

Developments in proton and light-ion therapy

Sandro Rossi

Fondazione CNAO

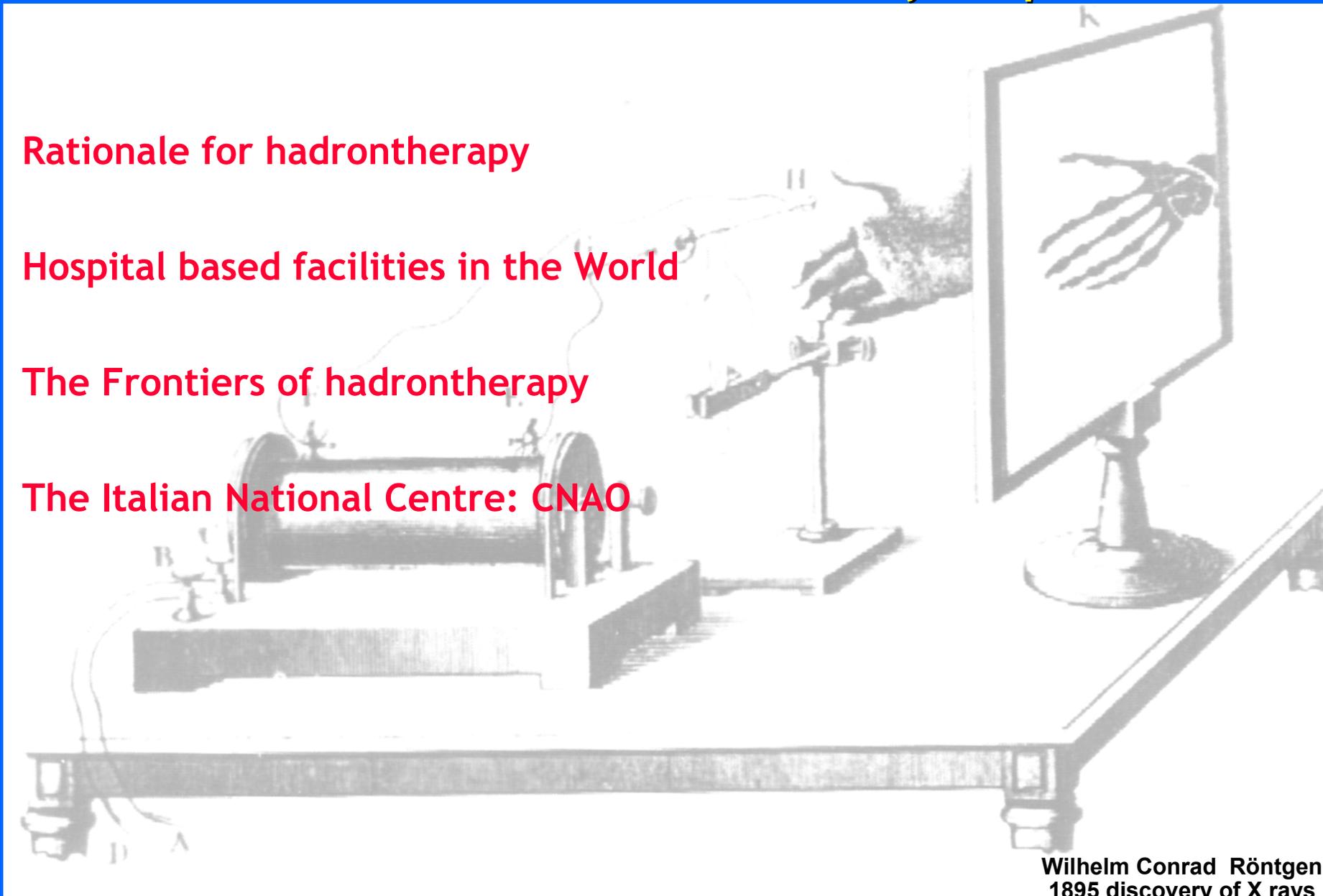
Outline of the presentation

Rationale for hadrontherapy

Hospital based facilities in the World

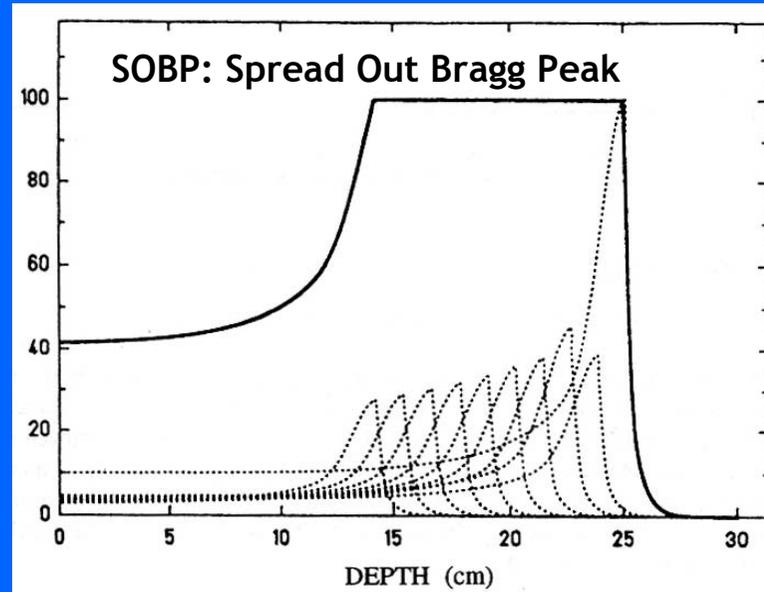
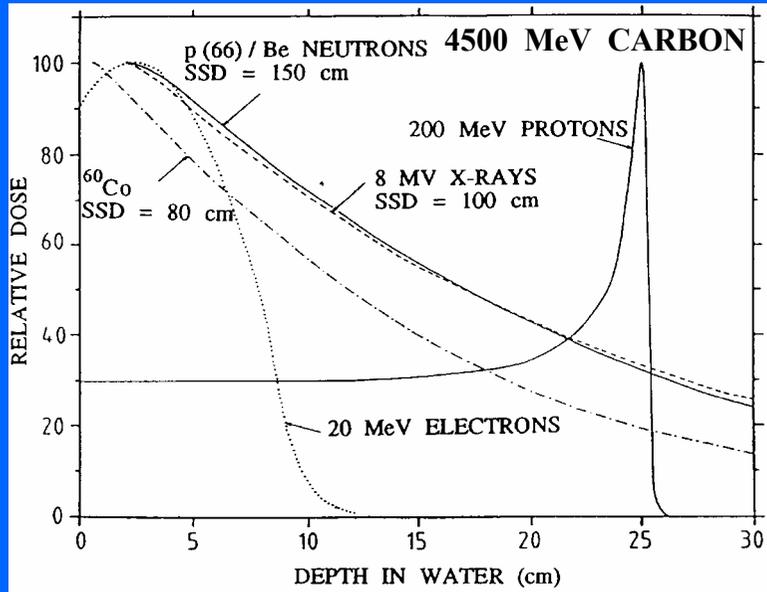
The Frontiers of hadrontherapy

The Italian National Centre: CNAO

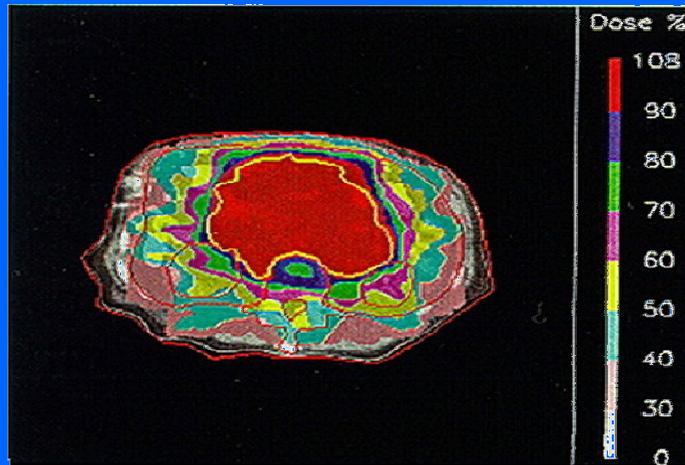


Wilhelm Conrad Röntgen
1895 discovery of X rays

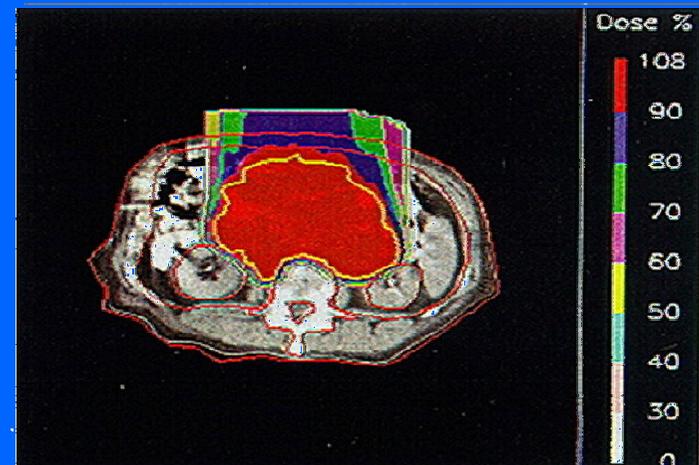
Hadrons: conformal dose irradiation



Abdomen



X-ray (IMRT) - 9 fields



Protons - 1 field

Results with protons

CHORDOMAS OF THE BASE OF THE SKULL		Number of Patients	OS 5 years	OS 10 years	PFS 5 years	PFS 10 years
X-rays	U. Michigan 1986	21	50 %	20 %	-	-
	R. Marsden 1988	25	44 %	17 %	33 %	20 %
	Mallinckrodt 1991	21	74 %	46 %	30 %	-
	Mayo Clinic 1993	51	51 %	35 %	33 %	24 %
	Princess Margaret 1996	13	-	-	15 %	-
Protons	MGH/HCL 1996	169	80 %	54 %	64 %	42 %

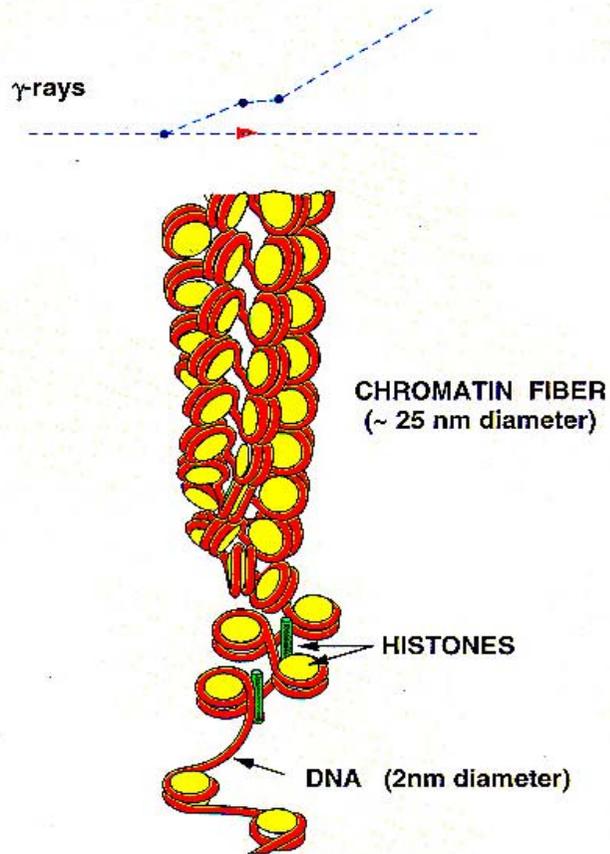
Indications for protontherapy

every 10 M EU citizens

12% of X-ray patients

2'400 pts/year

Carbon ions: biological efficacy

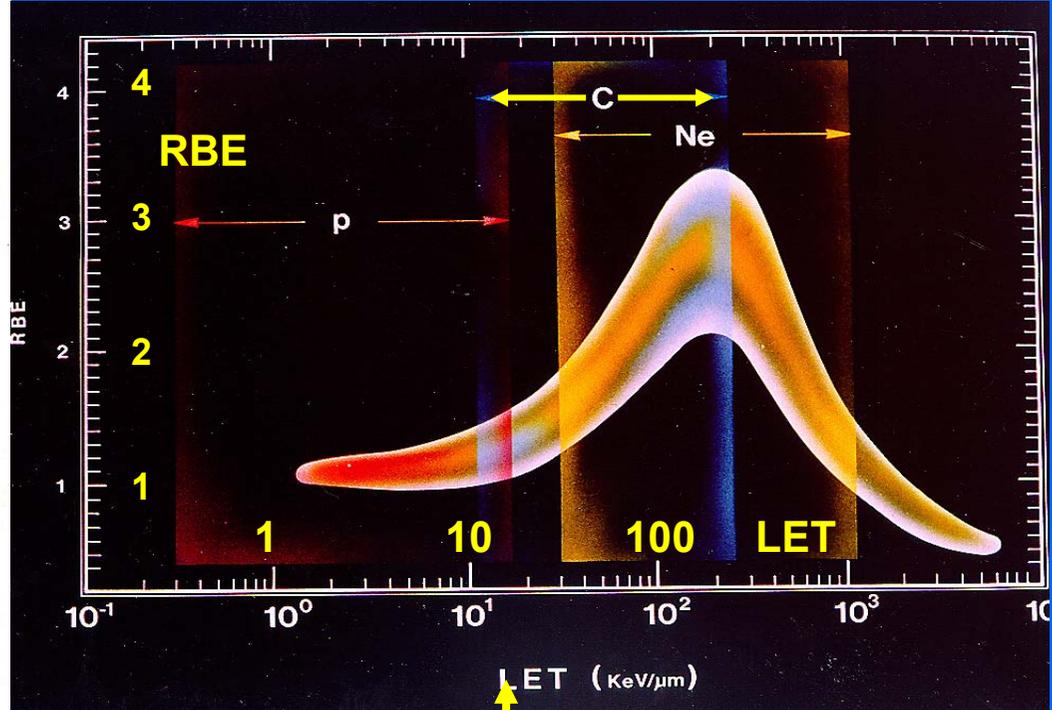


1 MeV protons

1 MeV/u α -particles

1 MeV/u C ions

10 nm



$$10 - 20 \text{ keV/mm} = 100 - 200 \text{ MeV/cm} = 20 - 40 \text{ eV/(2 nm)}$$

Reduced effect dependence from Oxygen content

Results with carbon ions (Chiba)

NSCLC = Non Small Cell Lung Cancer
182 patients treated with carbon ions

Comment:

The results with CIRT in early stage NSCLC are impressive with a local control rate ranging from 62% to 100% and a 3 year survival between 65 and 88%. Two Japanese studies with modern photon beam radiotherapy in early stage NSCLC can be mentioned in comparison. In the study reported in 1997 by Morita *et al*, 149 patients with stage I were treated with a total dose of 64.7 Gy in 32 fractions. The local control rate was 56% and 3 year survival 34.2%. In the study reported in 1999 by Hayakawa *et al*, 36 patients received 60-81 Gy in 2 Gy fractions. The local control rate was 80.6% and 3 year survival 42%.

Indications for carbontherapy

every 10 M EU citizens

3% of X-ray patients

600 pts/year

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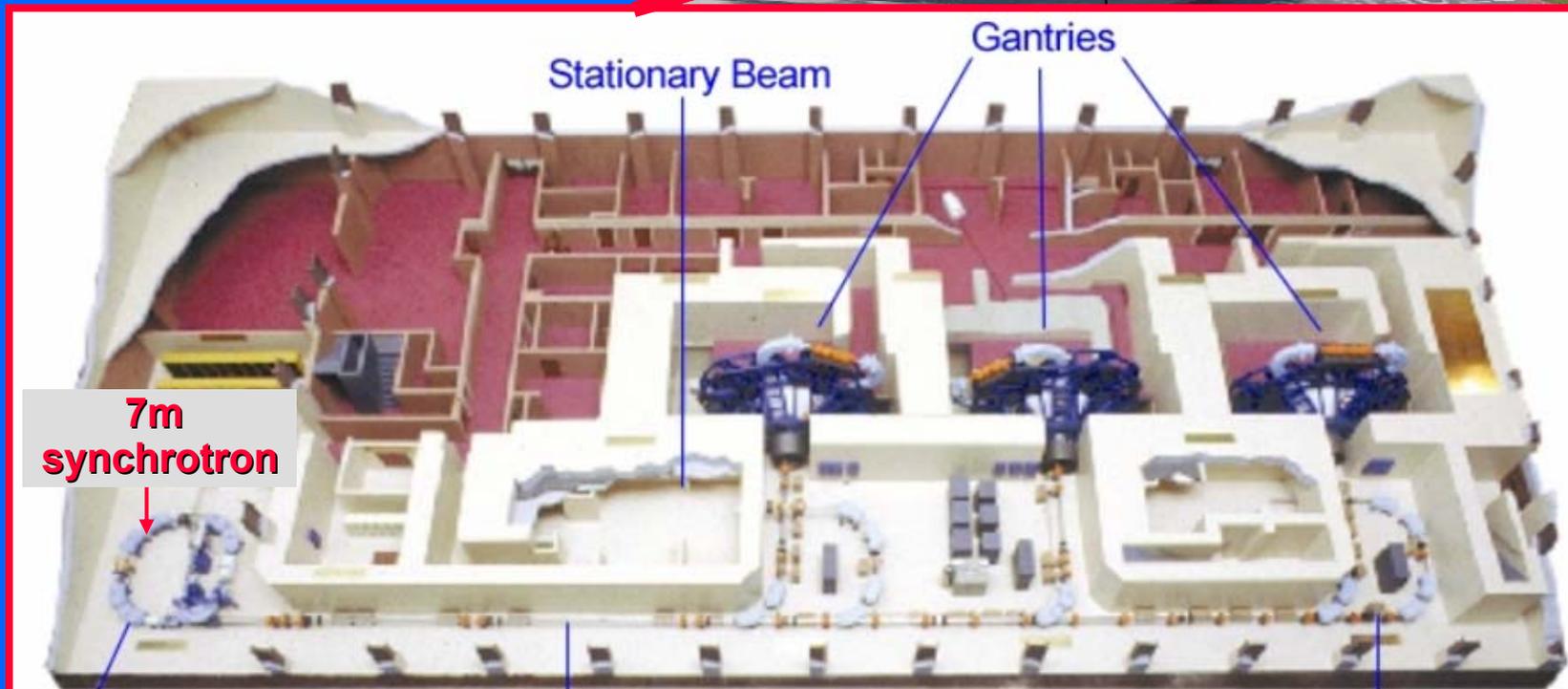
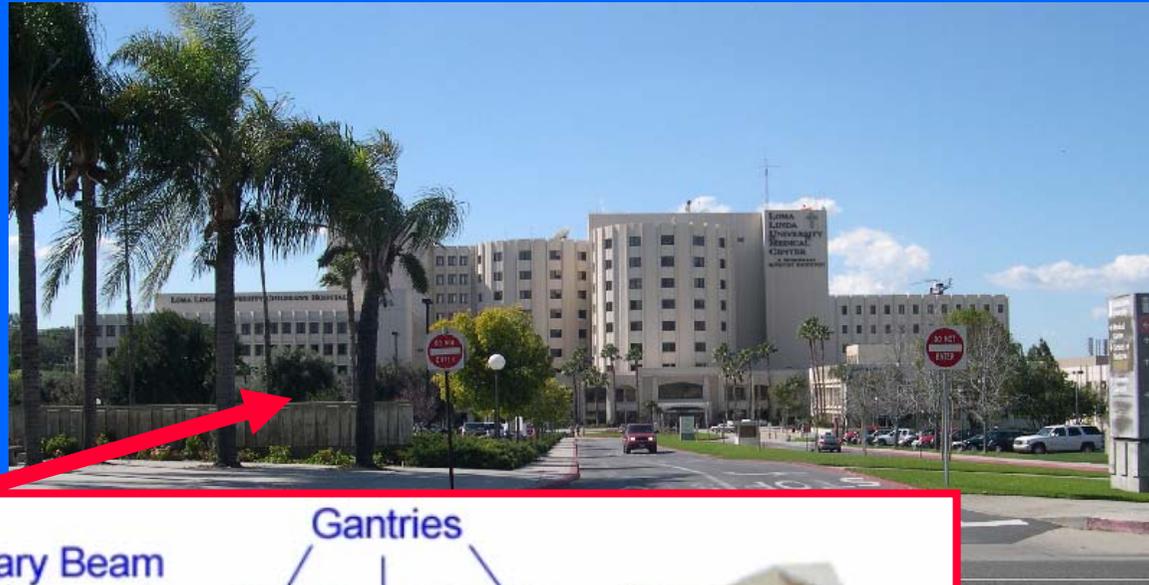
The Italian National Centre: CNAO



Wilhelm Conrad Röntgen
1895 discovery of X rays

Loma Linda University Medical Center: first patient 1992

- First hospital-based proton-therapy centre
- 2005:160 sessions/day



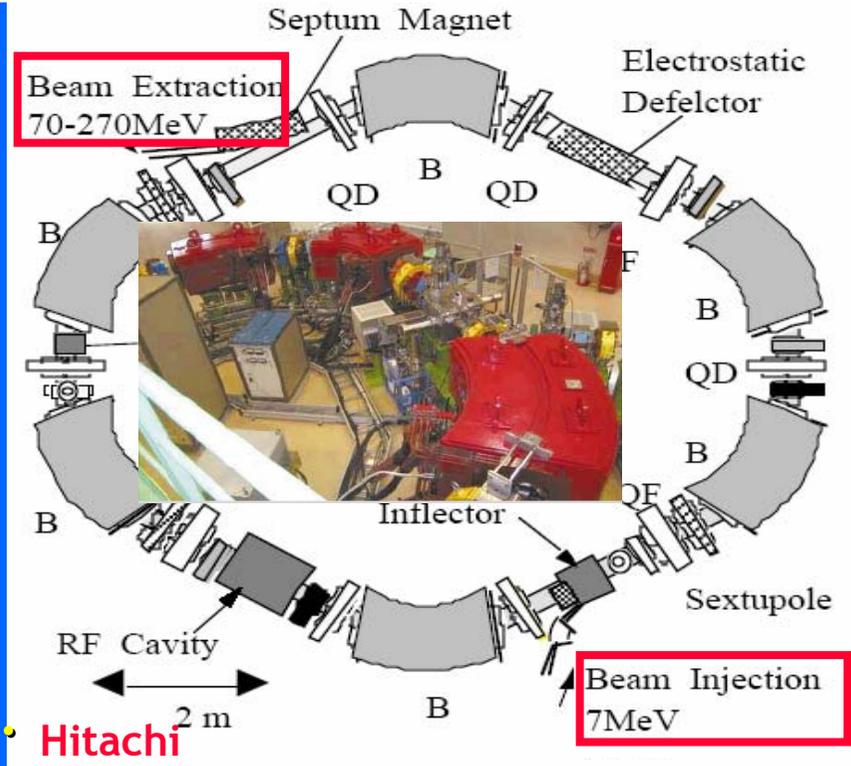
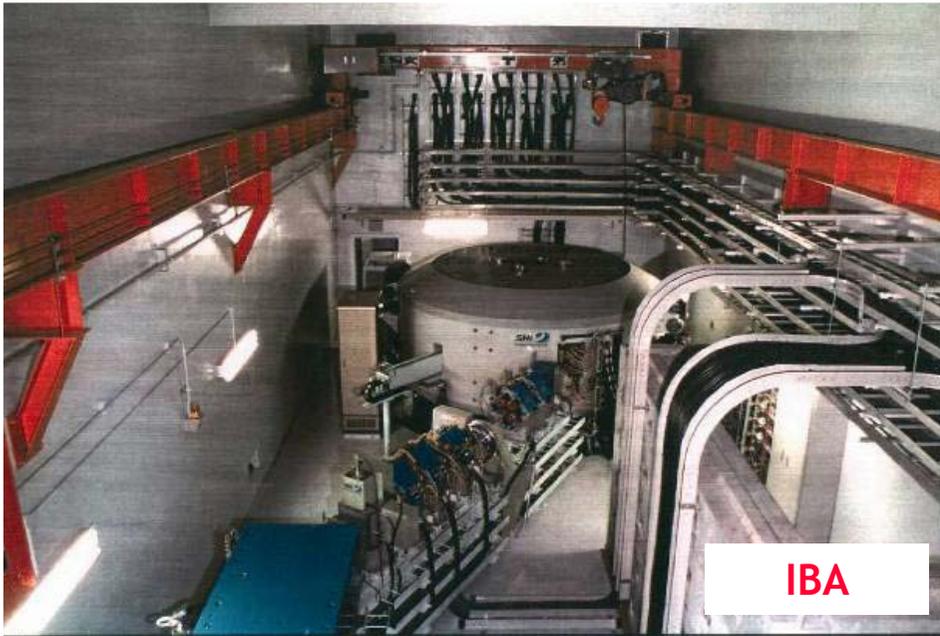
Protons: ten years of tenders

Hospital centres for deep protontherapy (>500 pts/year)

5 in USA, 4 in Japan, 2 in China, 1 in Korea, 1 in Switzerland, 1 in Germany, 1 in France and 1 in Italy

(running or financed)

Year	Customer	Provider
1995	MGH, Boston MA, USA	IBA
1996	NCC, Kashiwa, Japan	SHI-IBA
1996-99	Tsukuba University	Hitachi
	Wakasa Wan Energy Research Center	Hitachi
	Shizuoka Prefecture	Mitsubishi
2001	PSI – Villigen, Switzerland	ACCEL
	Wanjie Tumor Hospital – Zibo, China	IBA
	Chang An PMC – Beijing, China	IBA
2002	Rinecker PTC – Munchen, Germany	ACCEL
	Korean NCC - Seoul	IBA
	IUCF (MPRI), Bloomington IN, USA	IBA
	M.D. Anderson CC, Houston TX, USA	Hitachi
2004	University of Florida, Jacksonville FL, USA	IBA



Protontherapy: a mature market...



Carbon Ions: Japan 2+ centres

WAKASA BAY PROJECT
 by Wakasa-Bay Energy Research Center
 Fukui (2002)
 protons (≤ 200 MeV) synchrotron
 (Hitachi)
 1 h beam + 1 v beam + 1 gantry

TSUKUBA CENTRE
 Ibaraki (2001)
 protons (≤ 270 MeV)
 synchrotron (Hitachi)
 2 gantries
 2 beam for research

HYOGO MED CENTRE
 Hyogo (2001)
 protons (≤ 230 MeV) - He and C ions (≤ 320 MeV/u)
 Mitsubishi synchrotron
 2 p gantries + 2 fixed p beam + 2 ion rooms

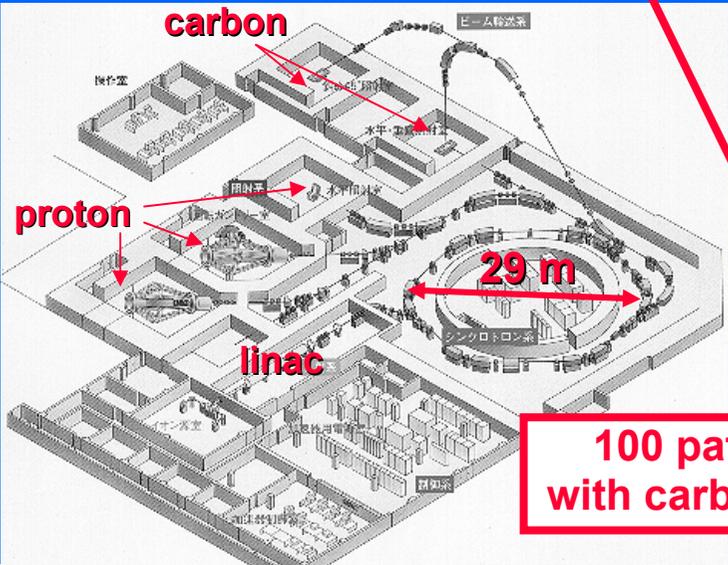
KASHIWA CENTER
 Chiba (1998)
 protons (≤ 235 MeV)
 cyclotron (IBA - SHI)
 2 Gantries + 1 hor. beam

HEAVY ION MEDICAL ACCELERATOR
 HIMAC of NIRS (1995)
 He and C (≤ 430 MeV/u) 2 synchrotrons
 2 h beams + 2 v beams
 + GUNMA UNIVERSITY

SHIZUOKA
 Shizuoka (2002)
 Proton synchrotron
 2 gantries + 1 h beam

2300 patients with carbon ions

100 patients with carbon ions



Mitsubishi: turn-key system

EUROPE: GSI pilot project and Enlight

**More than 200 patients treated
with carbon ions, active
scanning + In beam PET**



Approved projects (with ions): HIT (D)-CNAO (I)-MedAustron (A)-ETOILE (F)

Many projects around Europe...

***Coordination networks ENLIGHT (5 FP) and ENLIGHT++ (7 FP) :
European Network for LIGHT-ion Hadron Therapy ++***



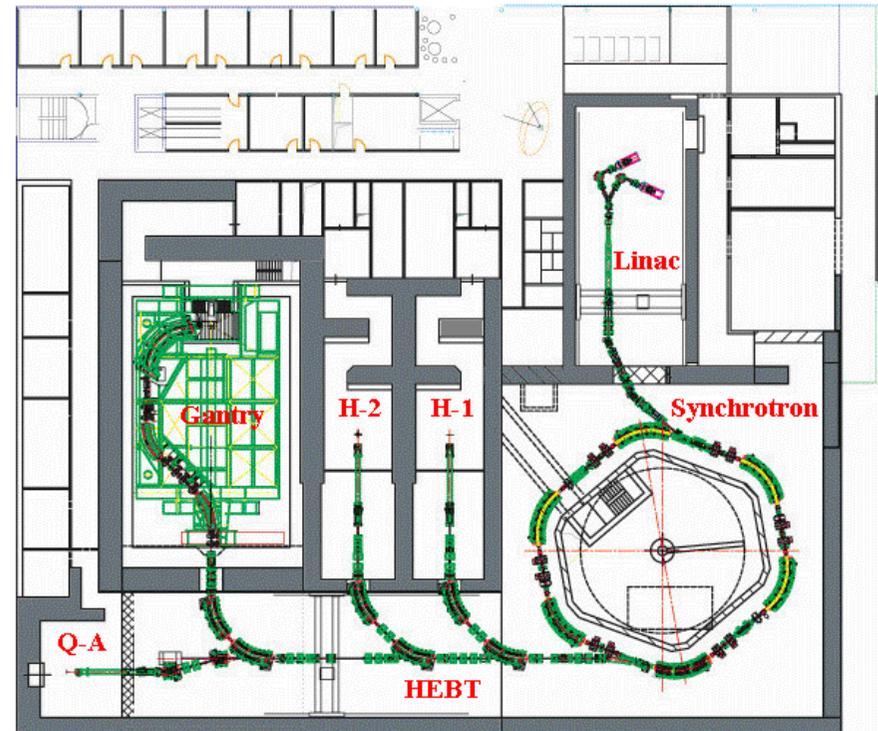
HIT

The Heidelberg Ion Therapy Center

First patient: end of 2007

- compact design
- rasterscanning only
- low-LET modality: Protons (later He)
- high-LET modality: Carbon (Oxygen)
- > 1000 patients/year
- > 15.000 fractions/year

(Pictures courtesy of T. Haberer)

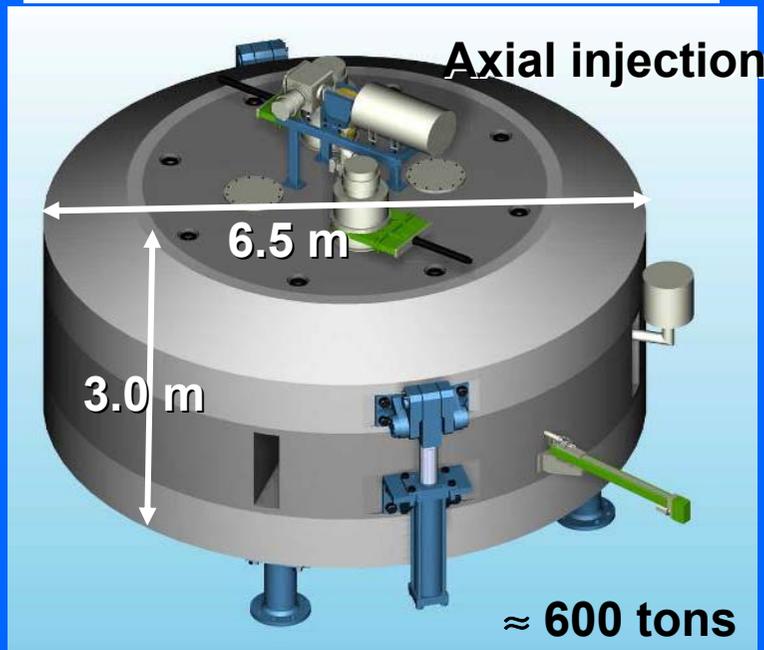


EU firms are interested, but "experts" support is still needed

Other firms:

ACCEL

IBA (400 MeV/U SC cyclotron)



(Picture courtesy of IBA/Y. Jongen)

GSI

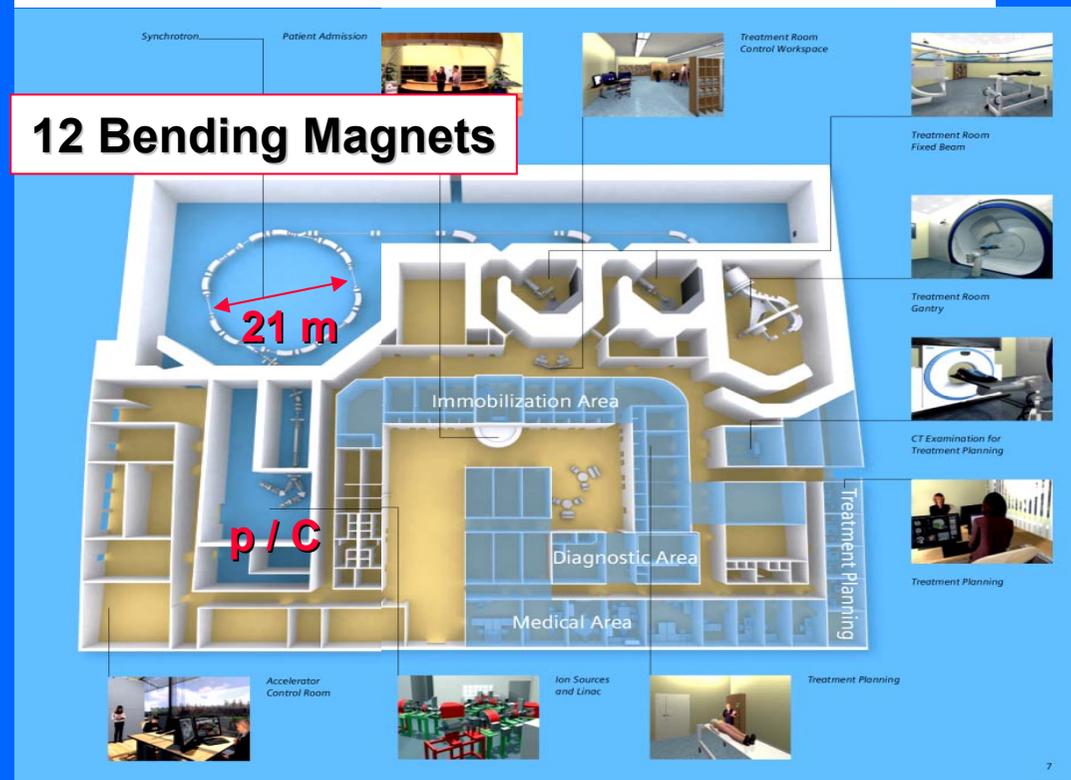
SIEMENS

Darmstadt, October 8, 2003

A decisive milestone in the battle against cancer

Siemens and GSI signed a contract for a new method irradiating tumors that also provides a larger patient cohort with access to this method.

12 Bending Magnets



(Picture courtesy of Siemens Medical)

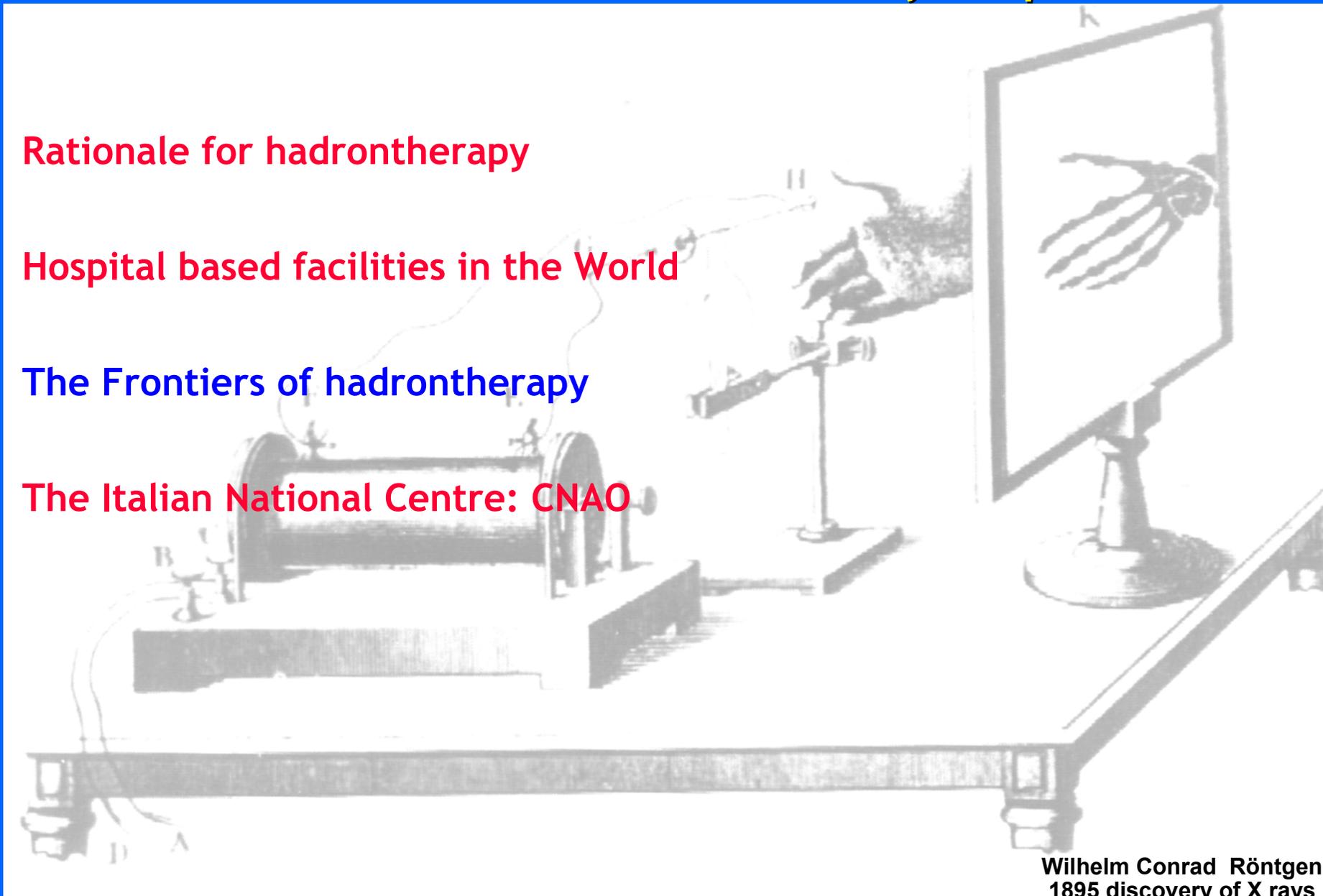
Outline of the presentation

Rationale for hadrontherapy

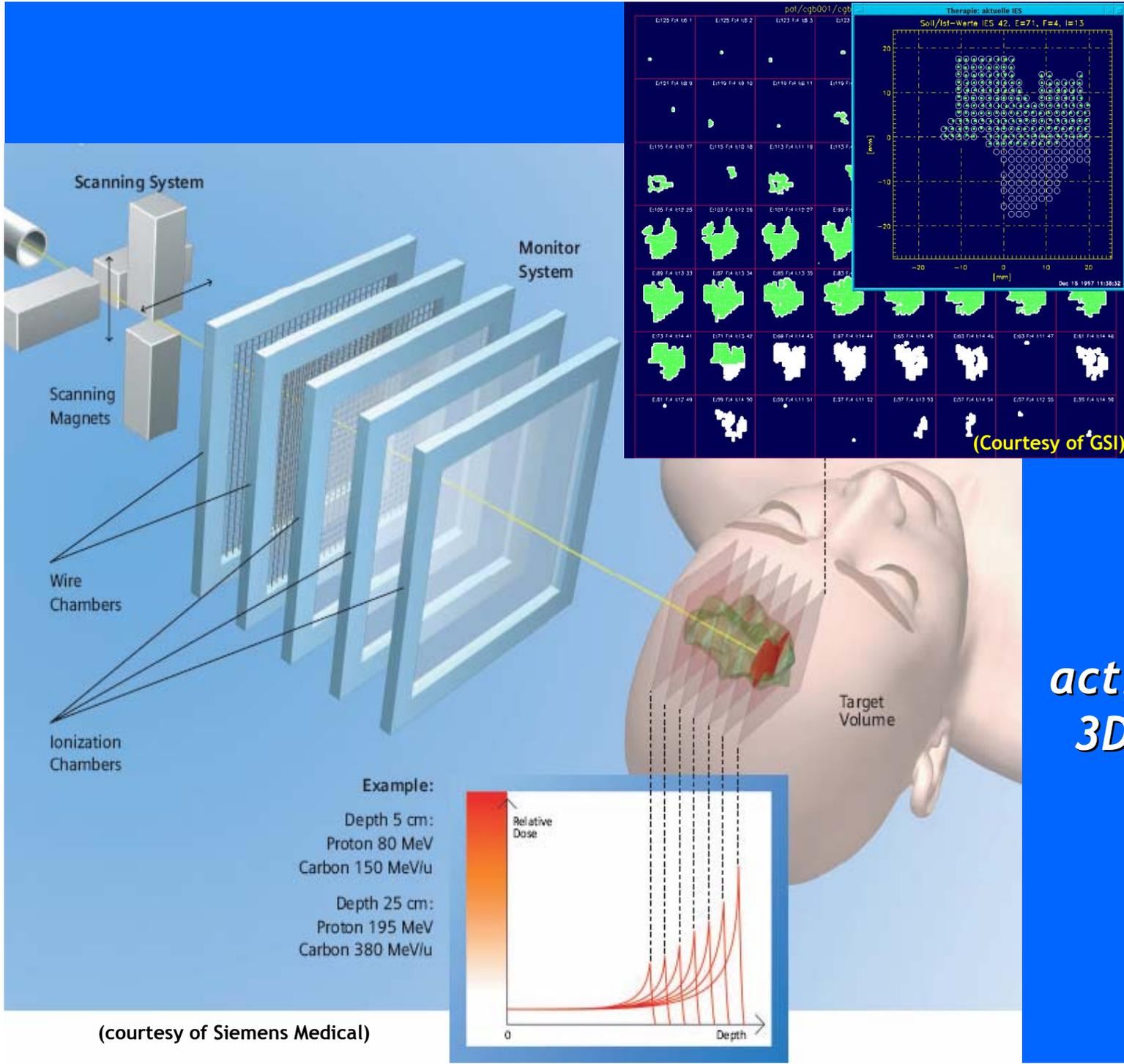
Hospital based facilities in the World

The Frontiers of hadrontherapy

The Italian National Centre: CNAO



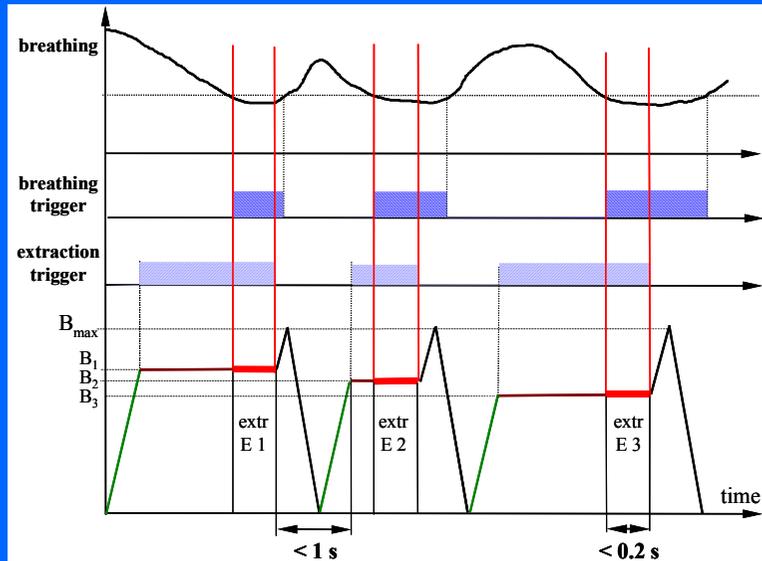
Wilhelm Conrad Röntgen
1895 discovery of X rays



(courtesy of Siemens Medical)

Tracking Optimisation (Time coordinate)

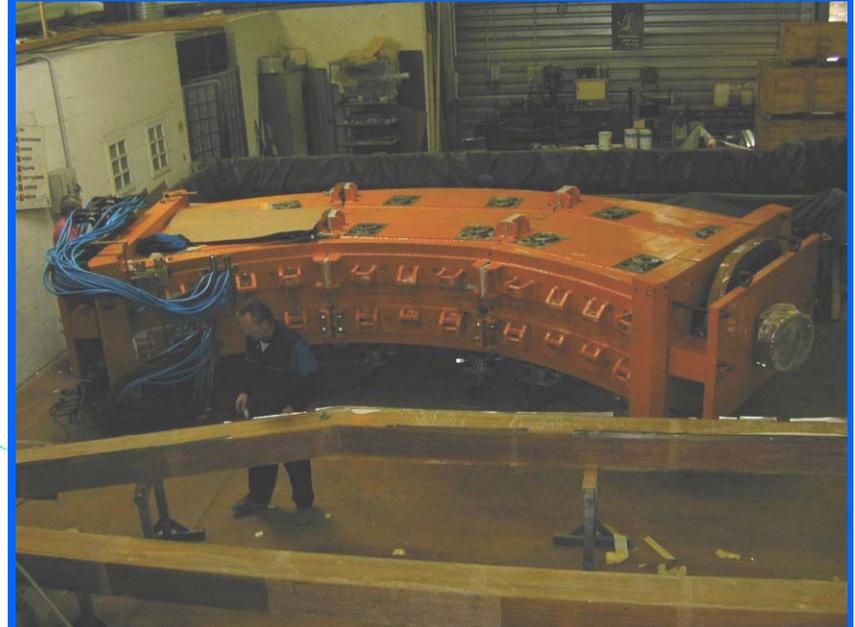
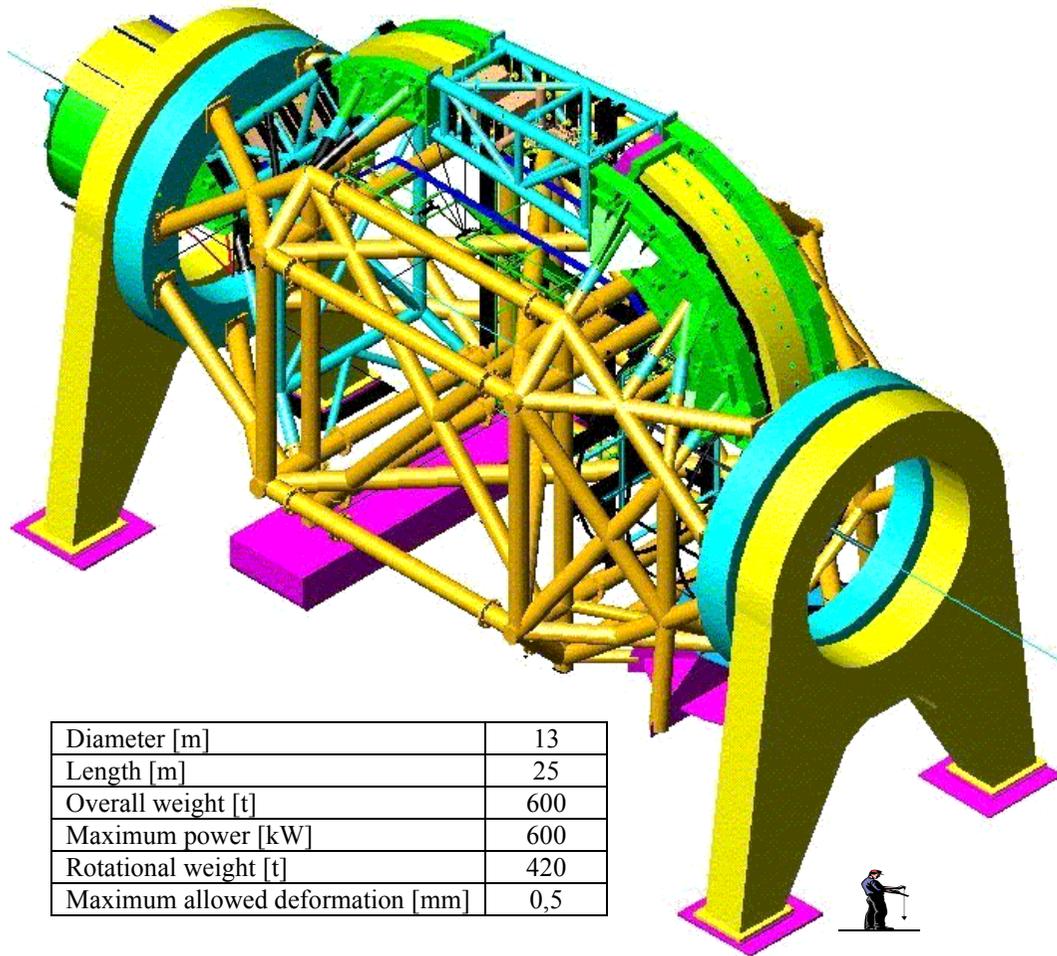
Already applied in Chiba:
breathing synchronisation



Interesting also for IMRT:
lots of efforts and devices

GSI Gantry - In construction for Heidelberg

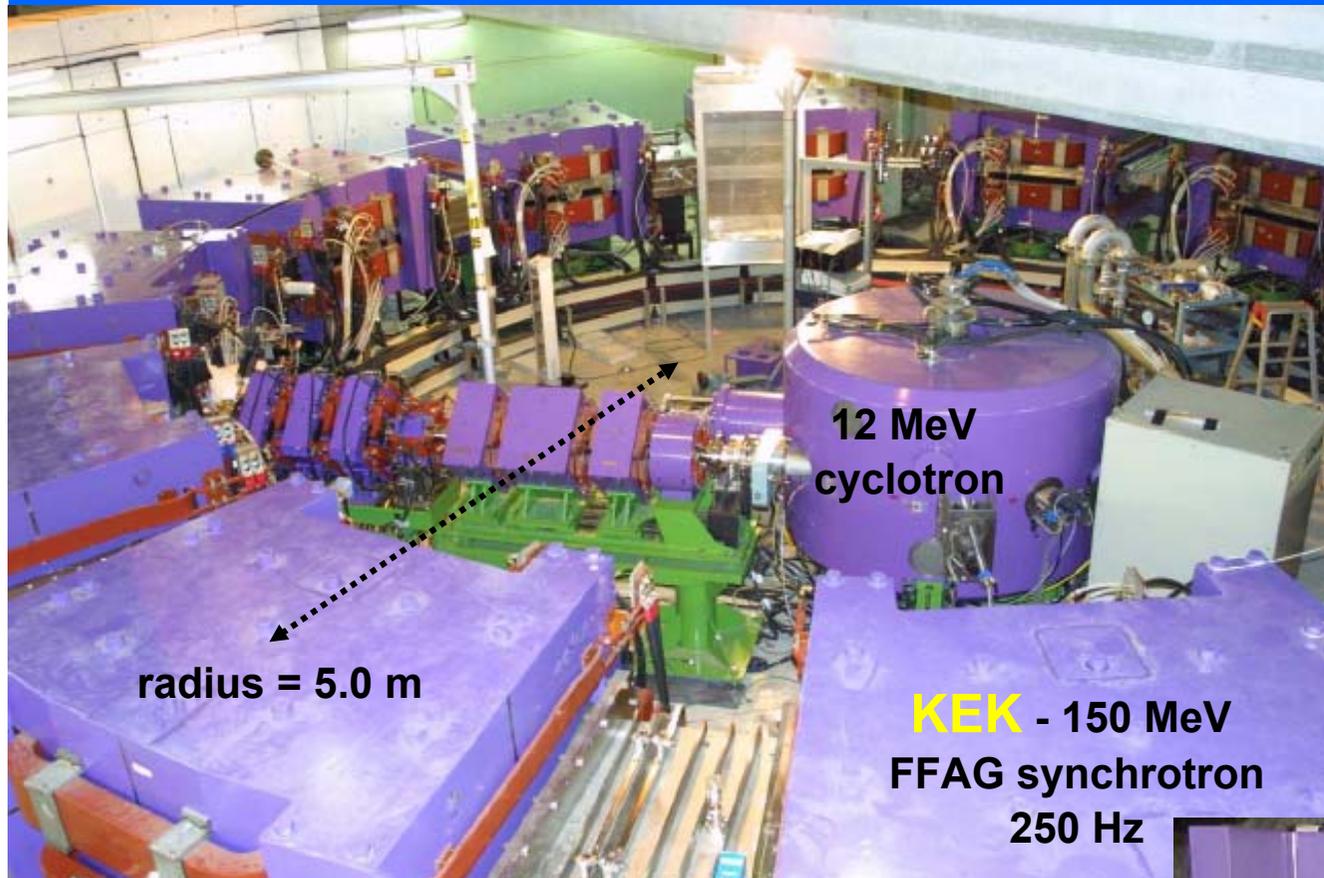
Adaptive radiotherapy: Gantry Optimisation



Superconductivity - FFAG magnets - ... ?

Integrated system: optics, technology, scanning, patient positioning

“Novel” accelerator concepts: FFAG synchrotron



Typical dFd dipole triplet
Return-yoke free magnet
easy injection+extraction



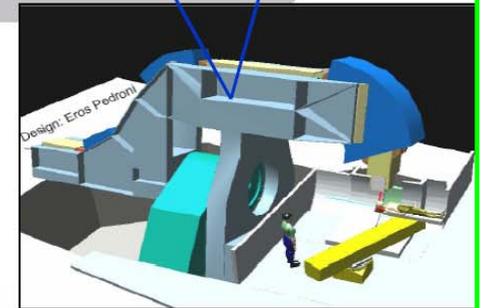
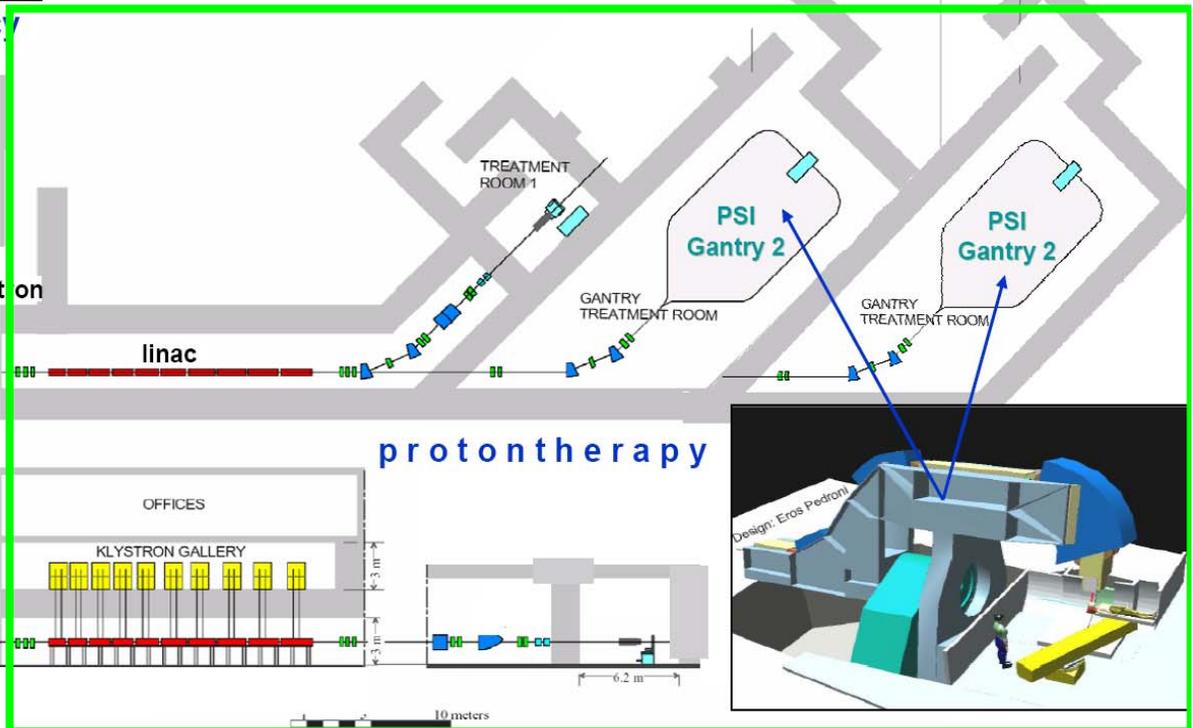
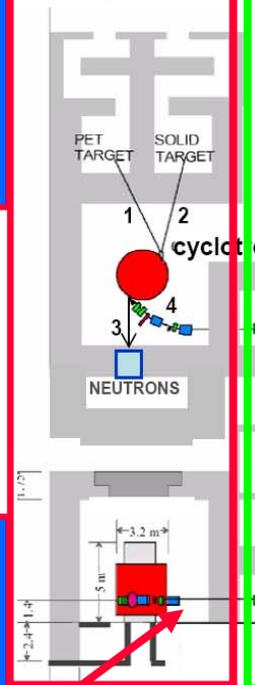
Aim: compact, high rep.rate (scanning),
cost-effective accelerators for hadrontherapy ?

IDRA = Istituto per la Diagnostica e la Radioterapia Avanzate
TERA - patent pending

Novel accelerator concepts: IDRA (cyclinac)

Radioisotopes etc.
Production
(PET, SPECT, BNCT...)
Commercial product

radiopharmacy



30 MeV
100 μ A – 1 mA



Linac Booster = LIBO (3 GHz) lenght = 16.4 m

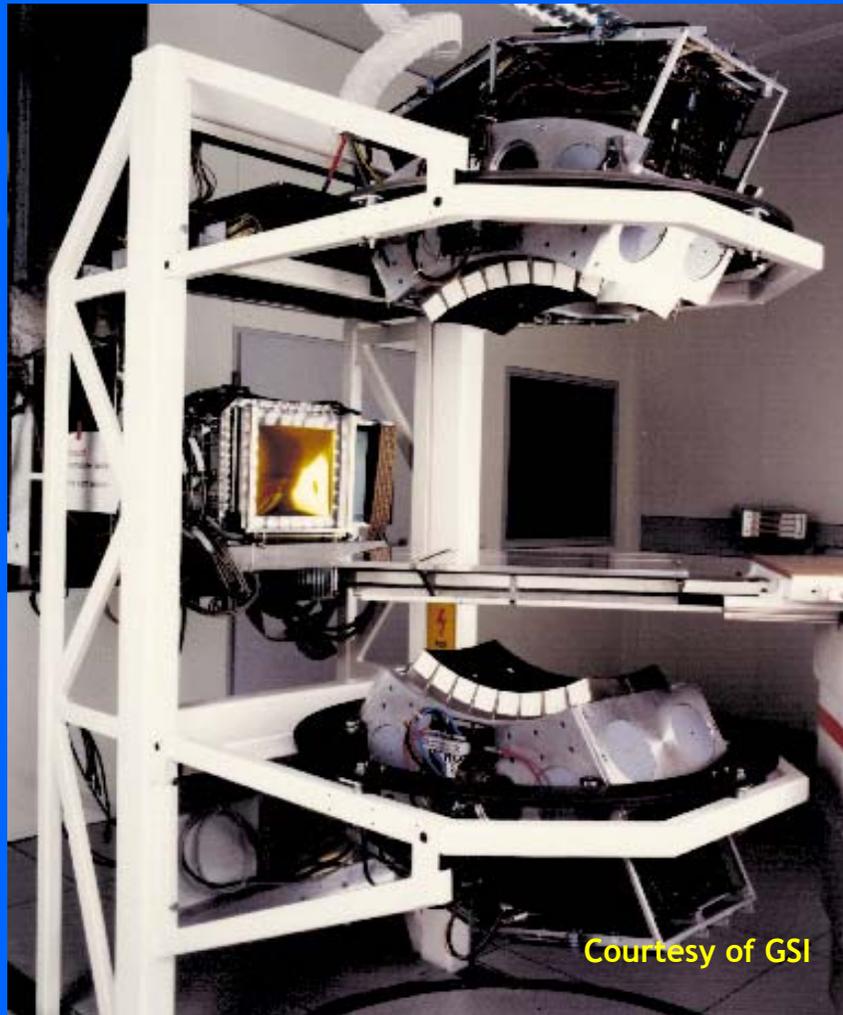
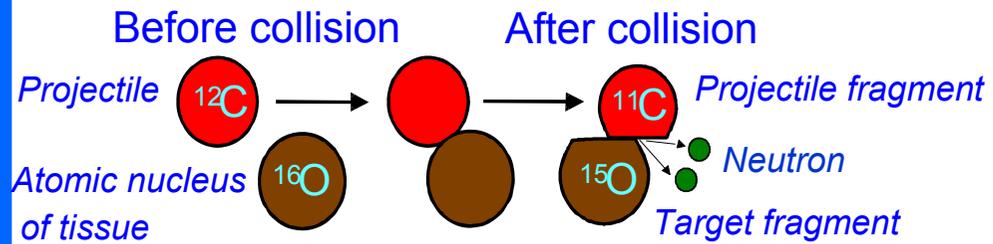
Protontherapy
Prototype successful

< 210 MeV
< 8 nA

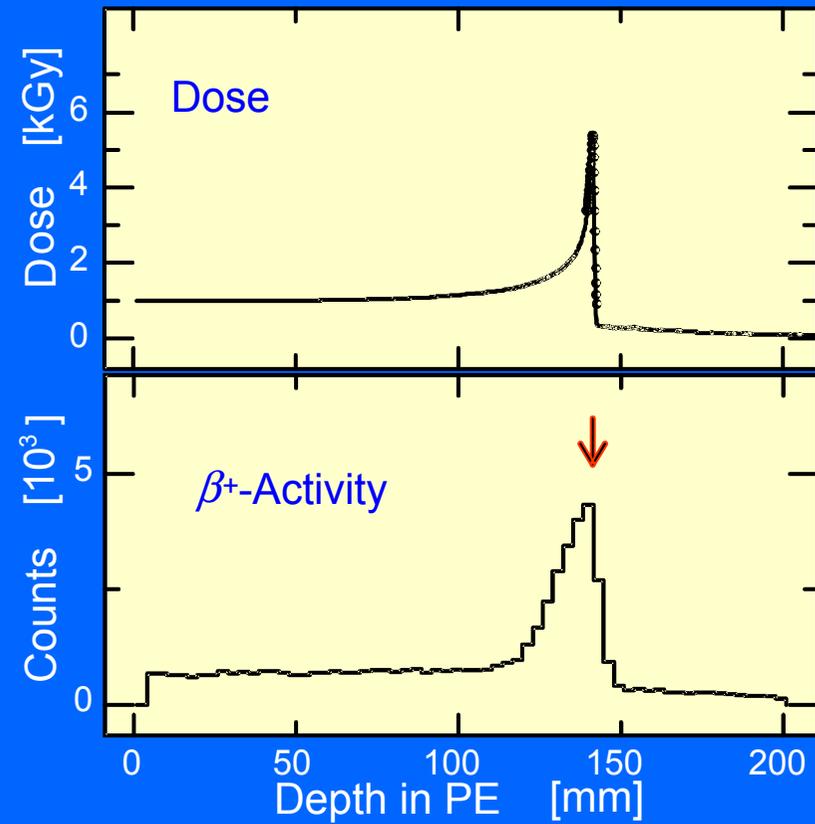
CABOTO
for carbon

(Pictures courtesy of U. Amaldi)

Imaging: quality assurance - In beam PET



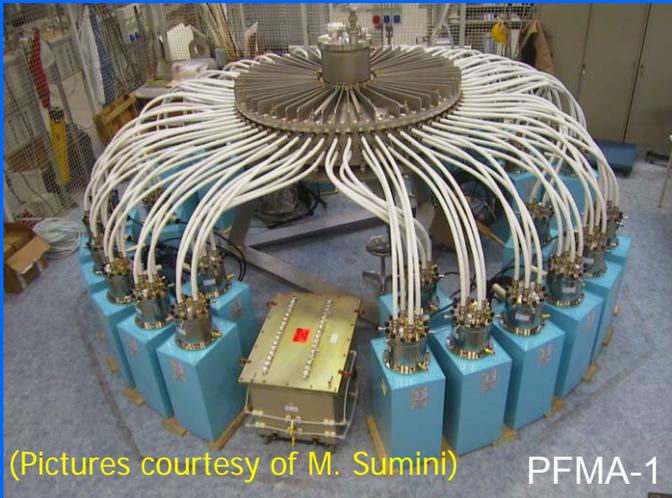
Courtesy of GSI



R&D Topics:

Increase efficiency, solid angle coverage,
new detectors (gas), quantitative
measurements of the dose

Long (?) range perspectives



(Pictures courtesy of M. Sumini) PFMA-1

PFMA-1

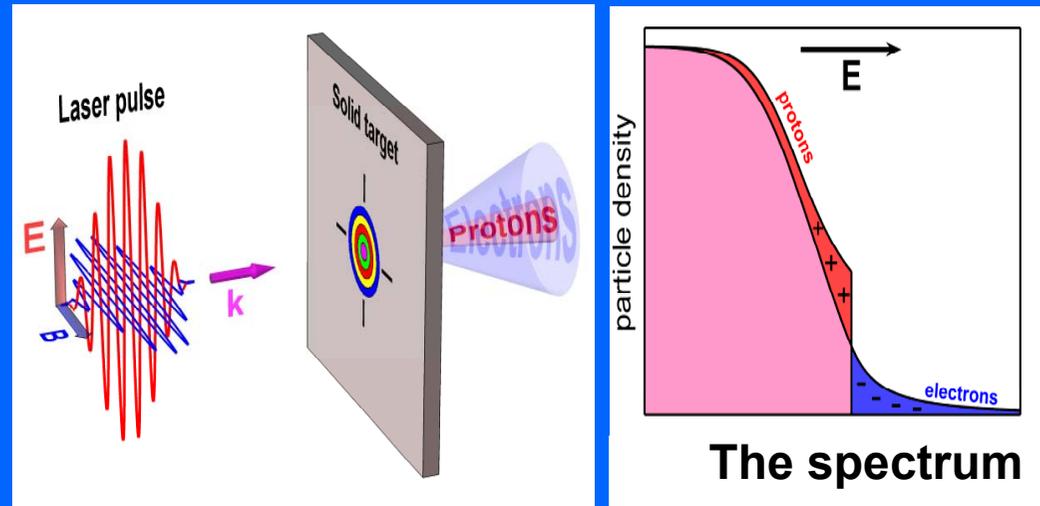
Plasma Focus for Medical Applications

Aim: device for ^{18}F production,

150 kJ (350 μF @ 30 kV)

1 Hz repetition frequency

To breed ~ 1 Ci of F^{18} in 2 hours.



- $\sim 10^{13}$ protons **measured**
- Proton energy: 58 MeV (LLNL)

SIMULATIONS

- Laser: 50 fs, 50 J (Petawatt!)
- $I = 10^{21}$ W/cm²
- $>10^{11}$ protons up to 300 MeV

+ mirrors transport and target close to patients
- broad spectrum, max. energy (C ?), rep. rate



UNIVERSITY
OF BOLOGNA

ABO
PROJECT



azienda Ulss 12 veneziana

UNIVERSITY
OF FERRARA



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Rationale for hadrontherapy

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Wilhelm Conrad Röntgen
1895 discovery of X rays

COLLABORATIONS TO BUILD THE CNAO

CNAO (almost 40 fte personnel) is coordinating the effort of many Institutions

NATIONAL

INFN: co-direction, involvement/responsibility in many technical issues (15), formation

Town of Pavia: land and authorisations

University of Milan: medical coordination and formation

Polytechnic of Milan: patient positioning, radioprotection and authorisations

University of Pavia: electrical plants, special power supplies and betatron, safety, formation

Province of Pavia: logistics and authorisation

INTERNATIONAL

CERN: special magnets, dipole measurements and diagnostics (+ PIMMS heritage)

GSI: linac and special components

LPSC: optics, betatron, low-level RF, control system

Overall: about 65 fte equivalent working for CNAO

82 firms working for CNAO

Hospital Building
Completion May 2007

High Technology Bld.
Delivery - October 2006

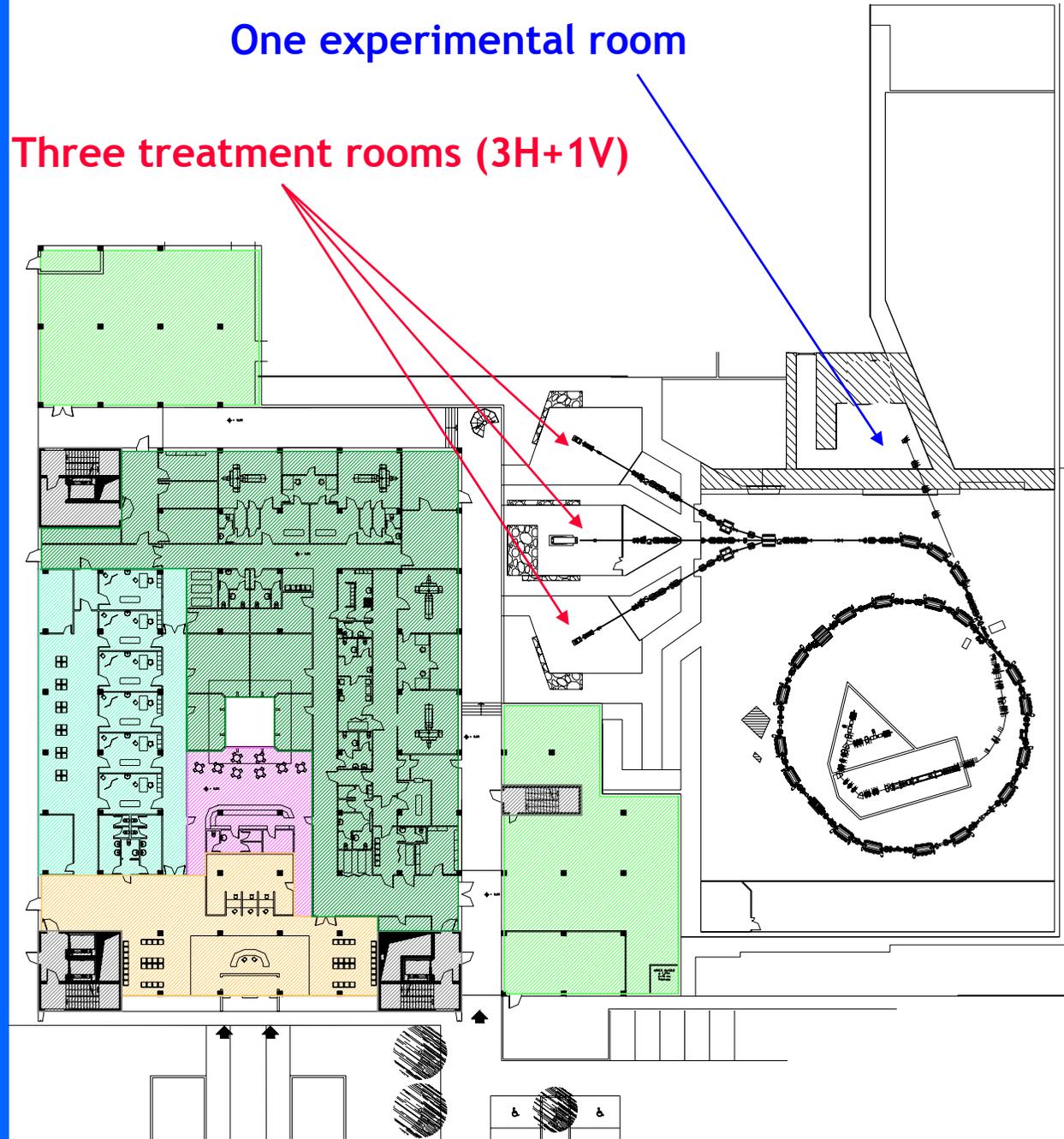


In construction in Pave
30 km south-west of Milan
northern part of Italy

CNAO

One experimental room

Three treatment rooms (3H+1V)



LEGENDA

AREE FUNZIONALI

- Ambulatori mq 453
- Area immagini mq 1146
- Ingresso accettazione mq 337
- Bar mq 144
- Centrali tecnologiche mq 736
- Collegamenti verticali mq 227

CNAO SITE
May 17th, 2006



CNAO
May 17th, 2006



CNAO
June 27th, 2006



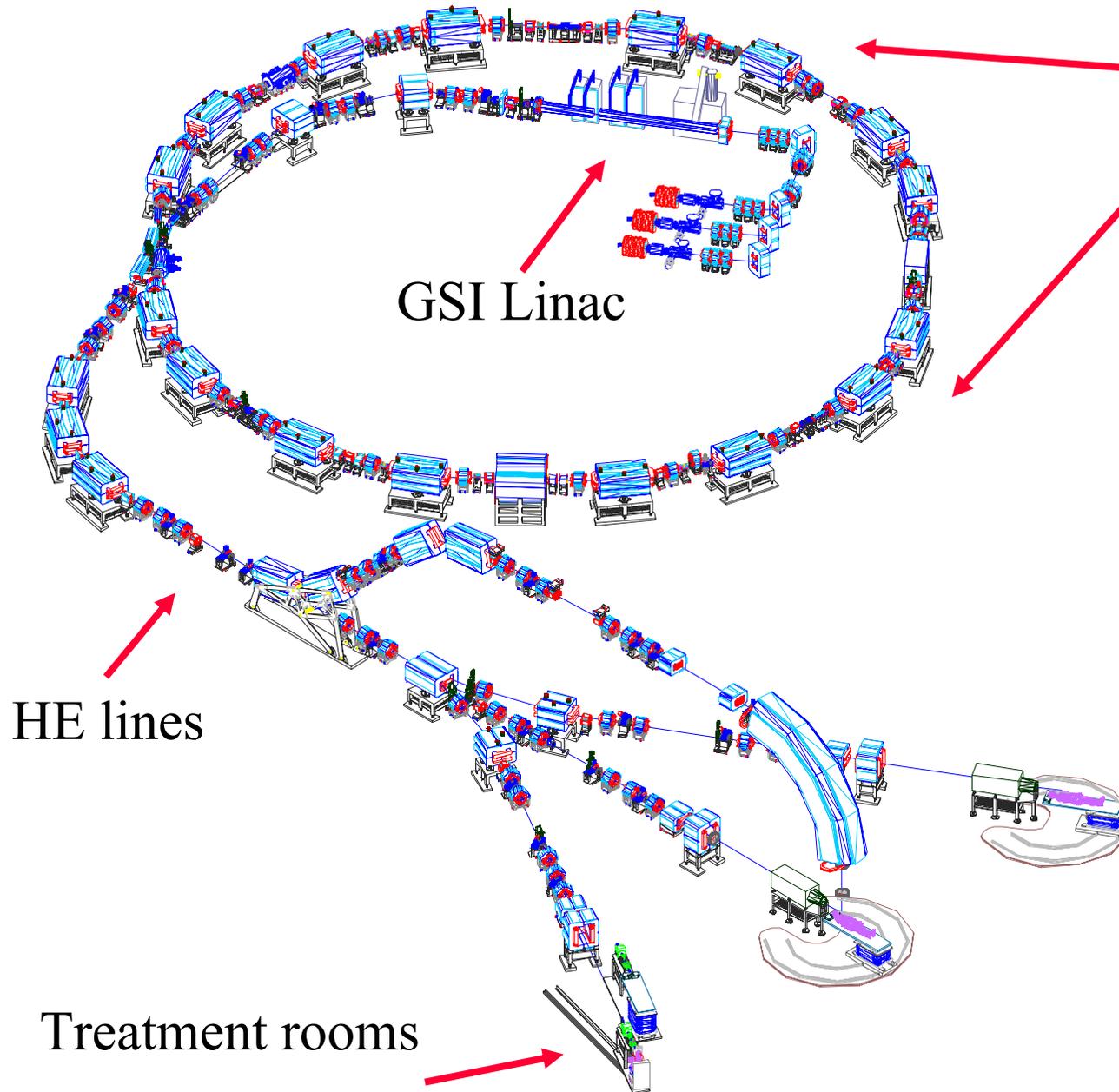
The heart of CNAO

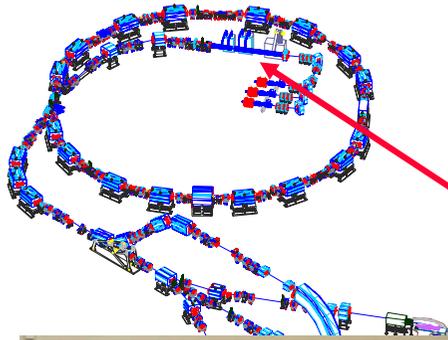
SYNCHROTRON

OPTIMIZED
for an hospital based
facility (all Ion-therapy
centres existing in the
World adopt it):

- Safety
- Efficiency
- Reliability
- Maintainability

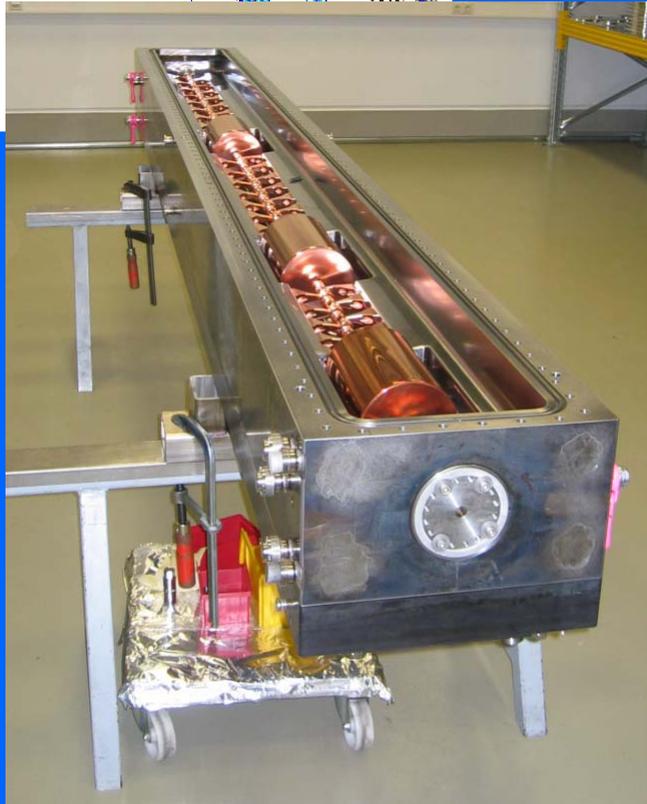
**MODULAR and
UPGRADABLE system:**
Up-to-date for
20-30 years





CNAO Tour - RFQ + IH-LINAC

Just after the RFQ there is a second linear accelerator, the IH-linac, that increases the energy till 7 MeV/u



Contract signed
15th July 2004

Sub-contracts with
firms:

Thales
Pink
Danfysik
Sigmaphy
Jaeger
NTG
Eckelmann

In collaboration with:



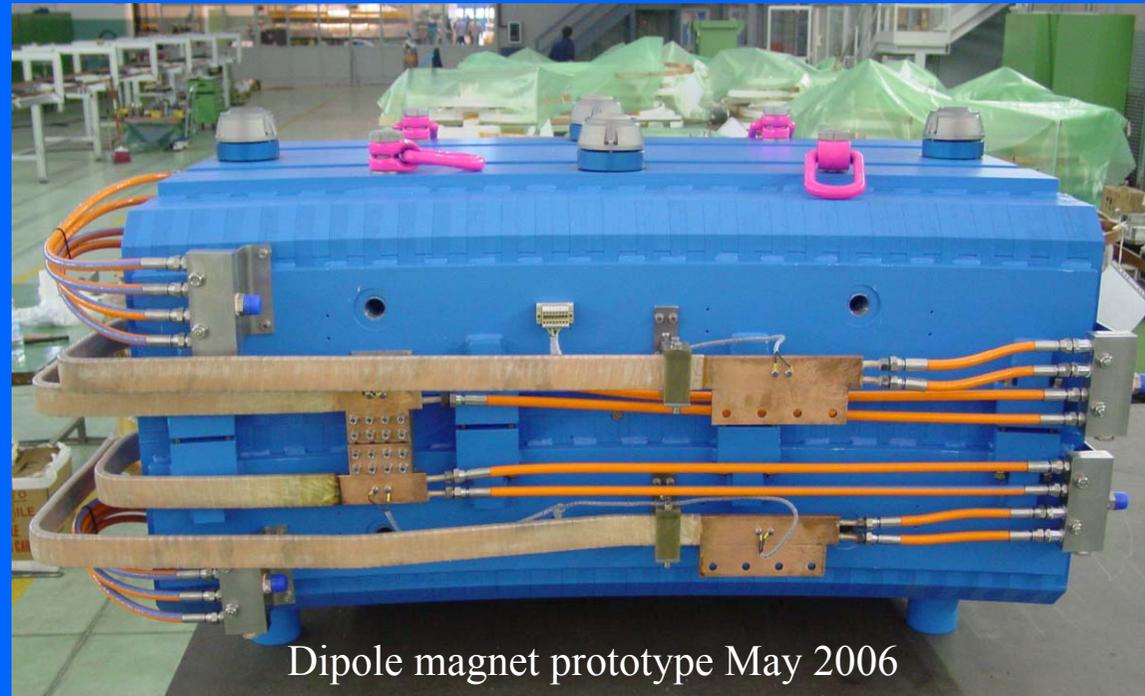
RFQ completed and presently at GSI – IH: mechanics completed, copperlating at GSI
Installation in Pavia: Feb – April 2007

CNAO Tour - Conventional Magnets

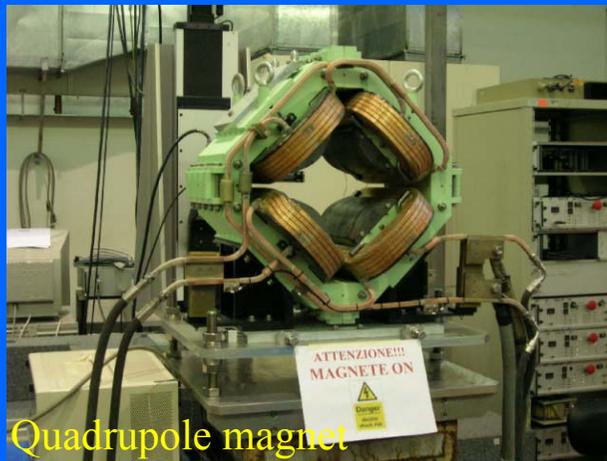
For a total of 180 magnets:
N. 32 Bending Dipoles
N. 85 Quadrupoles
N. 7 Sextupoles
N. 56 Correctors

**ANSALDO ASG
SIGMAPHI SA**

End of production Spring 07



Dipole magnet prototype May 2006

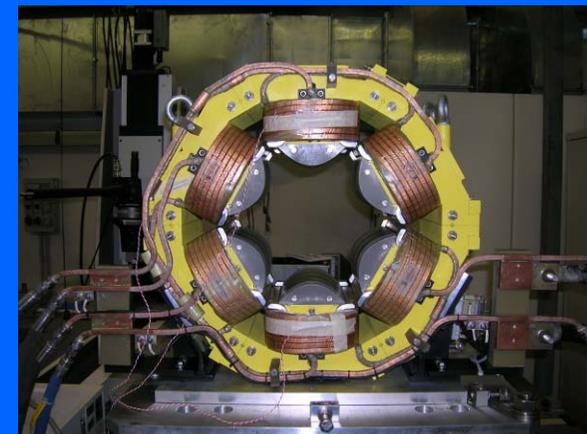


Quadrupole magnet
Prototype November 2005

In collaboration with:

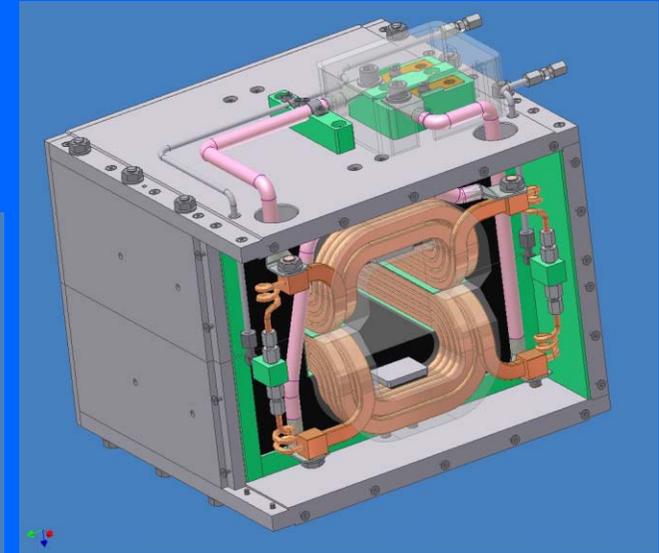
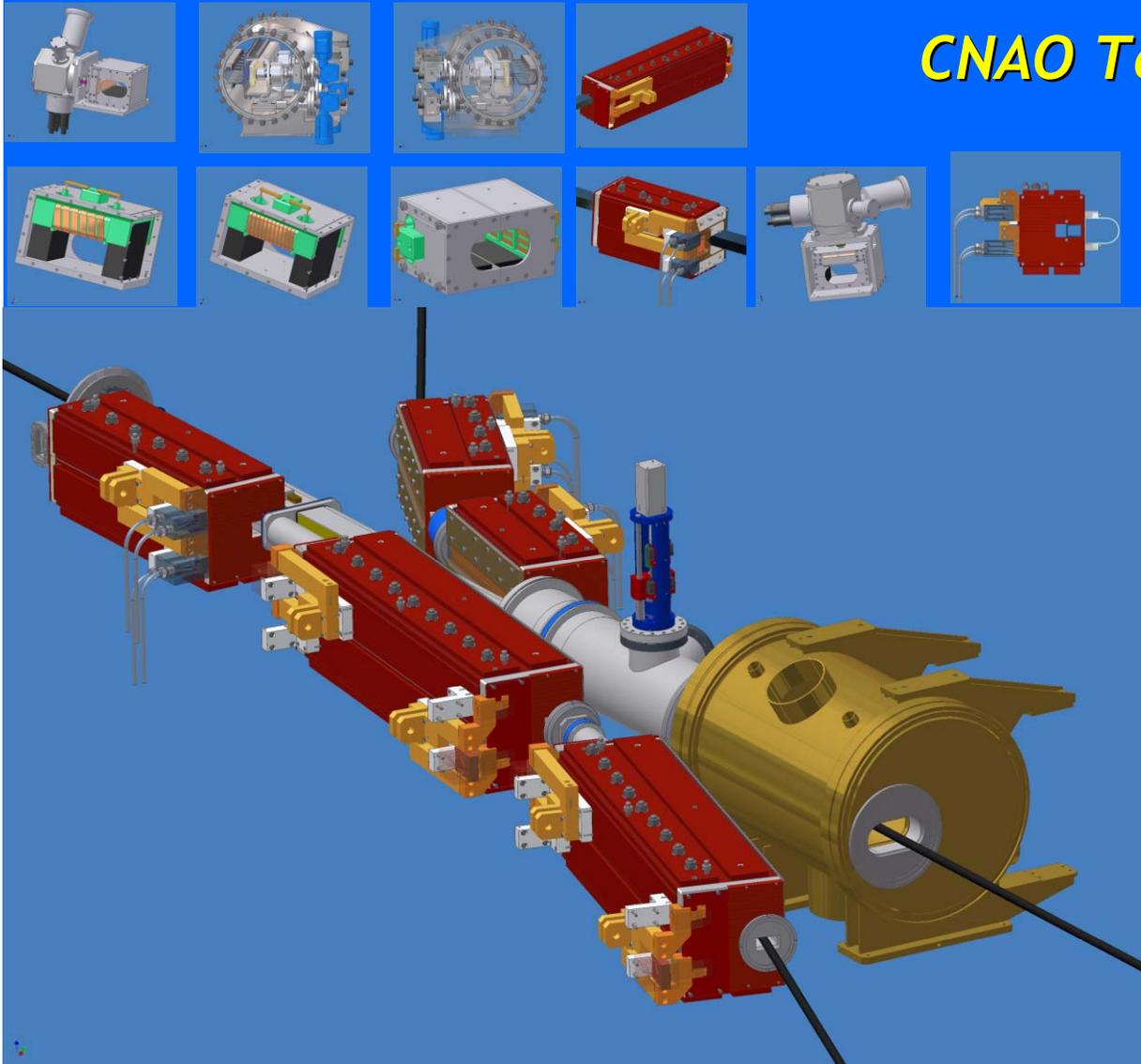


Corrector magnets
Production completed
October 2005.



Sextupole magnet prototype
December 2005

CNAO Tour - Special Magnets



- 1 Dump Bumper magnet
- 1 Dump Bumper magnet
- 2 Injection Bumper magnets
- 4 Chopper dipoles
- 1 Horizontal Tune kicker
- 1 Vertical Tune kicker
- 2 Injection Septa
- 1 Thin Extraction Septum
- 2 Thick Extraction Septa
- 1 Electrostatic Extraction Septum
- 1 Electrostatic Injection Septum

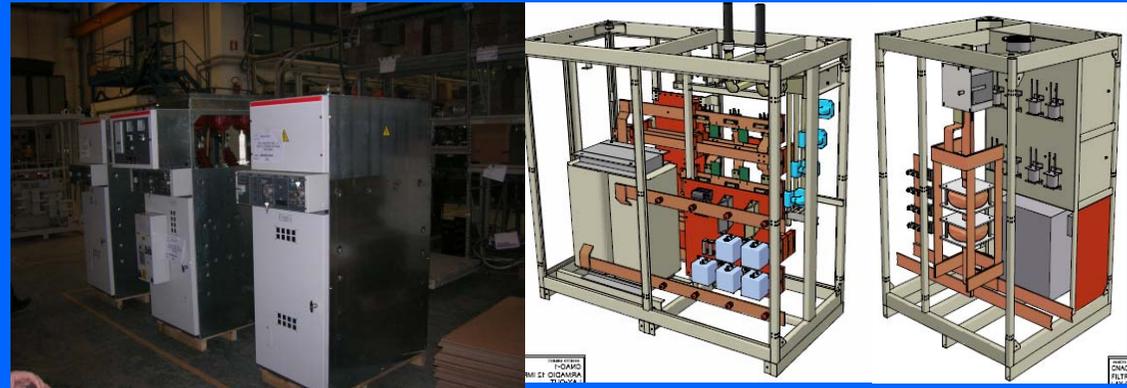
Under construction by Danfysik
End of production October 2006

In collaboration with:
CERN
UniPv

CNAO Tour - Power Supplies

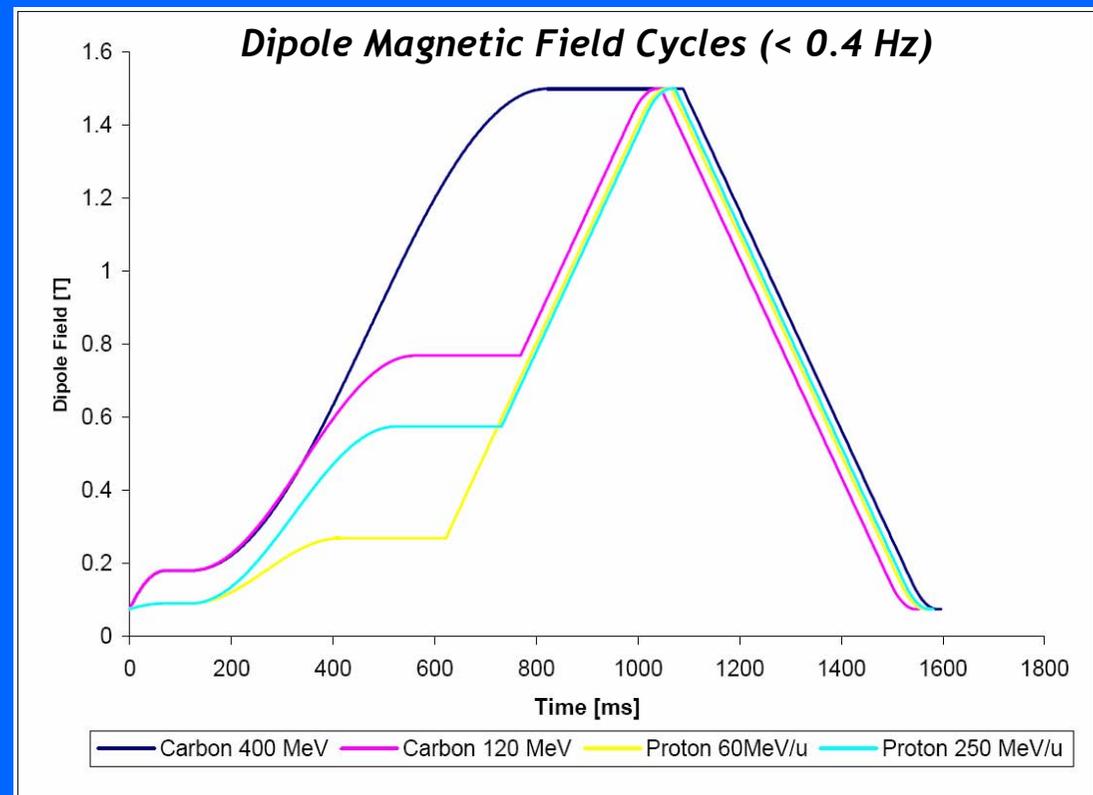
Synchrotron Dipole Power Supply

Current range 30 to 3000 A
Power range 0.9 to 5000 kW
Current stability $\pm 5 \cdot 10^{-6}$ to $\pm 5 \cdot 10^{-4}$
Current reproducibility $\pm 2.5 \cdot 10^{-6}$ to $\pm 2.5 \cdot 10^{-4}$



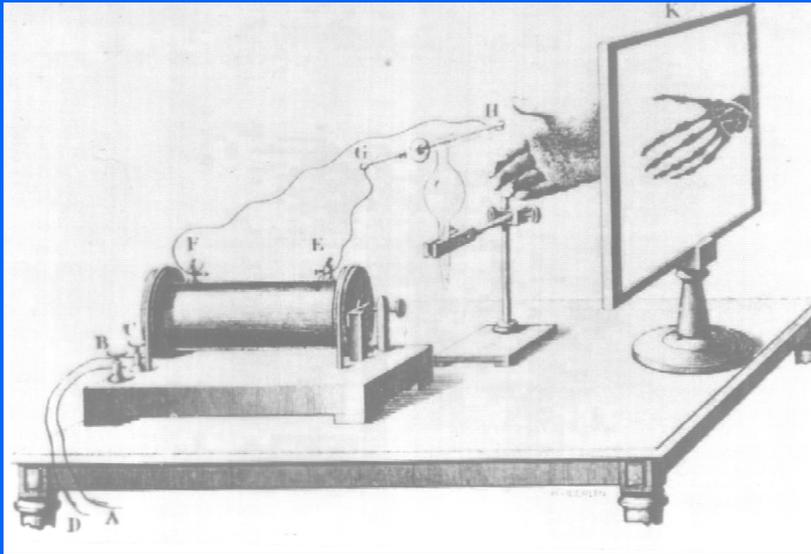
Power Supplies Total n. 187
N. 177 under construction
(OCEM - EEI)
N. 10 still to be ordered
Last delivery March 2007

In collaboration with:



Conclusion

Discovery of X-rays
1895



First patient
at HIT and CNAO
fall 2007 - start 2008

... a long way through
hadrontherapy is a further step,
with good results and
very promising developments,
in radiotherapy evolution

