, Koriyama- City	Japan	р	S 235	2 gantry, 1 horiz.	2008

## Particle therapy facilities in operation II.

WHO, WHERE	COUNTRY	PARTICLE	S/C*, MAX. ENERGY (MeV)	BEAM DIRECTION	START OF TREATMENT
WPTC, Zibo	China	р	C 230	2 gantry, 1 horiz.	2004
MD Anderson Cancer Center, Houston	TX.,USA	p***	S 250	3 gantry, 1 horiz.	2006
UFPTI, Jacksonville	FL.,USA	р	C 230	3 gantry, 1 horiz.	2006
NCC, Ilsan	South Korea	р	C 230	2 gantry, 1 horiz.	2007
RPTC, Munich	Germany	p**	C 250	4 gantry, 1 horiz.	2009
ProCure PTC, Oklahoma City	OK.,USA	р	C 230	1 gantry, 1 horiz, 2 horiz/60 deg.	2009
HIT, Heidelberg	Germany	p**	S 250	2 horiz.	2009
HIT, Heidelberg	Germany	C-ion**	S 430/u	2 horiz.	2009
UPenn, Philadelphia	PA.,USA	р	C 230	4 gantry, 1 horiz.	2010
GHMC, Gunma	Japan	C-ion	S 400/u	3 horiz., vertical	2010
IMP-CAS, Lanzhou	China	C-ion	S 400/u	1 horiz.	2006
CDH Proton Center, Warrenville	IL.,USA	р	C 230	1 gantry, 1 horiz, 2 horiz/60 deg.	2010
HUPTI, Hampton	VA., USA	р	C 230	4 gantry, 1 horiz.	2010
IFJ PAN, Krakow	Poland	р	C 60	1 horiz.	2011
Medipolis Medical Research Institute, Ibusuki	Japan	р	S 250	3 gantry	2011
CNAO, Pavia	Italy	C-lon, p	S 430/u	3 horiz/1 vertical	2011
ProCure Proton Therapy Center, New Jersey	NY., USA	р	C 230	4 gantry	2012

## Planned particle therapy facilities or under construction I.

WHO, WHERE	COUNTRY	PARTICLE	MAX. CLINICAL ENERGY (MeV)		NO. OF TREATMENT ROOMS	START OF TREATMENT PLANNED
PTC Czech s.r.o., Prague*	Czech Rep.	р	230 cyclotron	3 gantries, 1 horiz fixed beam	4	2012
Med-AUSTRON, Wiener Neustadt*	Austria	p, C-ion	430/u synchrotron	1 gantry (only for protons) 1 fixed beam, 1 fixed 0 + 90 deg	3	2015
ATreP, Trento *	Italy	р	230 cyclotron	2 gantries 1 horiz fixed beam	3	2013
Fudan University Shanghai CC*	China	p, C-ion	430/u synchrotron	3 fixed beams	3	2014
McLaren PTC, Flint, Michigan*	USA	р	250/330 synchrotron	3 gantries	3	2012
WPE, Essen*	Germany	р	230 cyclotron	3 gantries, 1 horiz fixed beam	4	2012
HITFil, Lanzhou*	China	C-ion	400/u synchrotron	4 horiz, vertical, oblique, fixed beams	4	2013
PTC, Marburg*	Germany	p, C-ion	430/u synchrotron	3 horiz fixed beams 1 fixed beam 0 + 45 deg	4	2012?
Northern Illinois PT Res.Institute, W. Chicago, IL*	USA	р	250 SC cyclotron	2 gantries, 2 horiz fixed beams	4	2012?
Chang Gung Memorial Hospital, Taipei*	Taiwan	р	235 cyclotron	4 gantries, 1 experimental room	4	2012
PMHPTC, Protvino*	Russia	р	250 synchrotron	1 horiz fixed beam	1	2012?
CCSR, Bratislava	Slovak Rep.	р	72 cyclotron	1 horiz fixed beam	1	?
CMHPTC, Ruzomberok*	Slovak Rep.	р	250 synchrotron	1 horiz fixed beam	1	?

## Planned particle therapy facilities or under construction II.

WHO, WHERE	COUNTRY	PARTICLE	MAX. CLINICAL ENERGY (MeV)	BEAM DIRECTION	NO. OF TREATMENT ROOMS	START OF TREATMENT PLANNED
SJFH, Beijing	China	р	230 cyclotron	1 gantry, 1 horiz fixed beam	2	?
Skandion Clinic, Uppsala*	Sweden	р	230 cyclotron	2 gantries	2	2013
Barnes Jewish St. Louis, MO*	USA	р	250 SC synchro- cyclotron	1 gantry	1	2012
Scripps Proton Therapy Center, San Diego, CA*	USA	р	250 SC cyclotron	3 gantries, 2 horiz fixed beams	5	2013
SCCA Proton Therapy, a ProCure Center, Seattle, WA*	USA	р	230 cyclotron	4 gantries	4	2013
Samsung Proton Center, Seoul*	South Korea	р	230 cyclotron	2 gantries	2	2014
Robert Wood Johnson, New Brunswick*	USA	р	250 SC synchro- cyclotron	1 gantry	1	2013
Oklahoma University, Oklahoma City, OK*	USA	р	250 SC synchro- cyclotron	1 gantry	1	2013
MD Anderson, Orlando, FL*	USA	р	250 SC synchro- cyclotron	1 gantry	1	2013
First Coast Oncology, Jacksonville, FLI*	USA	р	250 SC synchro- cyclotron	1 gantry	1	2013
IFJ PAN, Krakow*	Poland	р	235 cyclotron	1 gantry	1	2014?
PTC Zürichobersee, Galgenen	Switzerland	р	230 cyclotron	4 gantries, 1 horiz fixed beam	5	2016

## Hadron therapy patient statistics

Patient Statistics (for the facilities out of operation; end of 2010):

	WHERE	PARTICLE	FIRST	LAST	PATIENT	
			PAT	IENT	TOTAL	
Belgium	Louvain-la-Neuve	р	1991	1993	21	ocular tumors only
Canada	Vancouver (TRIUMF)	π_	1979	1994	367	ocular tumors only
Germany	Darmstadt (GSI)	C-ion	1997	2009	440	
Japan	Tsukuba (PMRC, 1)	р	1983	2000	700	
Japan	Chiba	p	1979	2002	145	ocular tumors only
Japan	WERC	р	2002	2009	62	
Russia	Dubna (1)	р	1967	1996	124	
Sweden	Uppsala (1)	p	1957	1976	73	
Switzerland	Villigen PSI (SIN-Piotron)	π_	1980	1993	503	
Switzerland	Villigen PSI (OPTIS 1)	р	1984	2010	5458	ocular tumors only
CA., USA	Berkeley 184	р	1954	1957	30	
CA., USA	Berkeley	He	1957	1992	2054	
CA., USA	Berkeley	ion	1975	1992	433	
IN., USA	Bloomington (MPRI, 1)	р	1993	1999	34	ocular tumors only
MA., USA	Harvard	p	1961	2002	9116	
NM., USA	Los Alamos	π_	1974	1982	230	
1 -		1		1	40700	T I

19790 Total

thereof

PTCOG: Particle Therapy Co-Operative Group

2054 He 1100 pions 440 C-ions 433 other ions

15763 protons

## Hadron therapy patient statistics

Patient Statistics (for the facilities in operation end of 2011):

	WHERE	PARTICLE	FIRST	PATIENT	DATE OF	
			PATIENT	TOTAL	TOTAL	
Canada	Vancouver (TRIUMF)	p	1995	161	Dec-11	ocular tumors only
China	Wanjie (WPTC)	p	2004	1078	Dec-11	no patients in 2011
China	Lanzhou	C ion	2006	159	Dec-11	
England	Clatterbridge	p	1989	2151	Dec-11	ocular tumors only
France	Nice (CAL)	p	1991	4417	Dec-11	ocular tumors
France	Orsay (CPO)	p	1991	5634	Dec-11	4540 ocular tumors
Germany	Berlin (HMI)	p	1998	1859	Dec-11	ocular tumors only
Germany	Munich (RPTC)	p	2009	895	Dec-11	
Germany	HIT, Heidelberg	C ion	2010	568	Dec-11	
Germany	HIT, Heidelberg	р	2010	94	Dec-11	
Italy	Catania (INFN-LNS)	p	2002	290	Dec-11	ocular tumors only
Italy	Pavia (CNAO)	C ion	2011	5	Dec-11	
Japan	Chiba (HIMAC)	C ion	1994	6569	Dec-11	11 with scanning
Japan	Kashiwa (NCC)	p	1998	870	Dec-11	estimated
Japan	Hyogo (HIBMC)	p	2001	3198	Dec-11	
Japan	Hyogo (HIBMC)	C ion	2002	1271	Dec-11	
Japan	Tsukuba (PMRC, 2)	p	2001	2166	Dec-11	
Japan	Shizuoka	p	2003	1175	Dec-11	
Japan	Koriyama-City	p	2008	1378	Dec-11	
Japan	Gunma	C ion	2010	271	Dec-11	
Japan	Ibusuki (MMRI)	p	2011	180	Dec-11	

March 2012 - Martin Jermann, PTCOG Secretary

to be continued

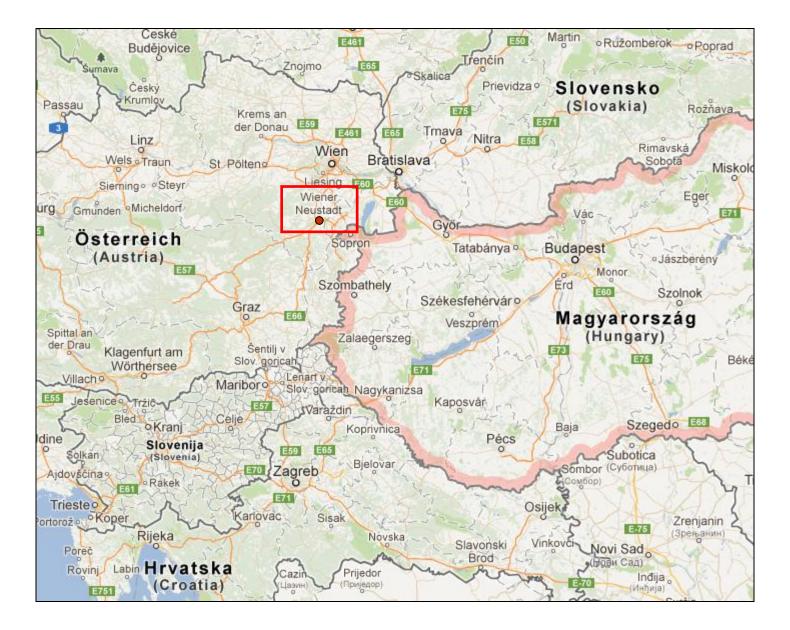
Korea	llsan, Seoul	p	2007	810	Dec-11	1
Poland	Krakow	p	2011	11	Dec-11	ocular tumors only
Russia	Moscow (ITEP)	p	1969	4300	Dec-11	estimated
Russia	St. Petersburg	p	1975	1372	Dec-11	
Russia	Dubna (JINR, 2)	p	1999	828	Dec-11	
South Africa	iThemba LABS	p	1993	521	Dec-11	
Sweden	Uppsala (2)	p	1989	1185	Dec-11	
Switzerland	Villigen PSI, incl OPTIS2	p	1996	1107	Dec-11	277 ocular tumors
USA, CA.	UCSF - CNL	p	1994	1391	Dec-11	ocular tumors only
USA, CA.	Loma Linda (LLUMC)	p	1990	16000	Dec-11	estimated
USA, IN.	Bloomington (IU Health PTC)	p	2004	1431	Dec-11	
USA, MA.	Boston (NPTC)	p	2001	5562	Oct-11	
USA, TX.	Houston (MD Anderson)	p	2006	3400	Feb-12	
USA, FL	Jacksonville (UFPTI)	p	2006	3461	Dec-11	
USA, OK.	Oklahoma City (ProCure PTC)	p	2009	623	Dec-11	
USA, PA.	Philadelphia Upenn)	p	2010	433	Dec-11	
USA, IL.	CDH Warrenville	p	2010	367	Dec-11	
USA, VA.	Hampton (HUPTI)	p	2010			no data available
				77191	Total	
			thereof	8843	C-ions	
					protons	
Total for all f	facilities (in operation and out of o	peration).		2054	He	
i otar i or an i		peration		1100		
					C-ions	
					other ions	
					protons	
					Grand Tot	al
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March 2012 - Martin Jermann, PTCOG Secretary

## How many patients eligible for proton therapy?

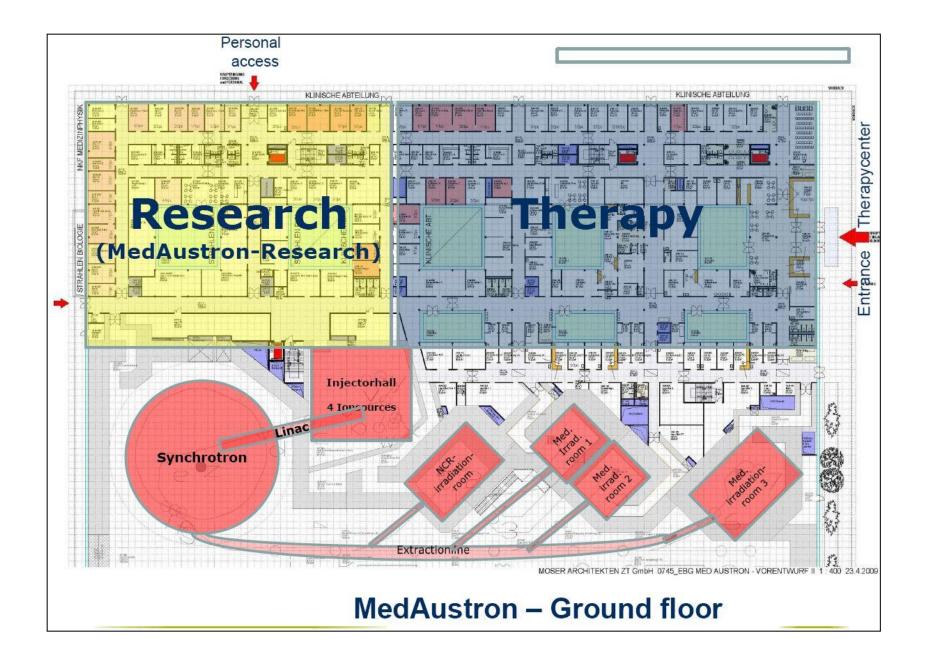
Country	No. of new cases	Pts. treated with RT	Suitable for proton	Percentage
Sweden* (9.3 millions)	31 050	16 000	2 350	15
Hungary** (10 millions)	58 305	23 119	2 865	13

\*Glimelius B, et al. Acta Oncol. 44:836-849. 2005 \*\*Mock U, ReglonCo project, 2011





Cost:	170 million EUR
Funds:	Federal State of Lower Austria
	Republic of Austria
	City of Wiener Neustadt



## **MedAustron – Wiener Neustadt**





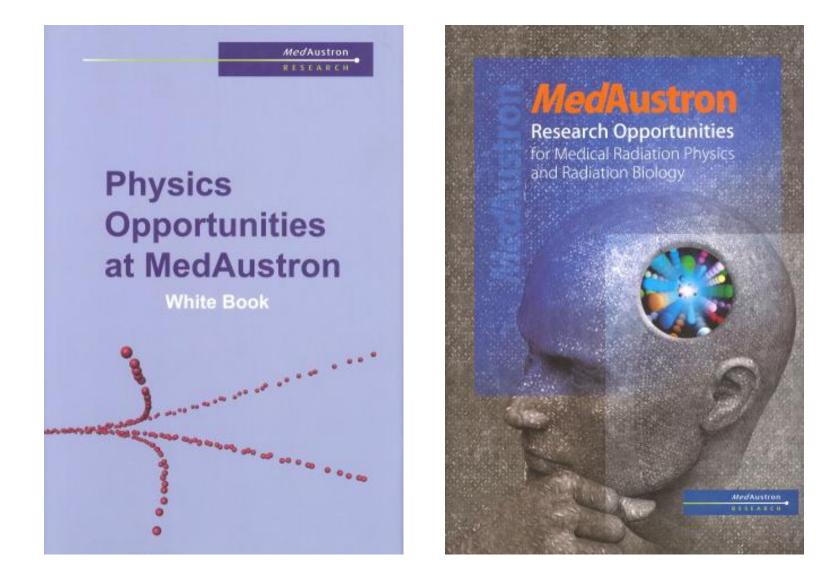
#### Timeline

Treatment of the first patient	2015
2014	Trial operation of the complete plant
Technical trial operation, installation of the medical equipment	2013
2012	Installation of the particle accellerator
Construction of the centre	2011

#### **Particle energy**

max. 800 MeV p for experiments max. 250 MeV p for treatments max. 400 MeV/A C for treatments

## **MedAustron books**



## **Regional Iontherapy Co-operation (RegIonCo)**



#### Projektpartnereink:

EBG MedAustron GmbH

PEG MedAustron GmbH

FOTEC Forschungsund Technologietransfer GmbH

Regionaler Entwicklungsverband Industrieviertel-Projektmanagement

Austrian Institute of Technology

#### EBG MedAustron Hungary Kft.

Vas Megyel Markusovszky Kórház Nonprofit Zrt.

Soproni Erzsébet Oktató Kórház

Országos Onkológiai Intézet Országos Korányi TBC és Pulmonológiai Intézet

# ReglonCo

www.regionco.eu

#### Regionális lonterápiás Kooperáció

Regionale Zusammenarbeit für Ionentherapie

#### Kapcsolat:

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Programm zur grenzüberschreitenden Kooperation ÖSTERREICH - UNGARN 2007-2013 AUSZTRIA - MAGYARORSZÁG Határon Átnyúló Együttműködési Program 2007-2013



## **RegionCo Project Partners**

Austria	Hungary
EBG MedAustron GmbH*	EBG MedAustron Hungary Kft.
FOTEC Forschungs- und Technologietransfer GmbH	Markusovszky Hospital Szombathely
PEG MedAustron GmbH	National Institute of Oncology Budapest
Regional Entwicklungsverband Industrieviertel-Projektmana- gement	National Korányi Institute Budapest
Austrian Institute of Technology	Erzsébet Hospital Sopron

\*Lead partner



## **RegionCo Work Packages**

- WP1 Project management
- WP 2.1 Principles of Treatment Planning with Helium ions
- WP 2.2 Molecular imaging
- WP 3.1 Transnational image management and IT networks
- WP 3.2 Joint utilization of MedAustron Research
- WP 4.1 Epidemiological survey in Hungary
- WP 4.2 Logistics of patient's management
- WP 5 Regional development
- WP 6 Communication and dissemination



## WP 2.1 Principles of Treatment Planning with Helium ions

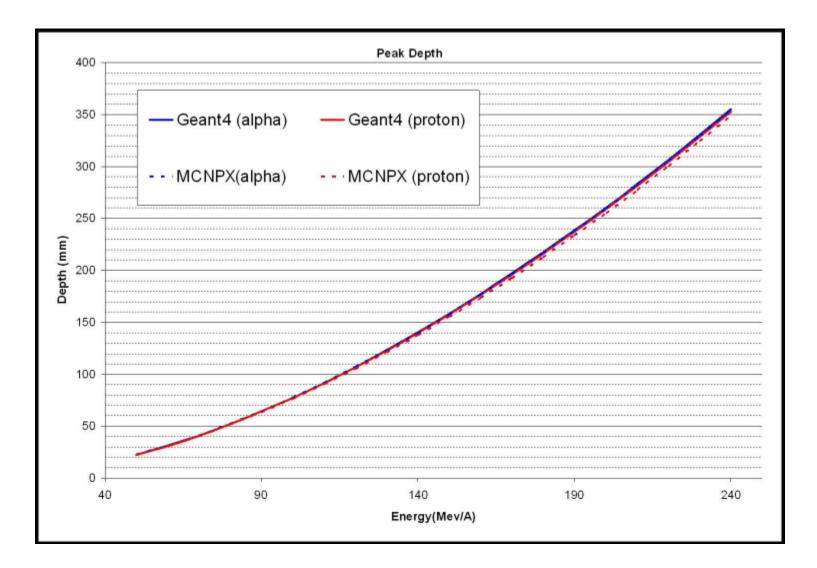
#### Monte Carlo simulations of proton and helium ion beams using

- GEANT4 (Geometry and Tracking, CERN)
- MCNPX (Monte Carlo N-Particle Transport Code, Los Alamos)

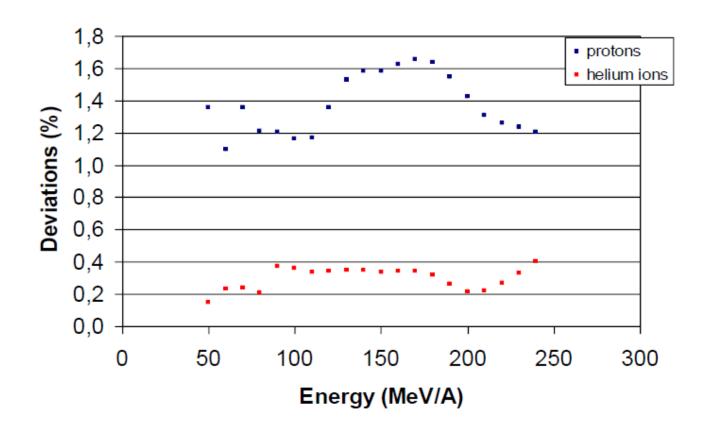
#### Two groups

- Wien EBG MedAustron GmbH (GEANT4)
- Budapest National Institute of Oncology (MCNPX)

## Bragg peak position calculated with MCNPX and GEANT4

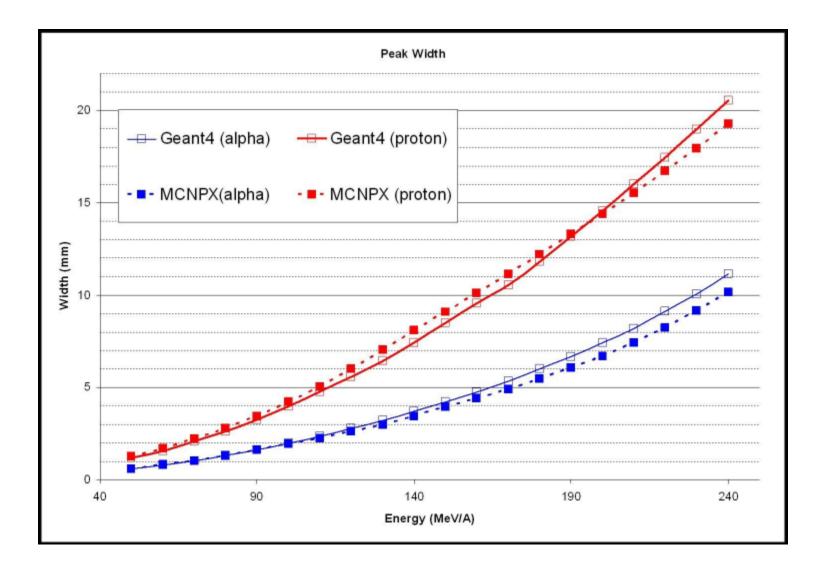


## Deviations of Bragg peak position calculated with MCNPX and GEANT4 for all initial kinetic energies

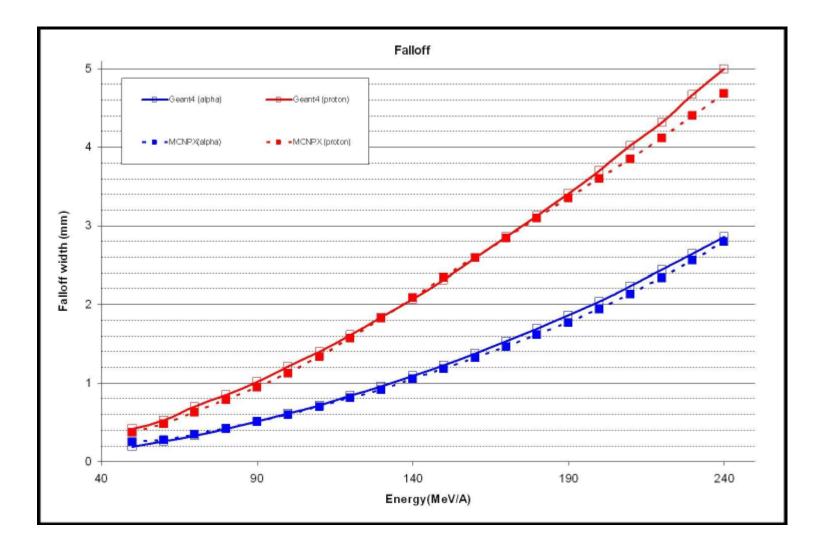


Peak Depth

## Bragg peak width calculated with MCNPX and GEANT4



## **Distal dose fall-off width calculated with MCNPX and GEANT4**





## Thank you for your attention!

