



## MedAustron Austria's Ion Therapy Center

Eugen Hug,  
Ulrike Mock, Johanna Salinger

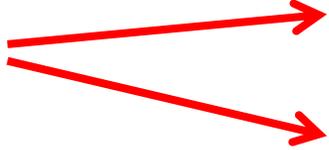
MedAustron  
Ion Therapy Center

Wiener Neustadt, Austria

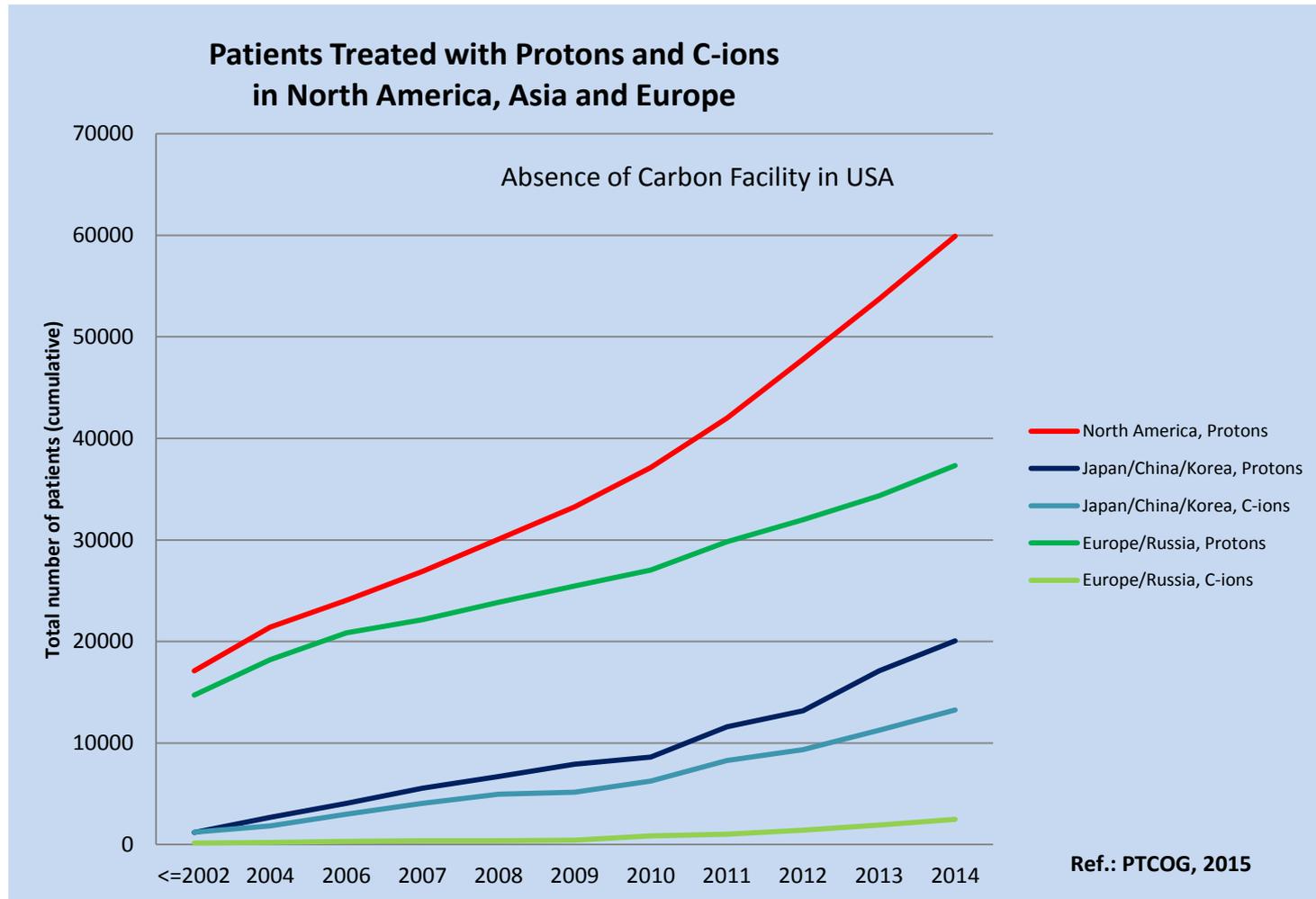
## Particle Therapy Worldwide – 2014 Survey by PTCOG Presented at PTCOG-54, San Diego, 2015 – E. Hug, M. Jermann

Number of Patients treated with Particle Therapy\* worldwide as of End 2014:

49 Centers operational

<b>P : C = 9:1</b> 	<b>Total of all facilities (in and out of operation):</b>	<b>He</b>	<b>2054</b>	1957-1992
		<b>Pions</b>	<b>1100</b>	1974-1994
		<b>C-ions</b>	<b>15736</b>	1994 -
		<b>Other ions</b>	<b>433</b>	1975-1992
		<b>Protons</b>	<b>118181</b>	1954 -
		<b>Grand Total</b>	<b>133931</b>	

\* Data collection = voluntary reporting by particle centers



## Particle Therapy –

### The Quest for the Clinically Meaningful Benefit

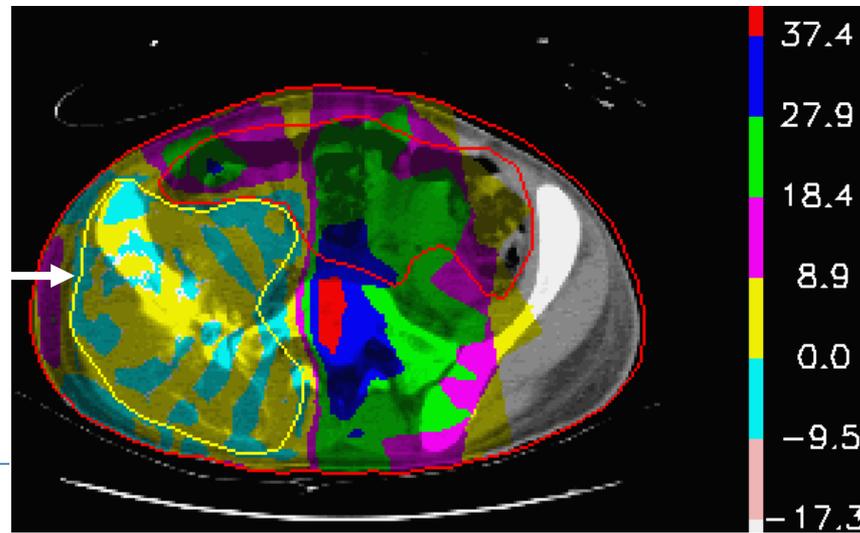
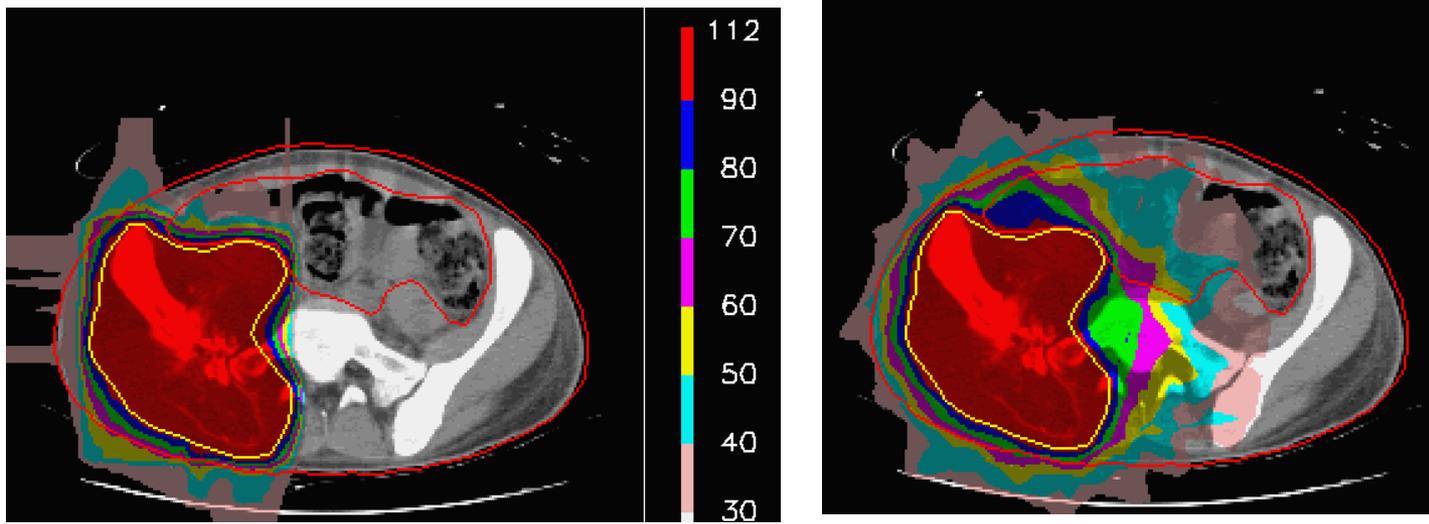


Dose to tumor



Dose to healthy tissue

# IMPT vs. IMXT : Ewing's Sarcoma



**Difference  
(IMXT – IMPT)**

# IMPT vs. IMXT : Ewing's Sarcoma

PROTONS

112

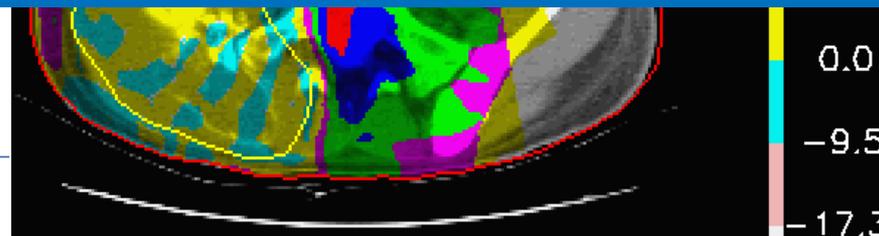
90

X-RAYS

Proton and Carbon ions will reduce the amount of integral dose delivered to normal tissues – the question is “clinical relevance”

- In children
- In young adults
- Adults
- Aging adults

Difference  
(IMXT – IMPT)



# Proton Indications: Large Target, High Complexity, Combined Modality Therapy

## Ewing's Sarcoma

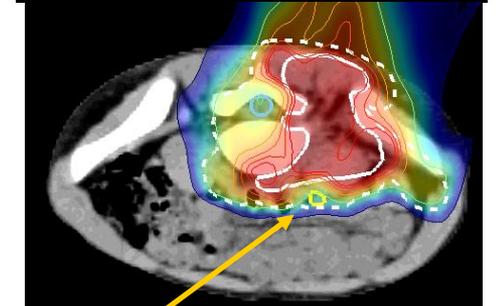
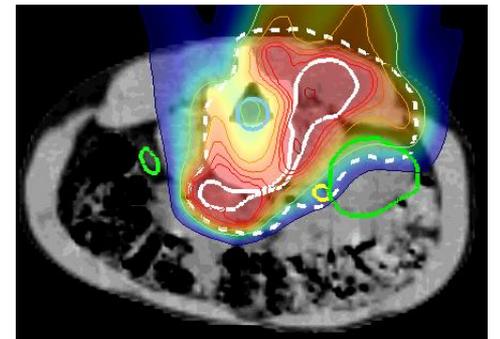
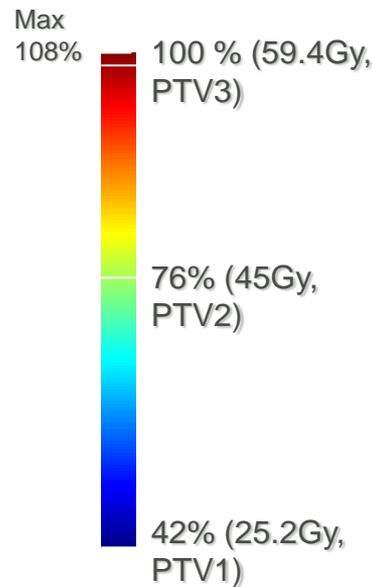
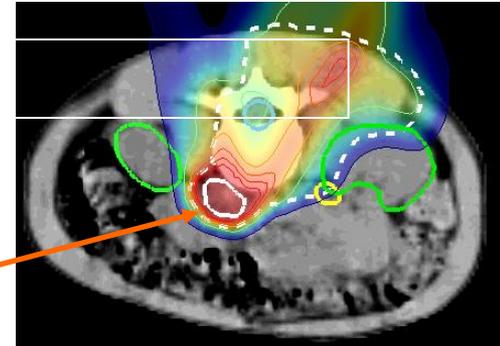
Isodose lines

- 42 %
- 76 %
- 90 %
- 95 %
- 98 %
- 100 %
- 105 %



PTV1 ..... PTV2 ----- GTV ———

GTV: Tumor thrombus in Vena Cava



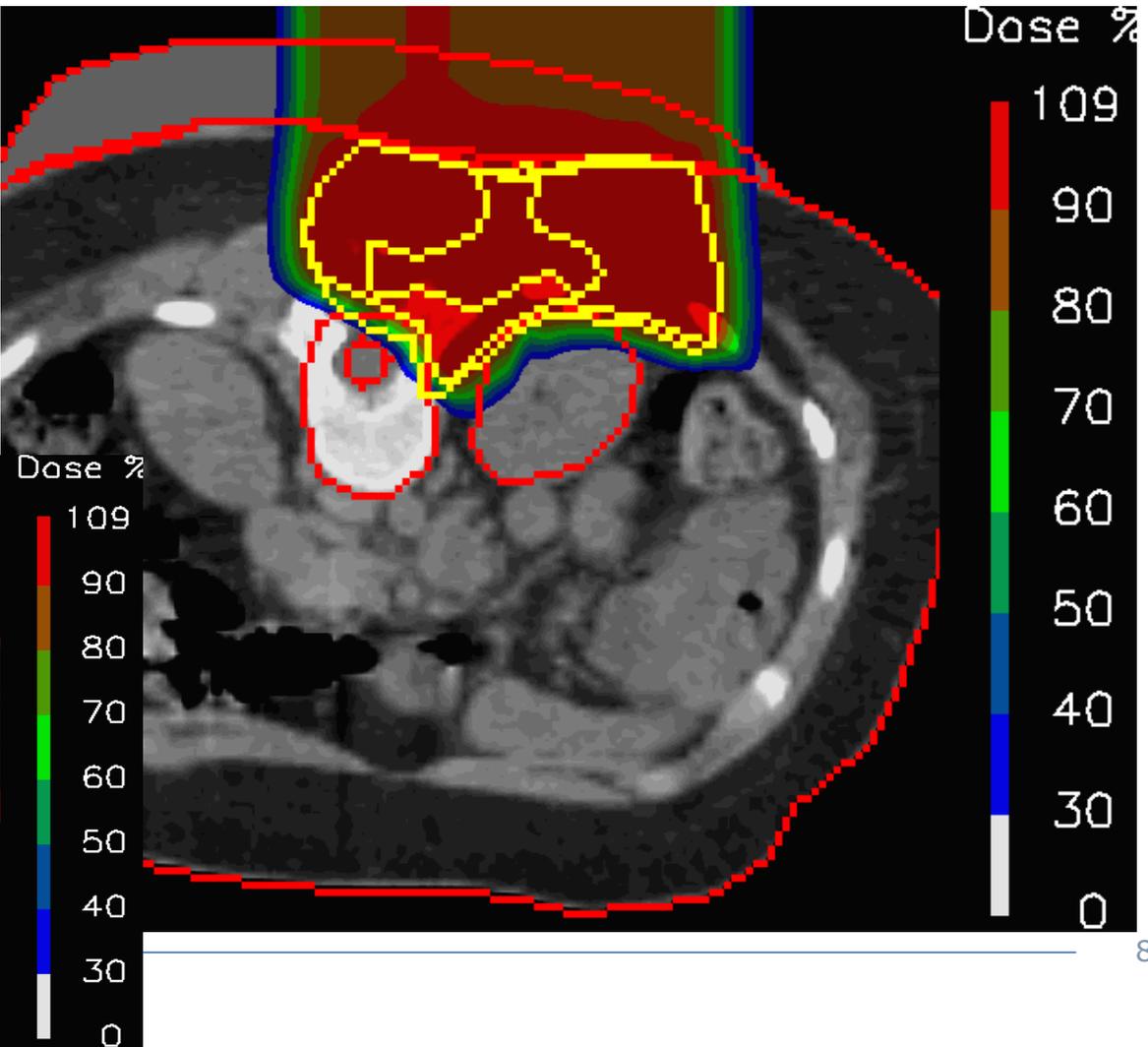
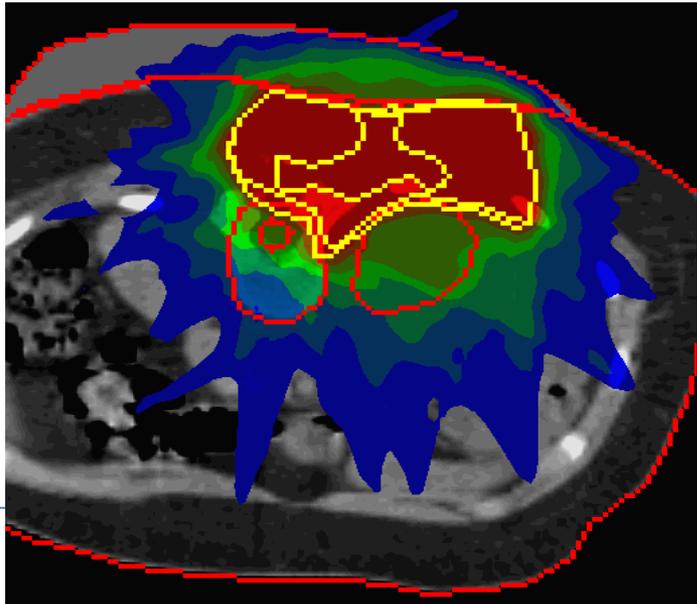
left Ureter

Note: Organ sparing in immediate proximity to the target

Dx: Rec. Desmoid,  
Boy, 2 yrs.

Dose:  
50.0 Gy + 10.0 Gy

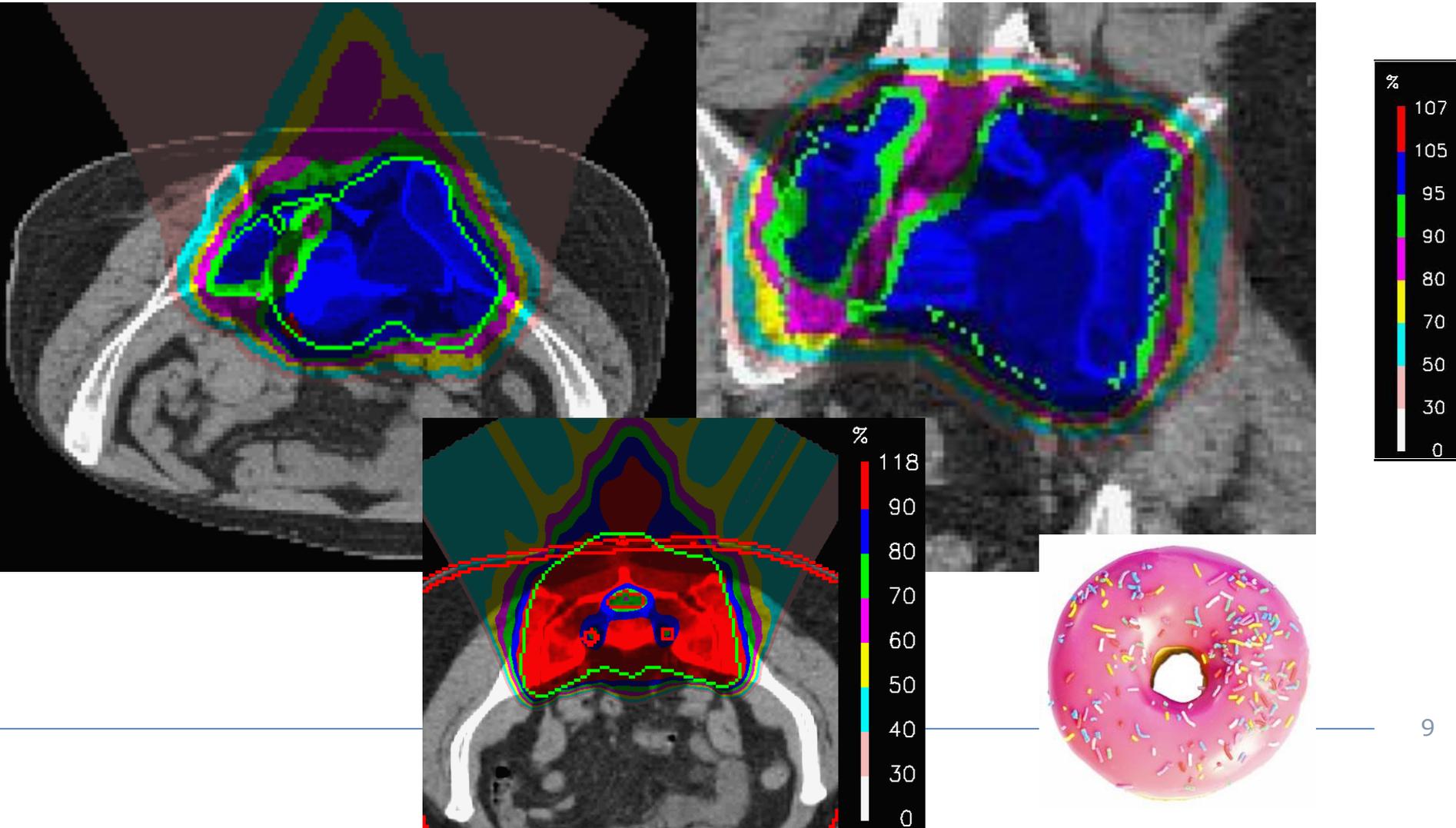
IMRT- Photons



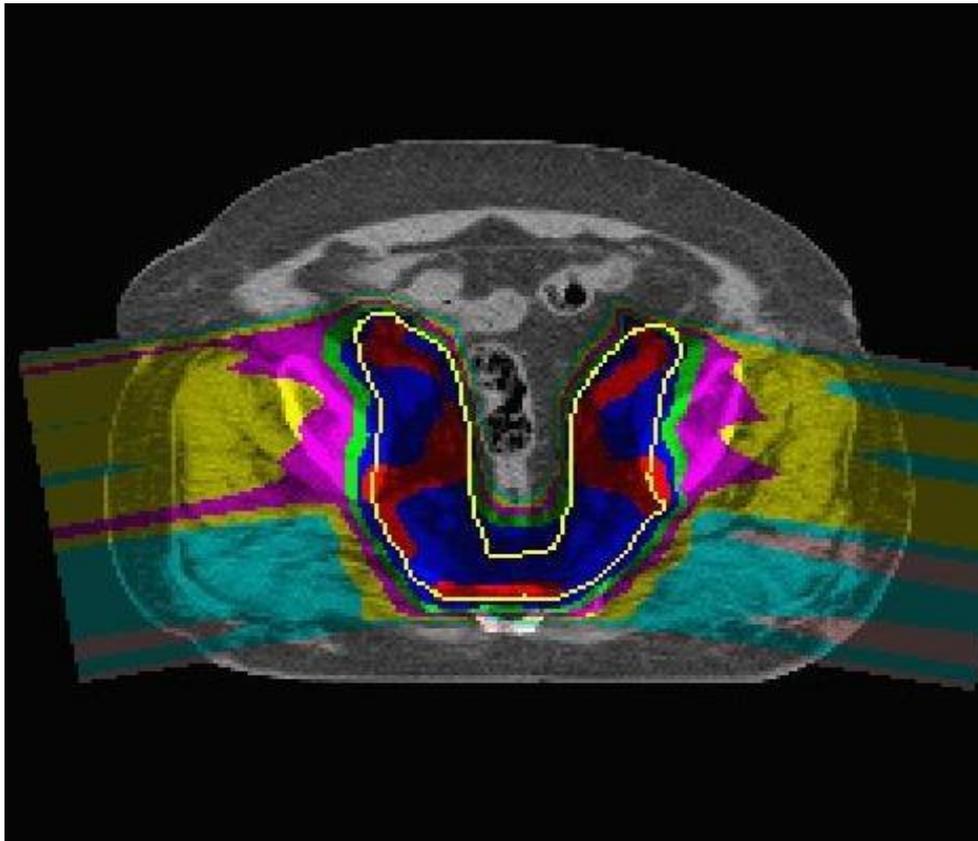
## Protons

**Active scanning: “sculpting” the dose:  
 Creating differential dose distributions within targets with  
 relative ease – hot (SIB) or cold regions (OAR)**

Example: sparing of selective nerve roots / cauda equina



Irradiation of Pelvic Lymph nodes with IMPT including selective boost.

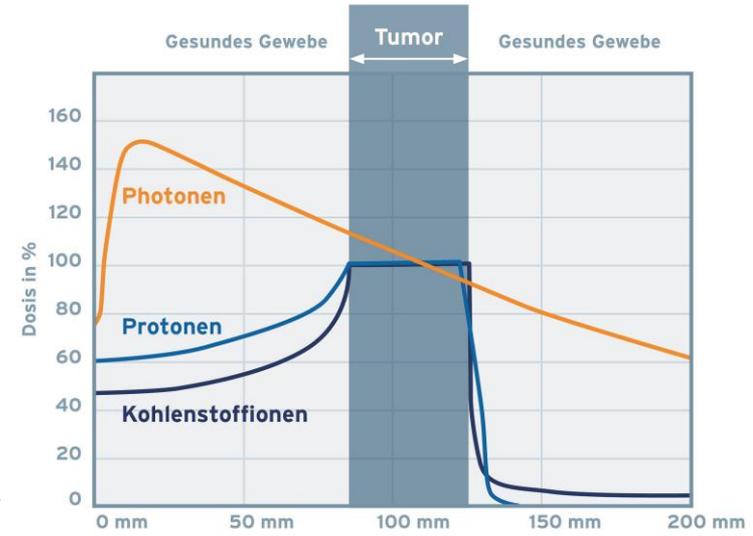
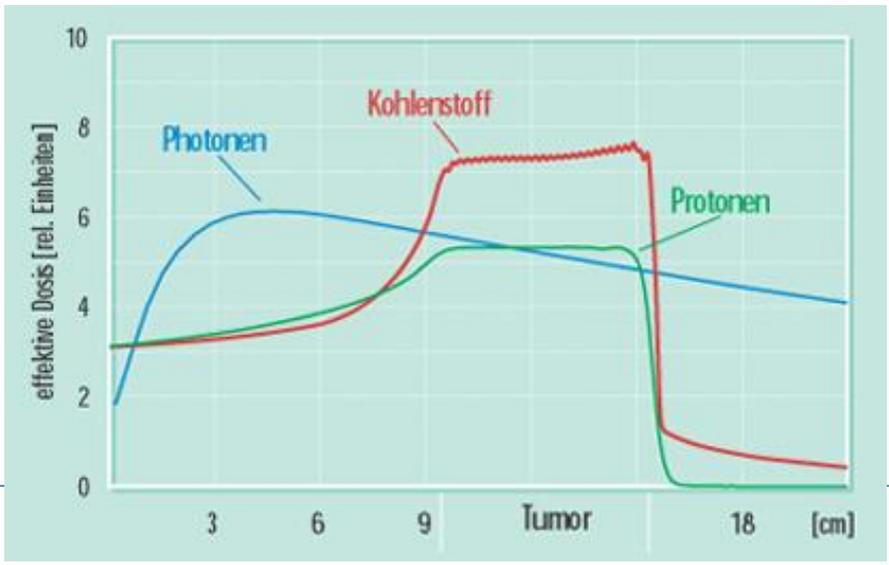


# Particle Therapy - The Quest for the Clinically Meaningful Benefit

Particle Community needs to focus disease-subpopulation-specific on realization of Proton's physical advantages and Carbon Ion's physical and radiobiological advantages

i.e.

**to identify the subpopulation of patients most likely to maximally benefit from use of particle therapy.**



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# From Construction to 1<sup>st</sup> Patient Treatment

# MedAustron

- Center for ion beam therapy and research
- Synchrotron allows protons and carbon ions
- „EBG MedAustron GmbH“ was founded in 2007
  - indirect ownership of the federal state of Lower Austria

# Investment

- Investment volume: € 200 million
  - 40% Accelerator
  - 30% Building
  - 10% Development & CE Certification
  - 10% Medical Infrastructure
  - 10% Others

# Financing

- Public sponsoring for „non clinical research“:
  - Republic of Austria
  - Federal State of Lower Austria
  - City of Wiener Neustadt
- Private financing of „clinic“
  - Bank loans
  - Equity/ guarantee from Federal State of Lower Austria

# Construction 2011- 2012

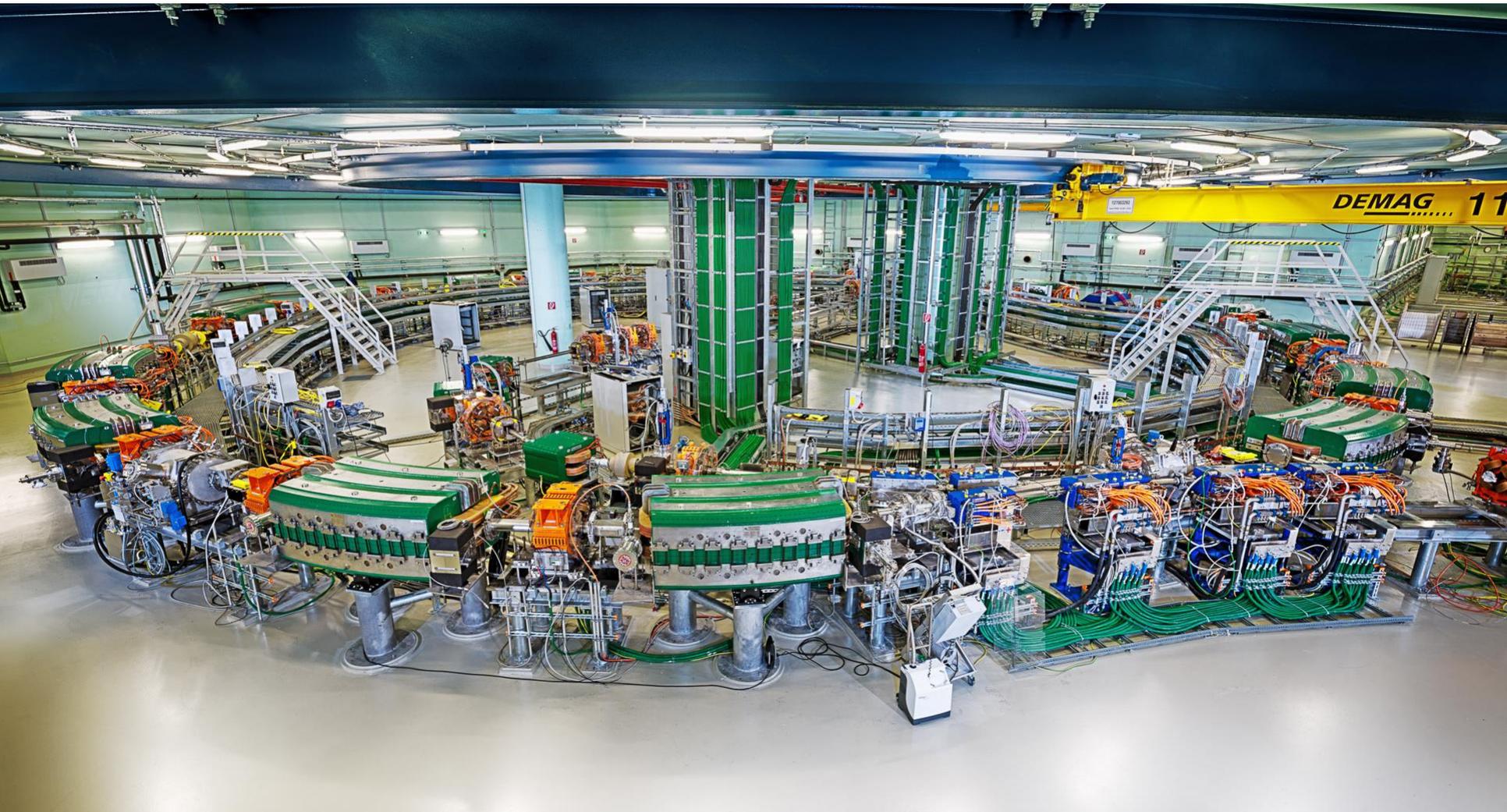




**March 2011: Construction Start**

# Oct 2012: Moving in







# Irradiation Rooms

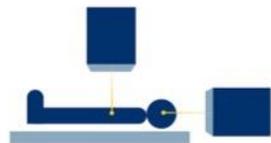


Horizontal Fixed Beam

**p** **c**

Protons & Carbon Ions

Patient Treatment



Horizontal and Vertical Fixed Beam

**p** **c**

Protons & Carbon Ions

Patient Treatment



Gantry

**p**

Protons

Patient Treatment



Horizontal Fixed Beam

**p** **c**

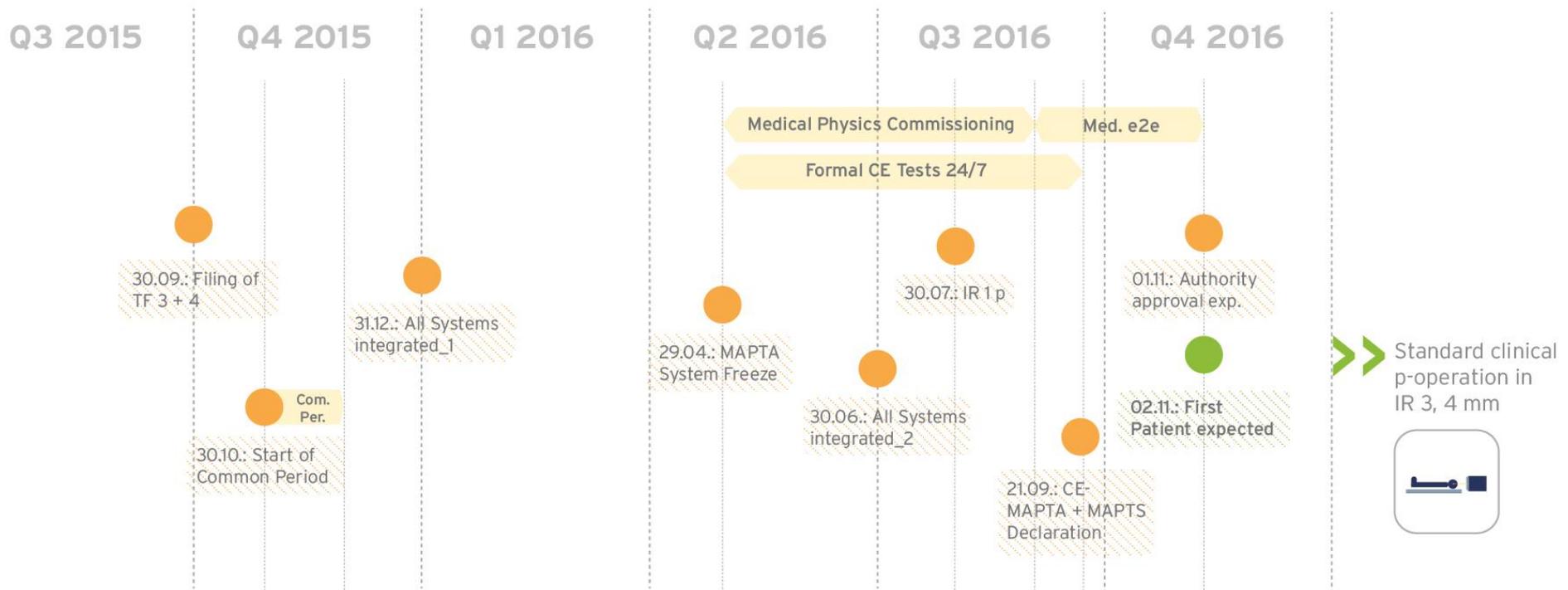
Protons & Carbon Ions

Non-clinical Research



## Unique System for Patient Positioning

## Timeline Short



## "Adapted" Ramp Up / Road Map

	2016	2017	2018	2019	2020	2021
Modalities	 H	 H+V	 H+V  H+V	 H+V+Gantry*  H+V	 H+V+Gantry  H+V	 H+V+Gantry  H+V
						
Rooms	1	2	2	3	3	3
Med. Shifts	1	1	1	2	2	2

\* Gantry cardinal angles in Q3/2019

## Start of patient treatments in 2016

- ❑ Start in one treatment room =
  - Horizontal fix beam line with protons**
- ❑ Parallel commissioning of the other treatment rooms
- ❑ Increasing functionality by adding vertical beam line to horizontal beam line
- ❑ Carbon ion treatment will be starting in 2018
- ❑ The gantry will be available by 2019/2020
- ❑ Continuous increase of patient numbers

# Research at MedAustron

- **Clinical Research**
- **Non-clinical research**
  - Radiobiology
  - Medical Translational Physics Research
  - Experimental Physics





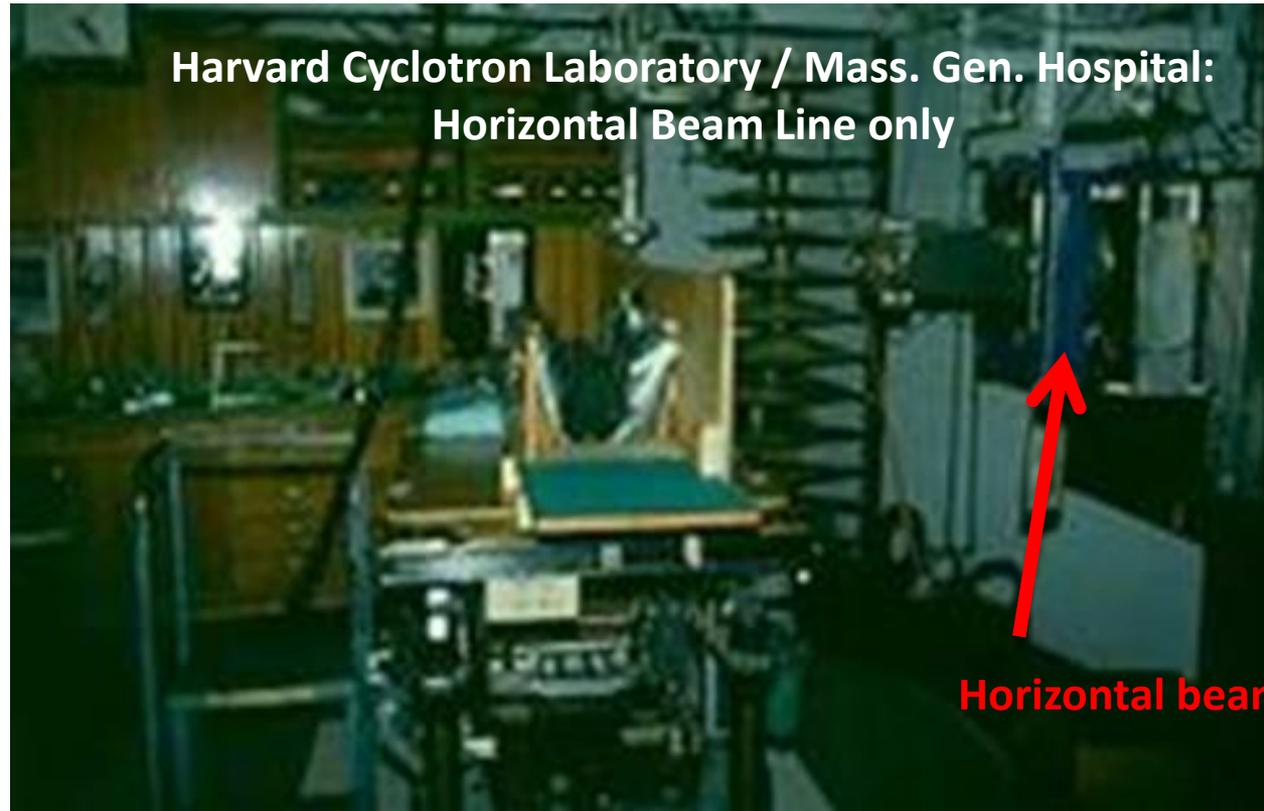
# MedAustron: Clinical Perspectives and Indications

## "Adapted" Ramp Up / Road Map

	2016	2017	2018	2019	2020	2021
Modalities	 H	 H+V	 H+V  H+V	 H+V+Gantry*  H+V	 H+V+Gantry  H+V	 H+V+Gantry  H+V
						
Rooms	1	2	2	3	3	3
Med. Shifts	1				2	2

Clinical Implications

\* Gantry cardinal angles in Q3/2019



- Many long term data of “established indications” are based on treatments performed using horizontal beam line (chordomas, chondrosarcomas, skull base, paraspinal, prostate, pediatrics) at HCL
- Scanning technology increases capabilities to deliver treatment in horizontal planes
- We expect that approx. 30% of particle-eligible initial indications will require added field entries (CSI a.o.)

# MedAustron – Indications 2017/18

(Note: Majority „established“ indications)

Skull Base (Chordomas, Chondrosarcomas)	RE-Irradiation
Pediatric Tumors (CNS, Sarcomas, other solid tumors)	Prostate
Head and Neck (Adenoid-Cystic, Paranasal Sinus)	
Sarcomas (Paraspinal, H&N, Truncal, Extremities)	
CNS (Meningiomas, Gliomas)	
Liver	

..... If suitable for horizontal and/or horizontal and vertical beam line

# MedAustron – Indications 2020

(Note: already presently treated internationally and under study evaluation)

Skull Base	RE-Irradiation
Pediatric Tumors	<b>Thoracic Malignancies</b> Non-small-cell Lung Cancer (central, Stage III) Lymphomas Thymomas, Mesotheliomas
Head and Neck (Adenoid cystic, Paranasal sinus) <b>Parotid, Oropharynx, Re-irradiation with larynx preservation</b>	<b>Breast-Ca</b> left-sided, complex lymphatic RT, unusual anatomy
Sarcomas	<b>Gastrointestinal (upper)</b> Esophagus, Pancreas
CNS <b>(high grade gliomas??)</b>	<b>Gastrointestinal (lower)</b> Anus, Rectum
Prostate	
Liver	

**Patient treatments  
at MedAustron  
will be embedded  
in clinical studies**

## Clinical Study Opportunities and Concepts

- **“Registry-Study” = prospective data evaluation with 90% - 100% recruitment of patients treated at MedAustron**
- **Disease-specific Clinical Studies implemented at MedAustron and developed in collaboration with Austrian Oncology Groups**
- **Multicenter Trial participation with international Proton Centers**
- **Multicenter Trial participation with international Carbon-Ion Centers (HIT, CNAO NIRS)**

# Registry Study

- an „umbrella“ study to prospectively capture data on all patients treated at the center
- Data capture:
  - Tumor-specific data (local control, disease progression)*
  - Patient Status (disease-specific survival, overall survival etc.)*
  - Acute and Late Side-Effects**
  - Optional: Quality of Life*
- Status: Conditional approval by Ethics-Committee

## **Disease-specific Study Protocols at MedAustron developed collaboratively with Austrian Rad Onc Centers (A-IONTREB) (P. Georg, Study Protocol Office)**

- Chordomas, Chondrosarcomas (Skull Base, Paraspinal, Sacrum)
- Soft Tissue Sarcomas (pre- and postoperative)
- Meningiomas (Grade I-III)
- Gliomas (low- and high-grade (III))
- Prostate Ca (definitive and postoperative)

## Potential International Collaborative Partners for clinical trials' participation

- NRG – formerly RTOG (Radiation Therapy Oncology Group) / USA
- COG (Children's Oncology Group) / USA and SIOP / Europe
- PCG: Proton Collaborative Group / USA;
- “single Institution-initiated” proton studies (MGH, MD Anderson, Univ. Florida, Univ. Penn., a.o.)
- Emerging European Initiatives: EORTC, IPACS
- Carbon Ion Studies: HIT (Heidelberg), CNAO (Pavia), NIRS (Japan)

# Challenges of Clinical Particle Therapy

## —● Protons:

- Paradigm: Particle Therapy has to demonstrate a significant clinical benefit over photon RT
- Define most suitable tumor entities and patient subpopulations = clinical studies
- Emerging Technology = further improvements on beam delivery, planning aspects, imaging etc.
- Refinement / revision of universal concept of proton RBE = 1.1

# Challenges of Clinical Particle Therapy

## —● Carbon-Ions:

- Will Carbon Ions add a significant advantage over protons?
- For which tumor entities and patient subgroups?
- Will specific radiobiological advantages in vitro/in silico /in vivo translate into clinical results?
- Will the international Carbon Ion Community be able to repeat single-institution results?
- Further refinements of RBE for tumors and normal tissues and therefore further refinements of biologically equivalent dose determination in treatment plans

# Present, Radiobiologic Challenges of Clinical Particle Therapy

## —● Protons:

- RBE optimization at end-of-range; revision/refinement of the universal RBE Concept of „1.1“
- Incorporation of optimized RBE into treatment planning system

## —● Carbon-Ions:

- RBE refinement of tumors and normal tissues
- Increase RBE optimization and therefore dose precision of Local Effects Model, i.e. Treatment Planning System



**MedAustron is a stand-alone Ion Therapy Facility within Austria**

**MedAustron will be integrated in the Radiation Oncology Community Austrias and the Particle Therapy Community worldwide based on Clinical, Translational and Basic Science Research Collaborations.**