

# ORIGINI E SVILUPPI DELL'ADROTERAPIA (\*) IN ITALIA

Ugo Amaldi

(\*) Termine collettivo che copre tutte le terapie basate su particelle fatte di quark:  
neutroni veloci, BNCT, pioni, protoni, ioni leggeri (C), alfa per alfa-immunotherapy, antiprotoni, ...

G. Tosi:

1964 in "Energia Nucleare" , CISE  
"Gli acceleratori di particelle in radioterapia"  
betatrons for electrons and synchrotrons for protons.

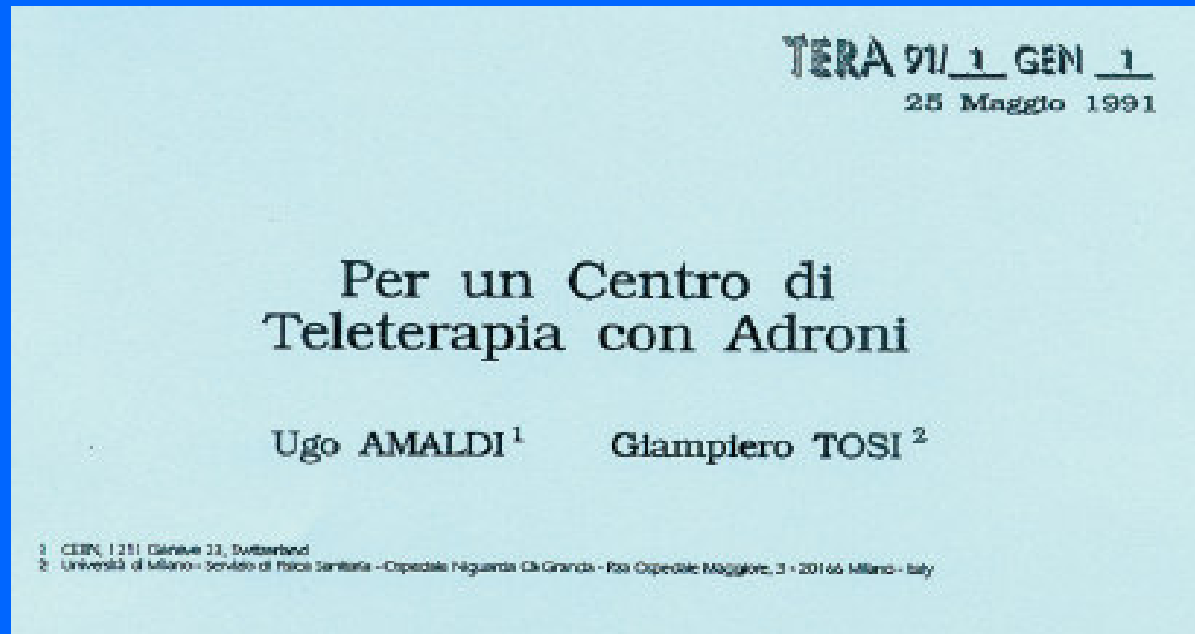
E. Fiorini:

1974 in "Giornale di Fis. San. e Prot. delle Radiazioni"  
"Nuovi mezzi strumentali per le applicazioni mediche delle particelle  
ad altissima energia"



## *My view of the Italian developments*

Discussions in the years 1988-1990 between G. Tosi and U.A. while meeting at Via Celoria when giving the courses to the students of **Scuola di Specializzazione in Fisica Sanitaria e Ospedaliera**



- September 1991      INFN finances ATER at INFN-Milano to design the ion accelerator
- October      1992      Creation of TERA foundation (U.A., E. Borgonovi, G. Tosi, G. Vanolo)
- 1992- 2000      INFN finances extensions of ATER up to 12 groups in 12 Sections

# The "Blu Book" and the "Green Book"

## THE TERA PROJECT AND THE CENTRE FOR ONCOLOGICAL HADRONTHERAPY

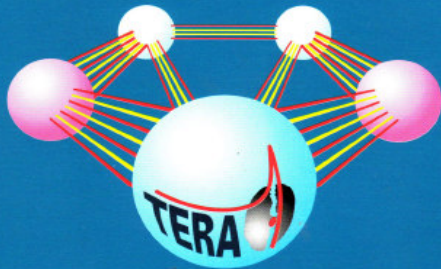
IL PROGETTO TERA E  
IL CENTRO DI ADROTERAPIA ONCOLOGICA

Vol. I

Second Edition

THE TERA COLLABORATION

U. AMALDI and M. SILARI editors



### PROGETTO ADROTERAPIA

INFN - ISTITUTO NAZIONALE DI FISICA NUCLEARE  
AIFB - ASSOCIAZIONE ITALIANA DI FISICA BIOMEDICA  
AIRO - ASSOCIAZIONE ITALIANA DI RADIOTERAPIA ONCOLOGICA  
CERN - EUROPEAN LABORATORY FOR PARTICLE PHYSICS  
CNR - CONSIGLIO NAZIONALE DELLE RICERCHE  
ENEA - ENTE PER LE NUOVE TECNOLOGIE, L'ENERGIA E L'AMBIENTE  
ISS - ISTITUTO SUPERIORE DI SANITÀ  
SIIRR - SOCIETÀ ITALIANA PER LE RICERCHE SULLE RADIAZIONI  
TERA - FONDAZIONE PER ADROTERAPIA ONCOLOGICA

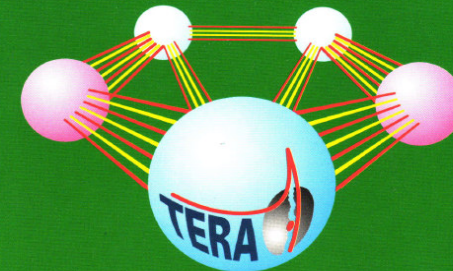
1995

## THE RITA NETWORK AND THE DESIGN OF COMPACT PROTON ACCELERATORS

LA RETE ITALIANA TRATTAMENTI ADROTERAPICI E  
IL PROGETTO DI ACCELERATORI COMPATTI DI PROTONI

THE TERA COLLABORATION

U. AMALDI, M. GRANDOLFO and L. PICARDI editors



### PROGRAMMA ADROTERAPIA

INFN - ISTITUTO NAZIONALE DI FISICA NUCLEARE  
AIFB - ASSOCIAZIONE ITALIANA DI FISICA BIOMEDICA  
AIFS - ASSOCIAZIONE ITALIANA FISICA SANITARIA  
AIRB - ASSOCIAZIONE ITALIANA DI RADIOBIOLOGIA  
AIRO - ASSOCIAZIONE ITALIANA DI RADIOTERAPIA ONCOLOGICA  
CERN - EUROPEAN LABORATORY FOR PARTICLE PHYSICS  
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TERA - FONDAZIONE PER ADROTERAPIA ONCOLOGICA

1996

# The "Red Book" and the "White Book"

## IL CENTRO NAZIONALE DI ADROTERAPIA ONCOLOGICA A MIRASOLE THE NATIONAL CENTRE FOR ONCOLOGICAL HADRON THERAPY AT MIRASOLE

FONDAZIONE TERA

A cura di Ugo Amaldi



### PROGRAMMA ADROTERAPIA

INFN - ISTITUTO NAZIONALE DI FISICA NUCLEARE  
AIFB - ASSOCIAZIONE ITALIANA DI FISICA BIOMEDICA  
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TERA - FONDAZIONE PER ADROTERAPIA ONCOLOGICA

1997

## THE PATH TO THE ITALIAN NATIONAL CENTRE FOR ION THERAPY

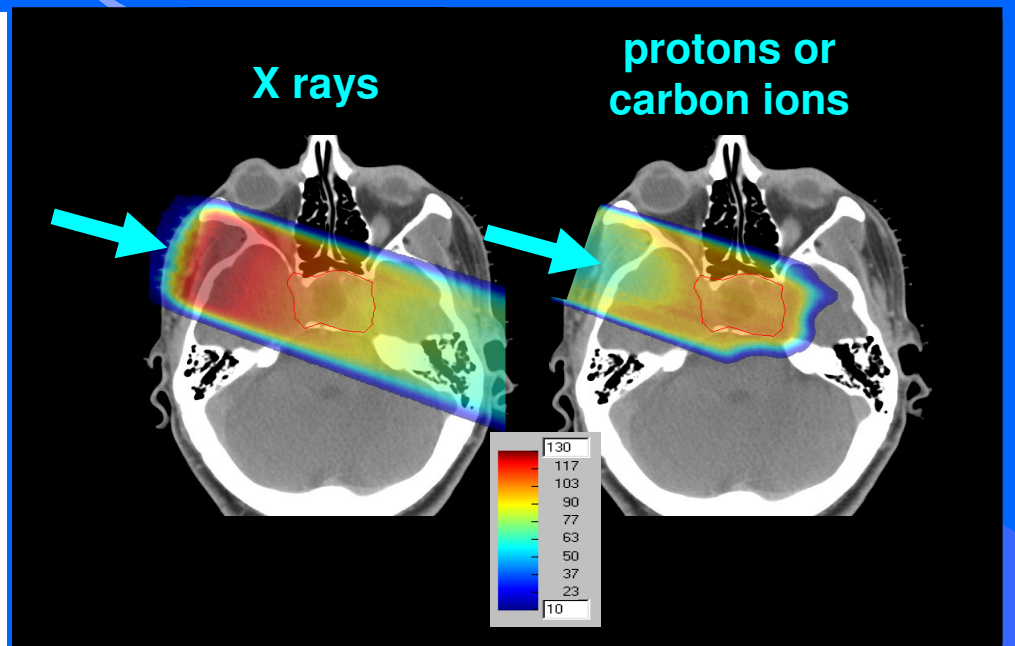
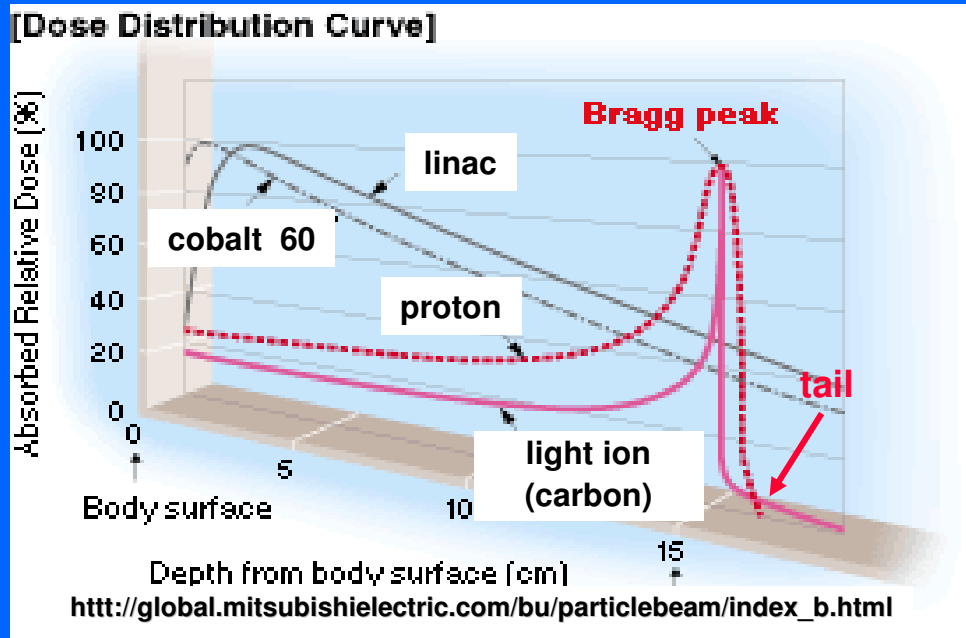
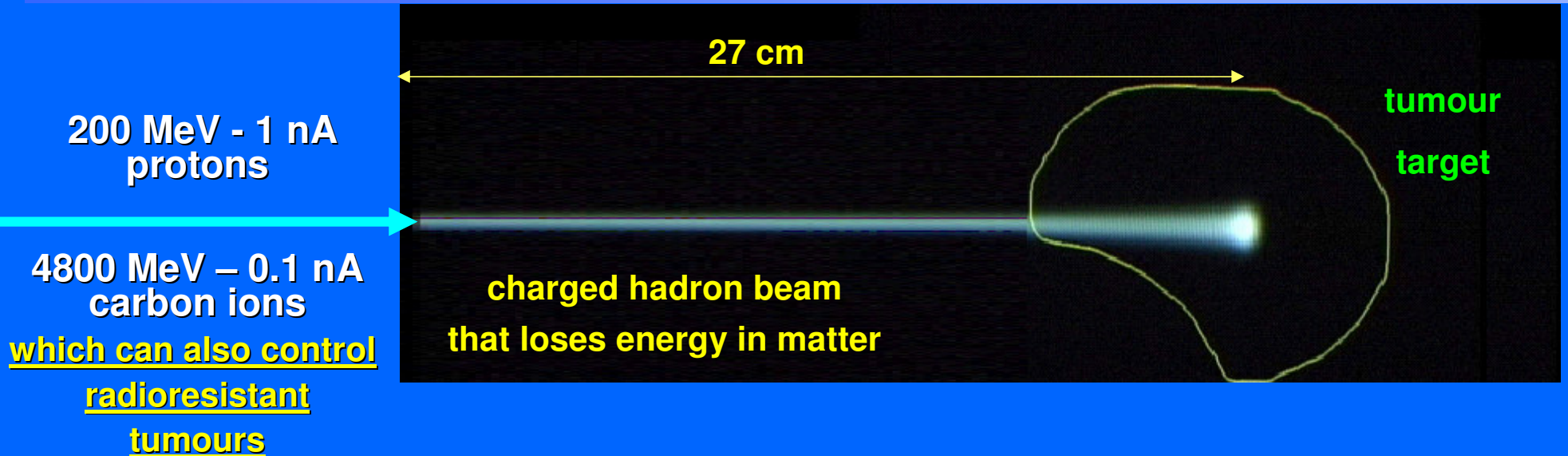
FONDAZIONE TERA

UGO AMALDI and GIULIO MAGRIN editors



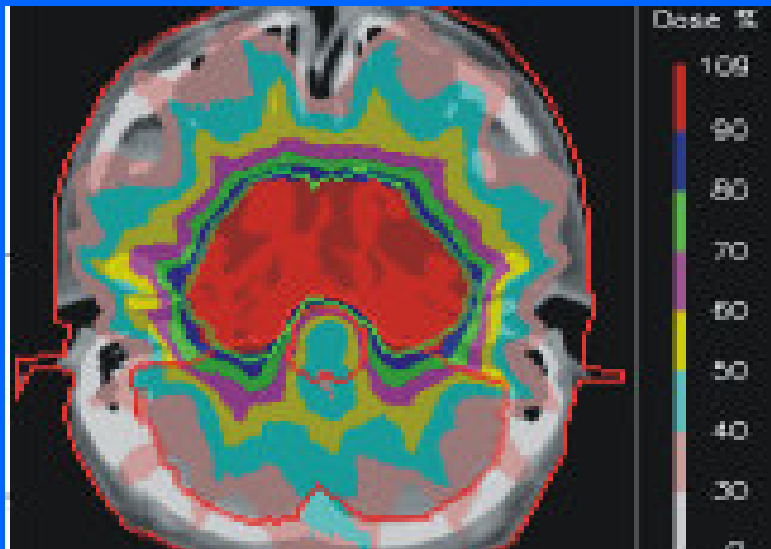
2005

# Basic facts: protons and ions spare healthy tissues

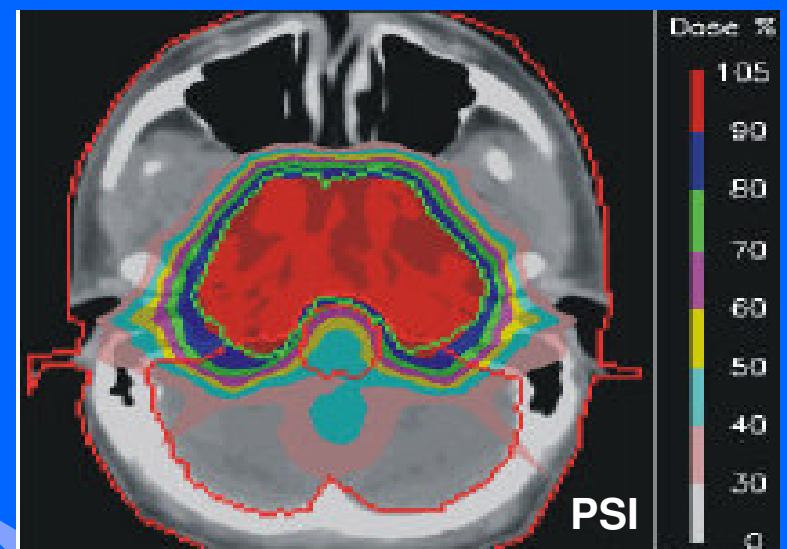


# Protons are quantitatively different from X-rays

## 9 X-ray fields

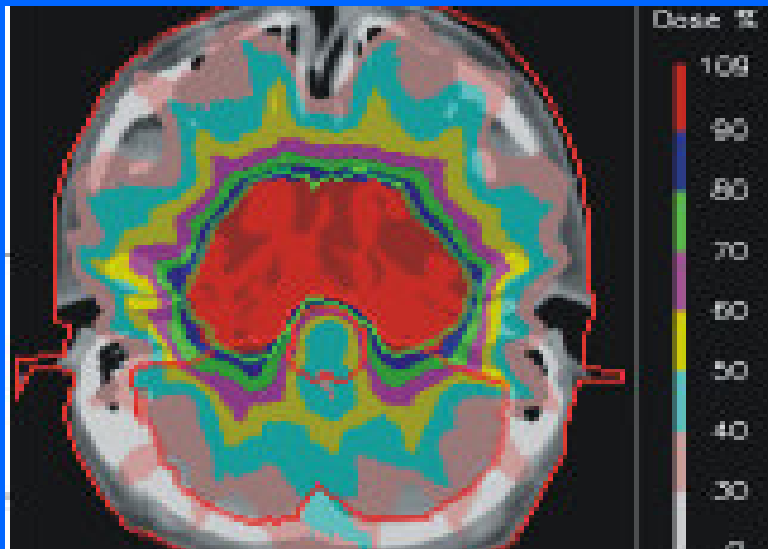


## 4 proton fields

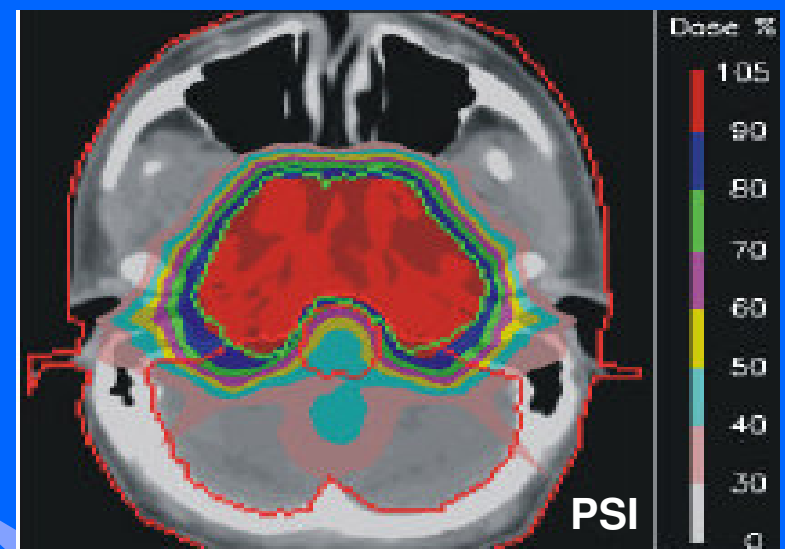


# ***Carbon ions are qualitatively different from X-rays***

**9 X-ray fields**



**4 proton fields**



**Carbon ions deposit in a cell 24 times more energy than a proton producing not reparable multiple close-by double strand breaks**

**Carbon ions can control radio-resistant tumours**

# The site treated with hadrons

Today in the world

protontherapy:  
50'000 patients

carbon ion  
therapy  
3500 patients

**BUT**

In 1991 about  
15'000 patients had  
been treated with p  
and none with C  
ions

## Eye and Orbit

- Choroidal Melanoma
- Retinoblastoma
- Choroidal Metastases
- Orbital Rhabdomyosarcoma
- Lacrimal Gland Carcinoma
- Choroidal Hemangiomas

## Head and Neck Tumors

- Locally Advanced Oropharynx
- Locally Advanced Nasopharynx
- Soft Tissue Sarcoma  
Recurrent or Unresectable
- Misc. Unresectable or Recurrent Carcinomas

## Chest

- Non Small Cell Lung Carcinoma  
Early Stage—Medically Inoperable
- Paraspinal Tumors  
Soft Tissue Sarcomas, Low Grade  
Chondrosarcomas, Chordomas

## Abdomen

- Paraspinal Tumors
- Soft Tissue  
Sarcomas,  
Low Grade  
Chondrosarcomas,  
Chordomas

## Pelvis

- Early Stage Prostate Carcinoma
- Locally Advanced Prostate Carcinoma
- Locally Advanced Cervix Carcinoma
- Sacral Chordoma
- Recurrent or Unresectable  
Rectal Carcinoma
- Recurrent or Unresectable  
Pelvic Masses

## Central Nervous System

- Adult Low Grade Gliomas
- Pediatric Gliomas
- Acoustic Neuroma  
Recurrent or Unresectable
- Pituitary Adenoma  
Recurrent or Unresectable
- Meningioma  
Recurrent or Unresectable
- Cranio-pharyngioma
- Chordomas and  
Low Grade Chondrosarcoma  
Clivus and Cervical Spine
- Brain Metastases
- Optic Glioma
- Arteriovenous Malformations

## ***Numbers of potential patients (2004)***

### X-ray therapy ( $\geq 40$ linacs)

every 10 million inhabitants: 20'000 pts/year

### Protontherapy

12% of X-ray patients 2'400 pts/year

### Therapy with Carbon ions for radio-resistant tumour

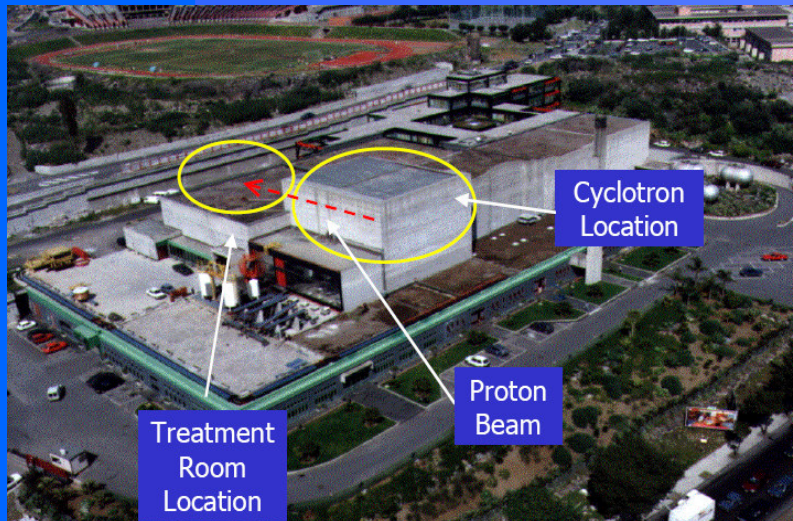
3% of X-ray patients 600 pts/year

TOTAL every 10 M about 3'000 pts/year (\*)

(\*) In Italy : 17'000 patients  
5 proton centres and 1 carbon centre (AIRO-2004)

## *Italian activities*

# Càtana at “Laboratori Nazionali del Sud” of INFN is described by. G. Battistoni



LNS Accelerator Layout

## ***15 years of TERA: 1992-2007***

**TERA has proposed and produced 3 designs for the National Centre for carbon ions (and p) to be built on 3 sites: Novara (1993-1995), Milano (1996-2000) and Pavia**



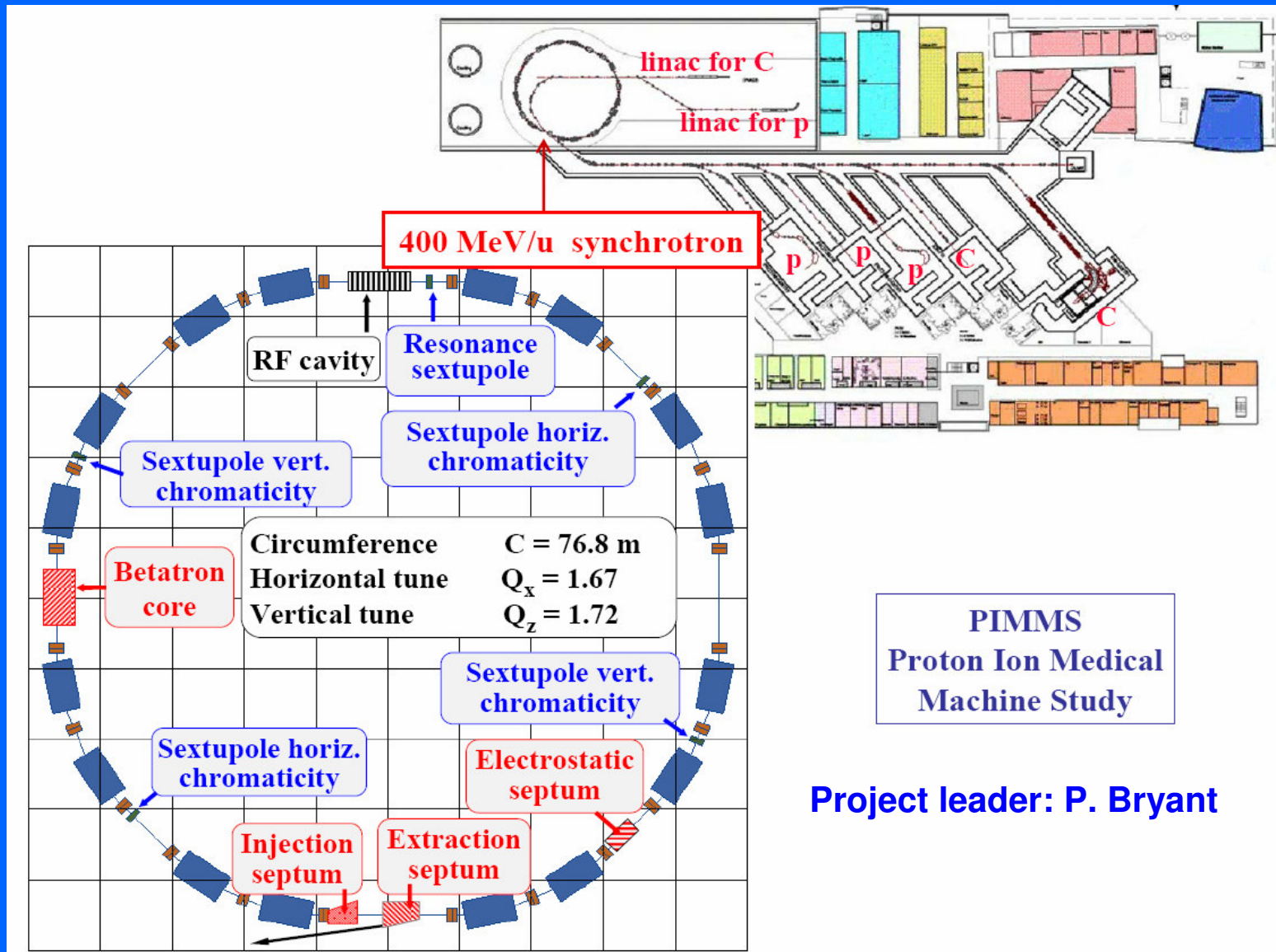
**1. CNAO is being completed in Pavia by the CNAO Foundation together with INFN**

**TERA has developed (1993-2006) a novel type of accelerator: the “cyclinac” for protons and carbon ions**



**2. IDRA (2001)**  
**3. CABOTO (2005)**

# Basic PIMMS design: CERN-TERA- MedAustron 1996-2000



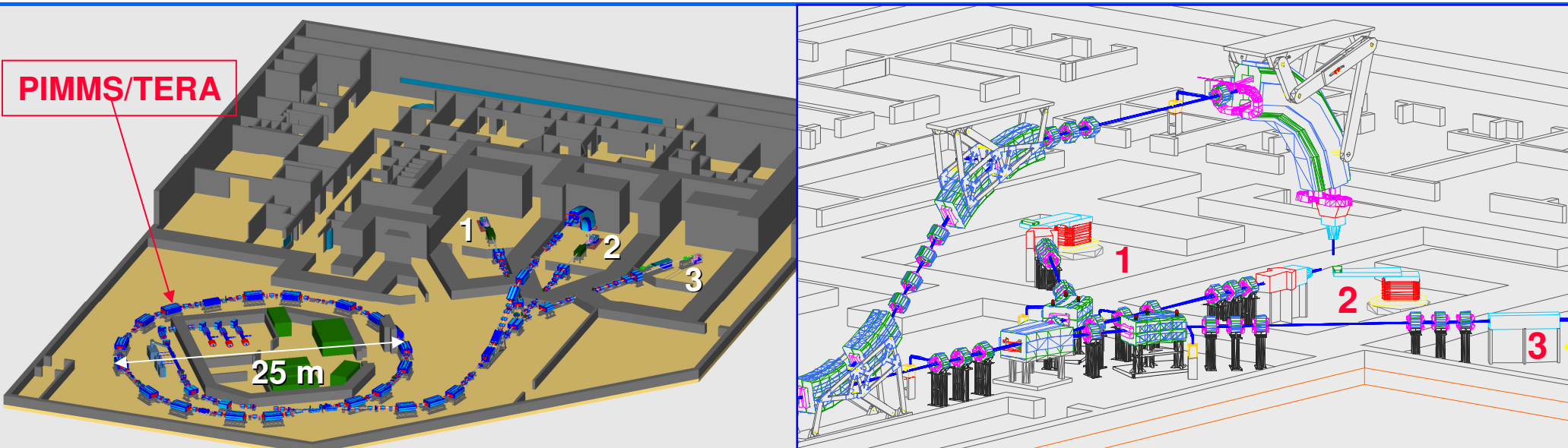
# **CNAO = Centro Nazionale di Adroterapia**

CNAO Foundation created by the Italian Government in 2001:  
“to construct and manage  
the ‘Centro Nazionale di Adroterapia Oncologica’ designed by TERA”

**Founders: Osp. Maggiore, San Matteo, INT, IEO, Besta and TERA**

Since 2004 INFN is Institutional Participant with construction  
responsibilities as described by G. Battistoni

**Modifications to PIMMS: multi-turn injection, linac inside the ring, short beam lines**



## *The surface buildings*

**Project: Calvi –TEKNE**

**In October 2003 TERA  
passed to CNAO  
specifications +  
drawings (3000 pages)  
and 25 people**

**Main source of 90 M€:  
Italian Health Ministry**

**First beam:  
spring 2008**



**CNAO Foundation**

**President:**

**E. Borloni**

**Medical Director:**

**R. Orecchia**

**Technical Director:**

**S. Rossi**

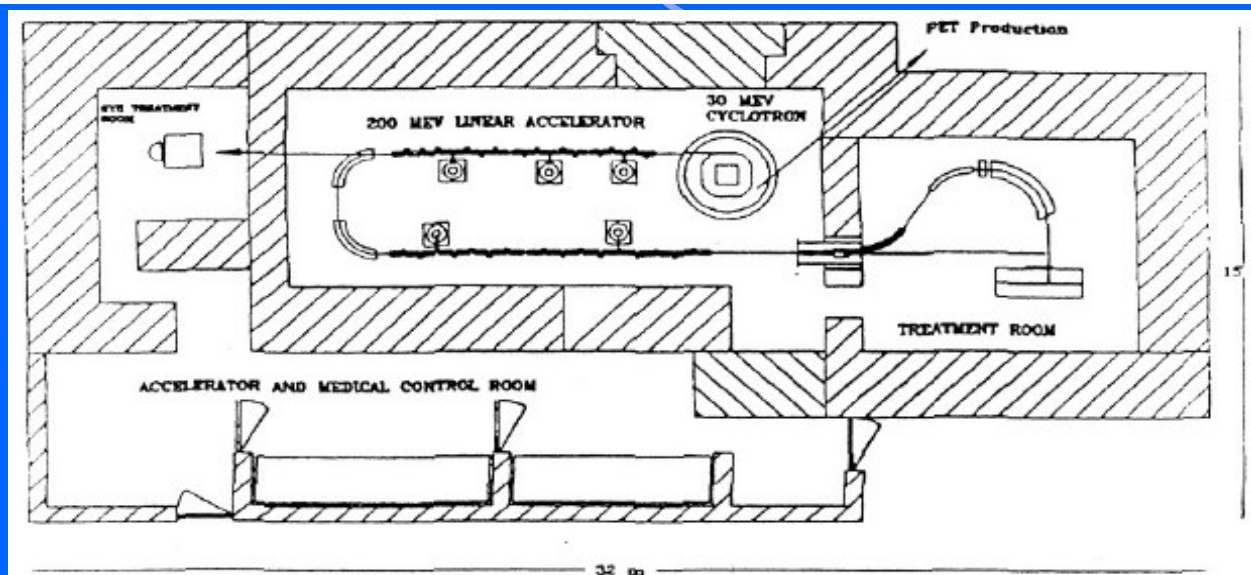
## *Linacs in hadrontherapy*

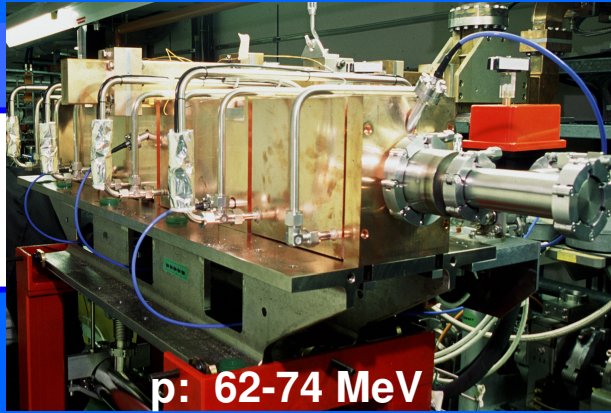
## ***The first proposal of the “cyclinac” accelerator: 1993***

**UA in “Hadrontherapy in Oncology”, UA and B. Larsson eds., Elsevier, 1994**

A preliminary solution foresees the use of a 30 MeV cyclotron as an injector, with other options under study.

**TERA 94/13 GEN 11: “Il progetto adroterapia tre anni dopo” 25.5.1994**





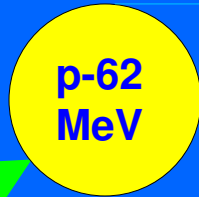
LIBO prototype  
CERN-INFN-TERA  
1998-2002

p: 62-74 MeV

## 2 routes to light ion therapy

TERA "cyclinacs" for p and C

Green  
Book  
1996

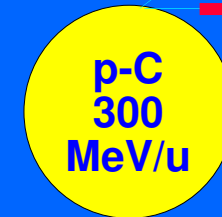


LIBO



LIBO

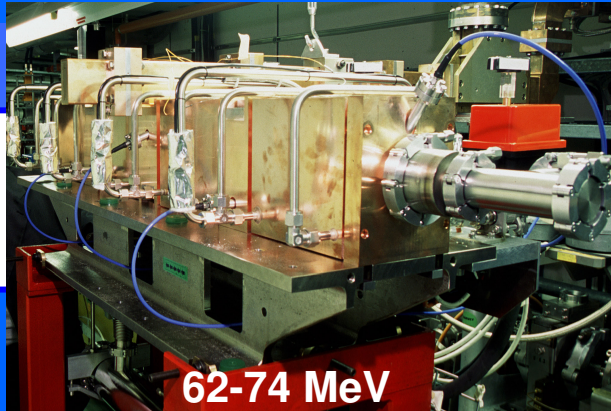
230 MeV



LIBO

435 MeV/u

## 2 routes to light ion therapy



LIBO prototype  
CERN-INFN-TERA  
1998-2002

62-74 MeV

TERA "cyclinacs" for p and C

Green  
Book  
1996

p-62  
MeV

LIBO

p-24  
MeV

LIBO

230 MeV

p-C  
300  
MeV/u

LIBO

435 MeV/u

SCDTL

Linac: 7 MeV

62 MeV

LIBO

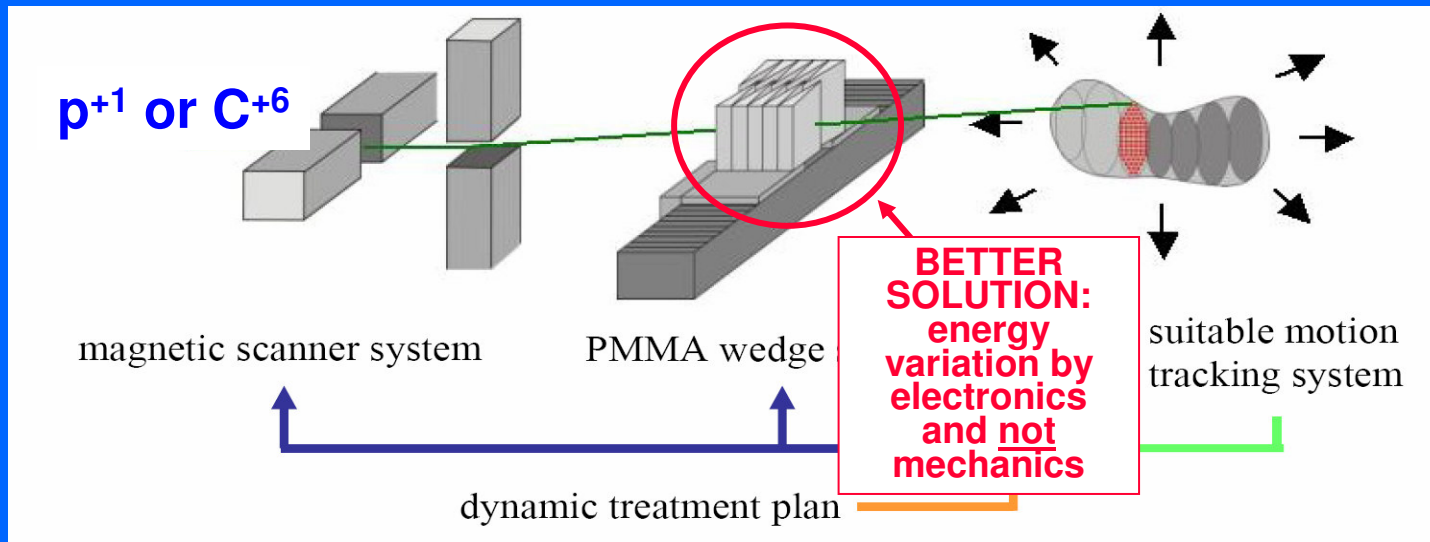
TOP = 'Terapia Oncologica con Protoni'  
ENEA and Istituto Superiore di Sanità

SCDTL prototype  
G. Picardi et al  
2000-2005



7-11 MeV

# WHY? The challenge of hadrontherapy: treat moving organs



Sven O. Grözinger, GSI Darmstadt

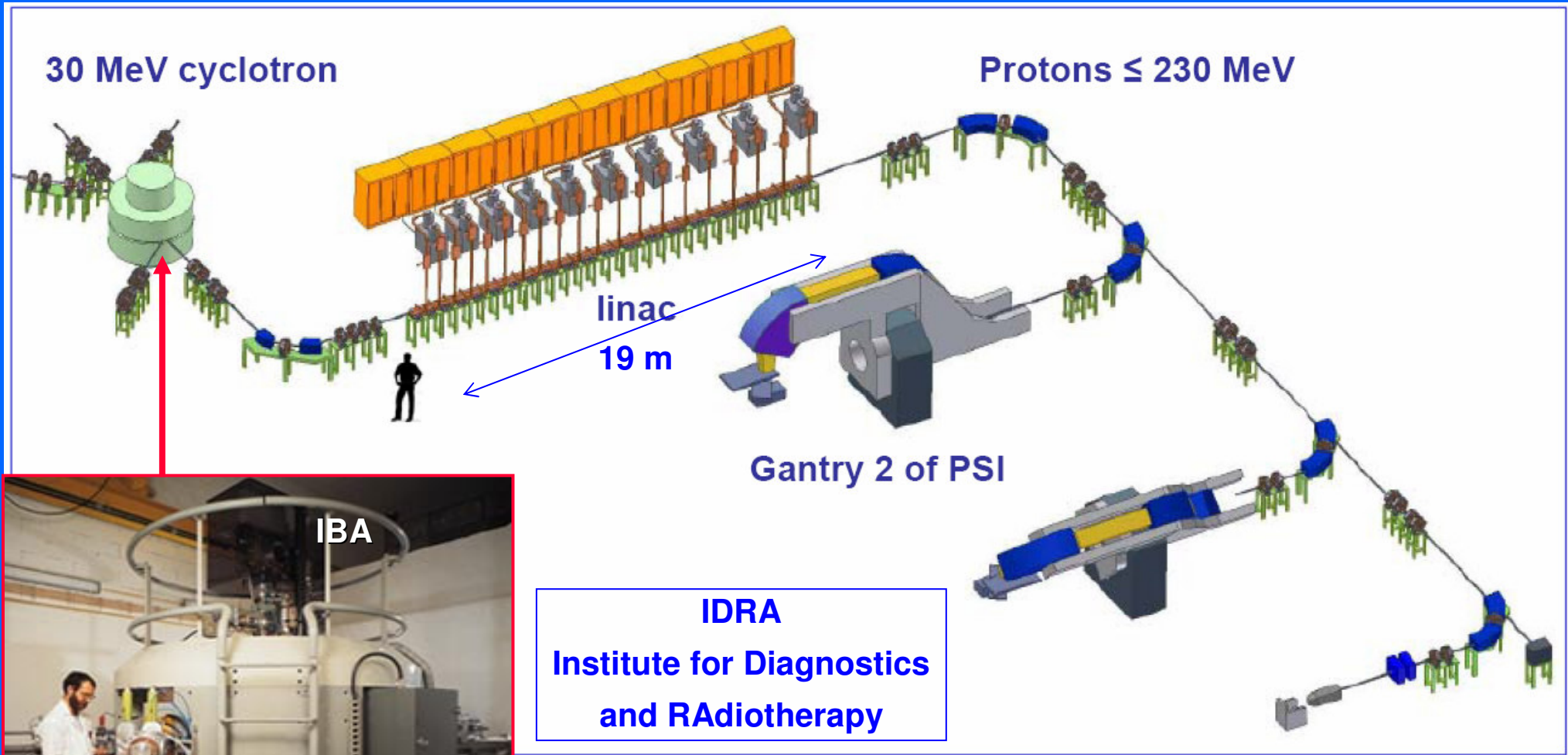
static

moving,  
non-compensated

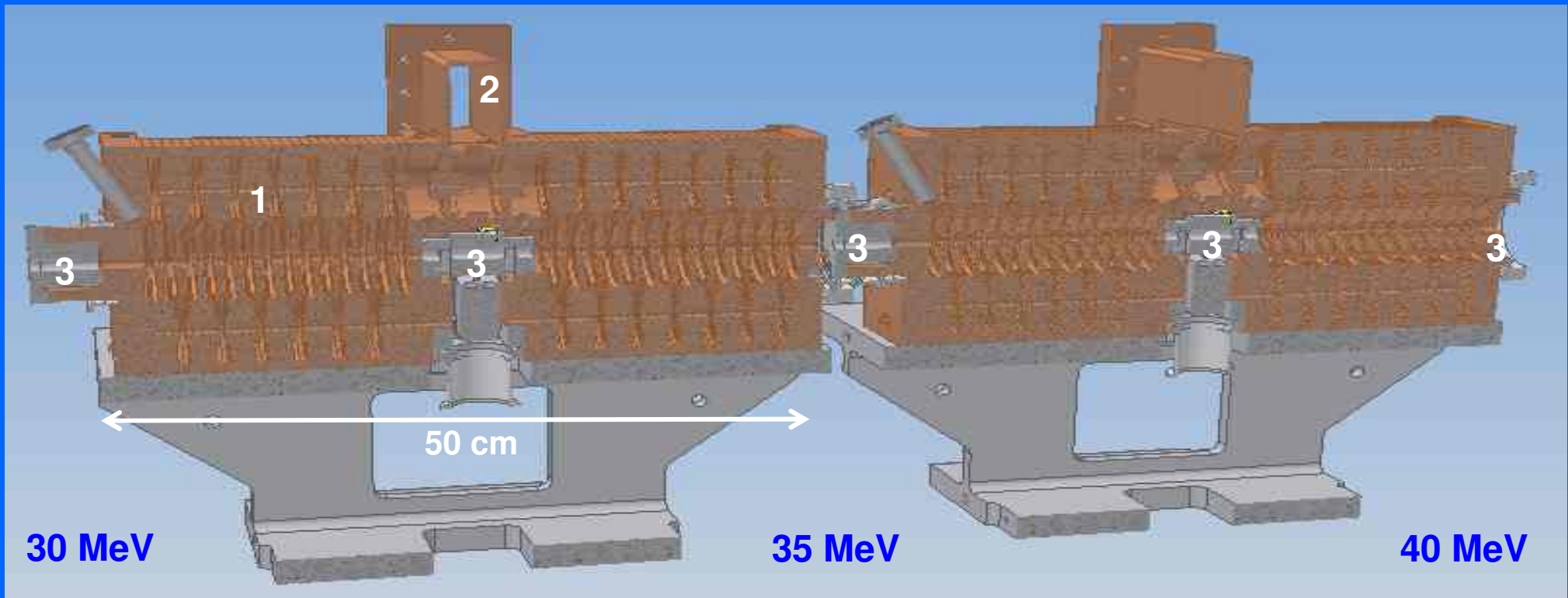
moving,  
compensated

**PSI and GSI approach : depth adjusted by moving absorbers**

# The proton 'cyclinac' concept faces the challenge : IDRA



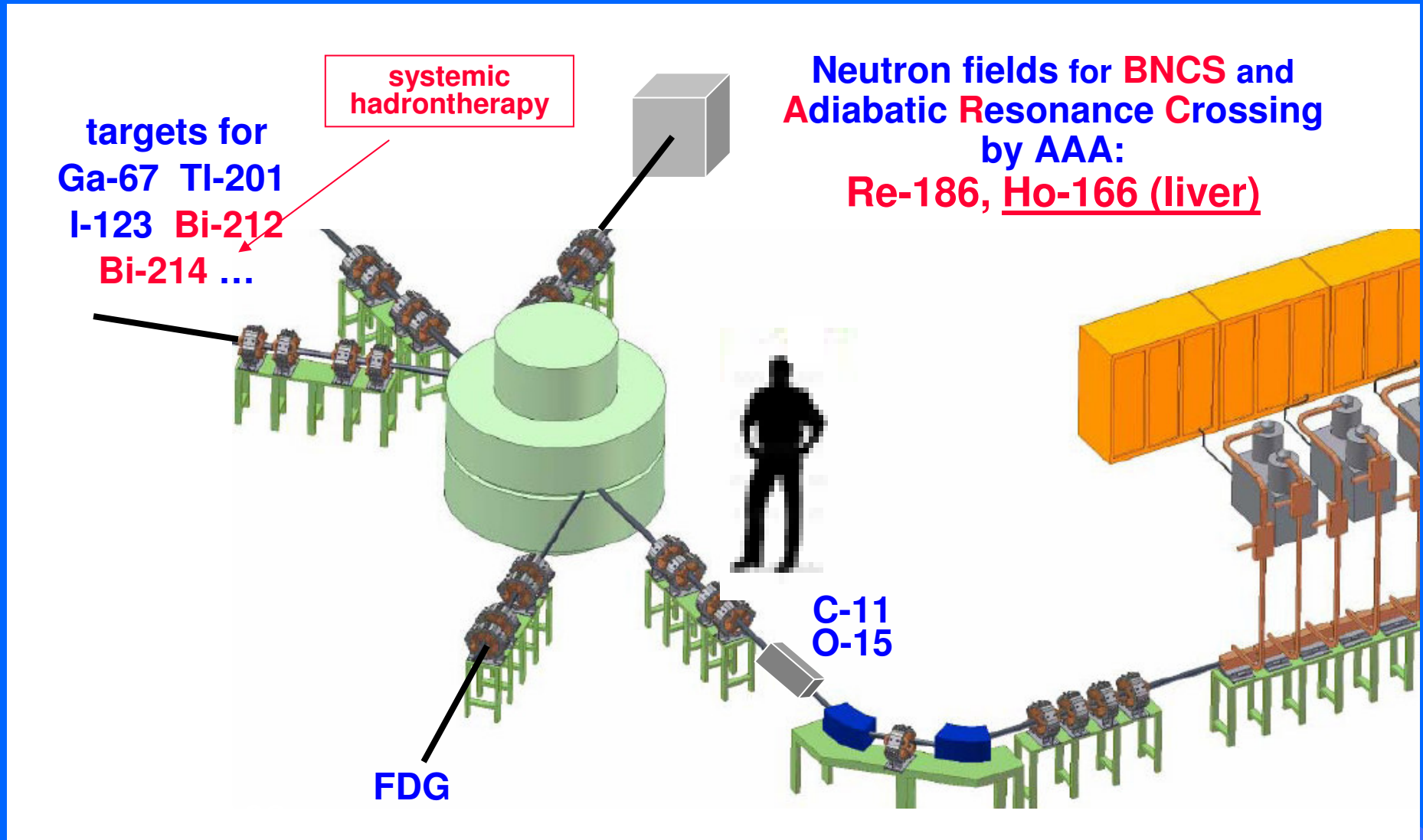
***The linac is made by 20 separately powered modules***



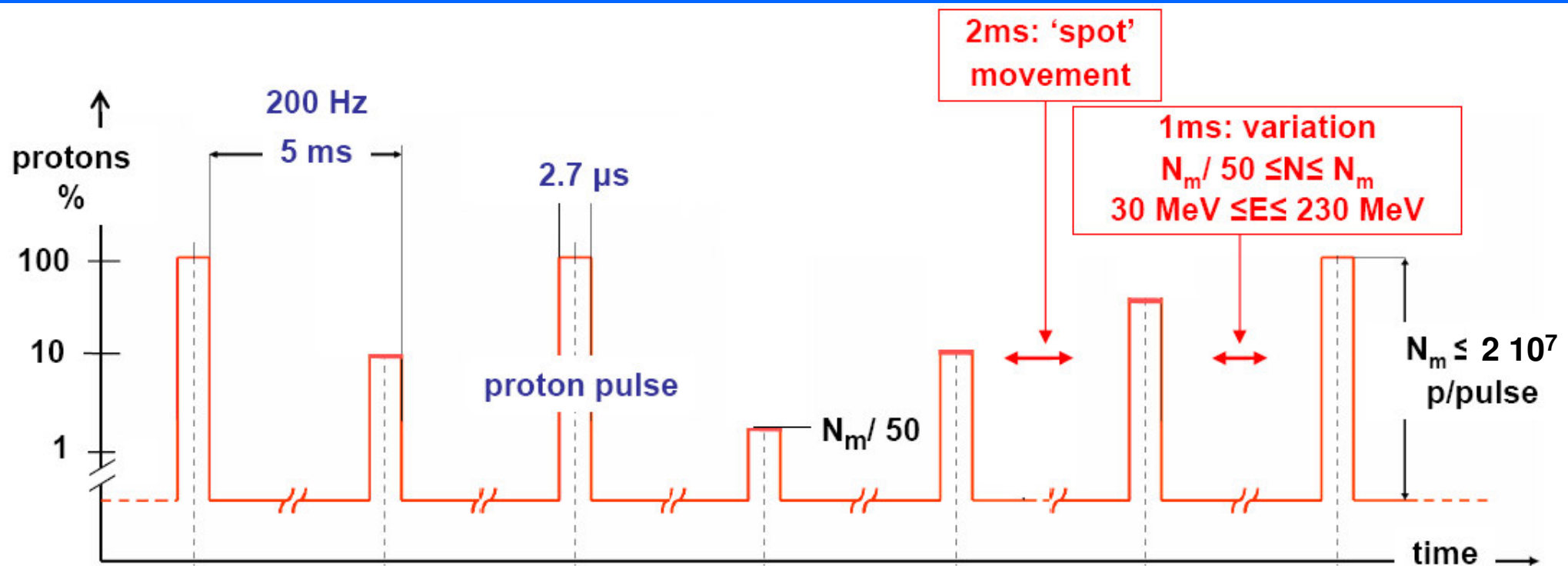
**The first two modules:**

**3 = locations of the permanent quadrupoles**

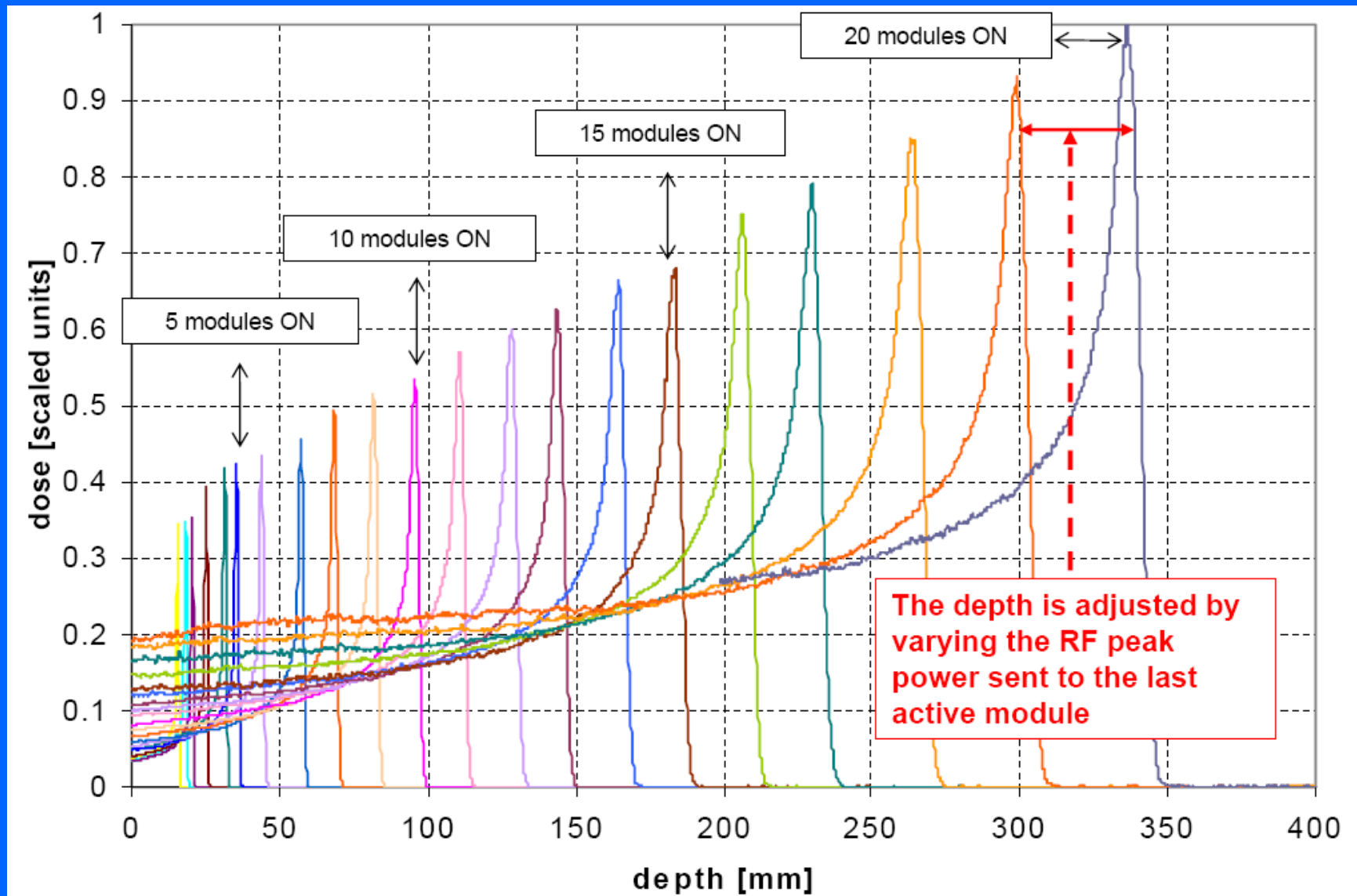
**The cyclotron has 4 high-current beam lines (30 MeV,  $\leq 750 \mu\text{A}$ ) to produce radioisotopes for diagnostics and endotherapy**



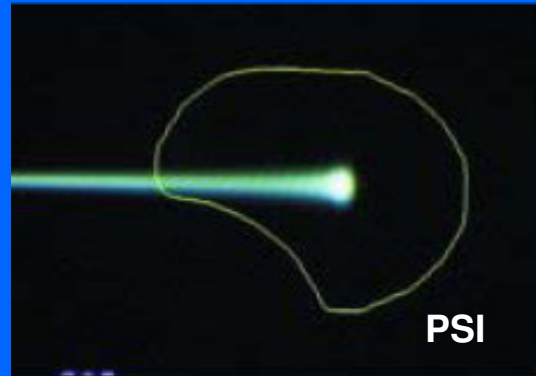
# Time and amplitude structure of the IDRA beam



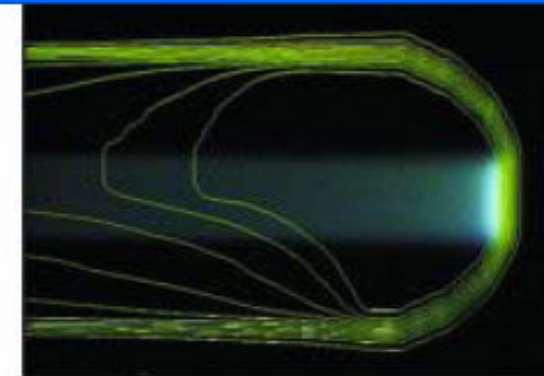
# Depth as a function of the number of active klystrons



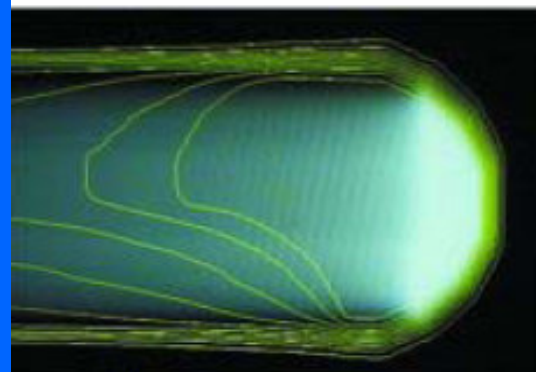
# Use of the cyclinac approach with the PSI technique



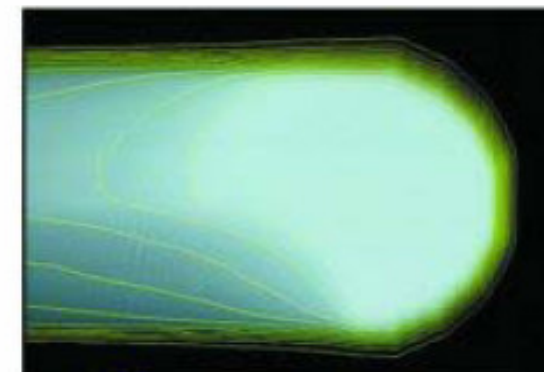
Single 'spot'



Lateral scanning with magnet: 2 ms/step



Depth scanning: vary the power to the 20 accelerating modules



Third scanning by a bending magnet and movable bed

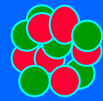
Fast variation by  $\pm 15$  mm  
if the magnetic channel has  
 $\pm 2\%$  momentum acceptance

## *Cyclinac for carbon ions*

# For C ions the TERA solution is based on the 300 MeV/u SC cyclotron designed by LNS

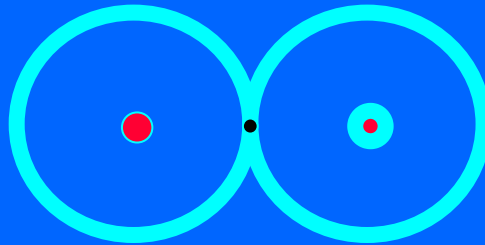
The superconducting cyclotron accelerates particles with  $Q/A = 1/2$

$^{12}\text{C}^{6+}$



$6p^+ + 6n$

$\text{H}_2^+$



$2p^+ + 1e^-$

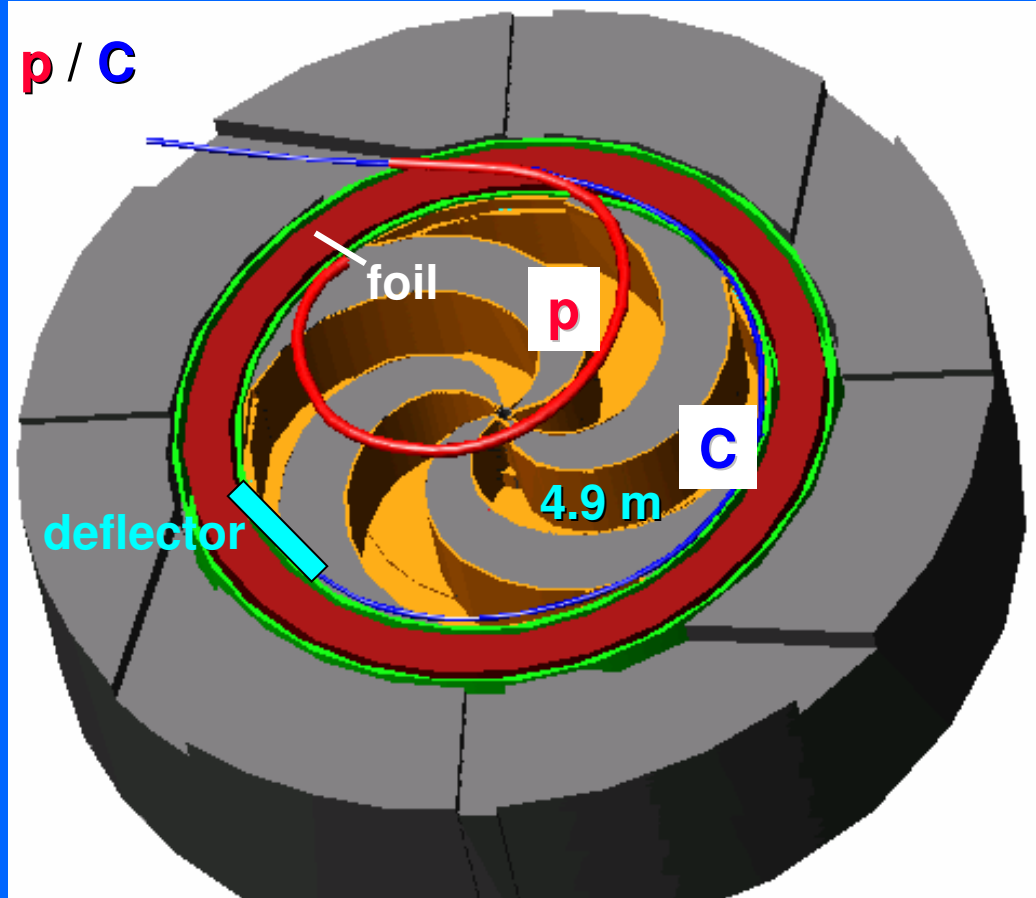
Output energies:

protons

300 MeV

carbon ions

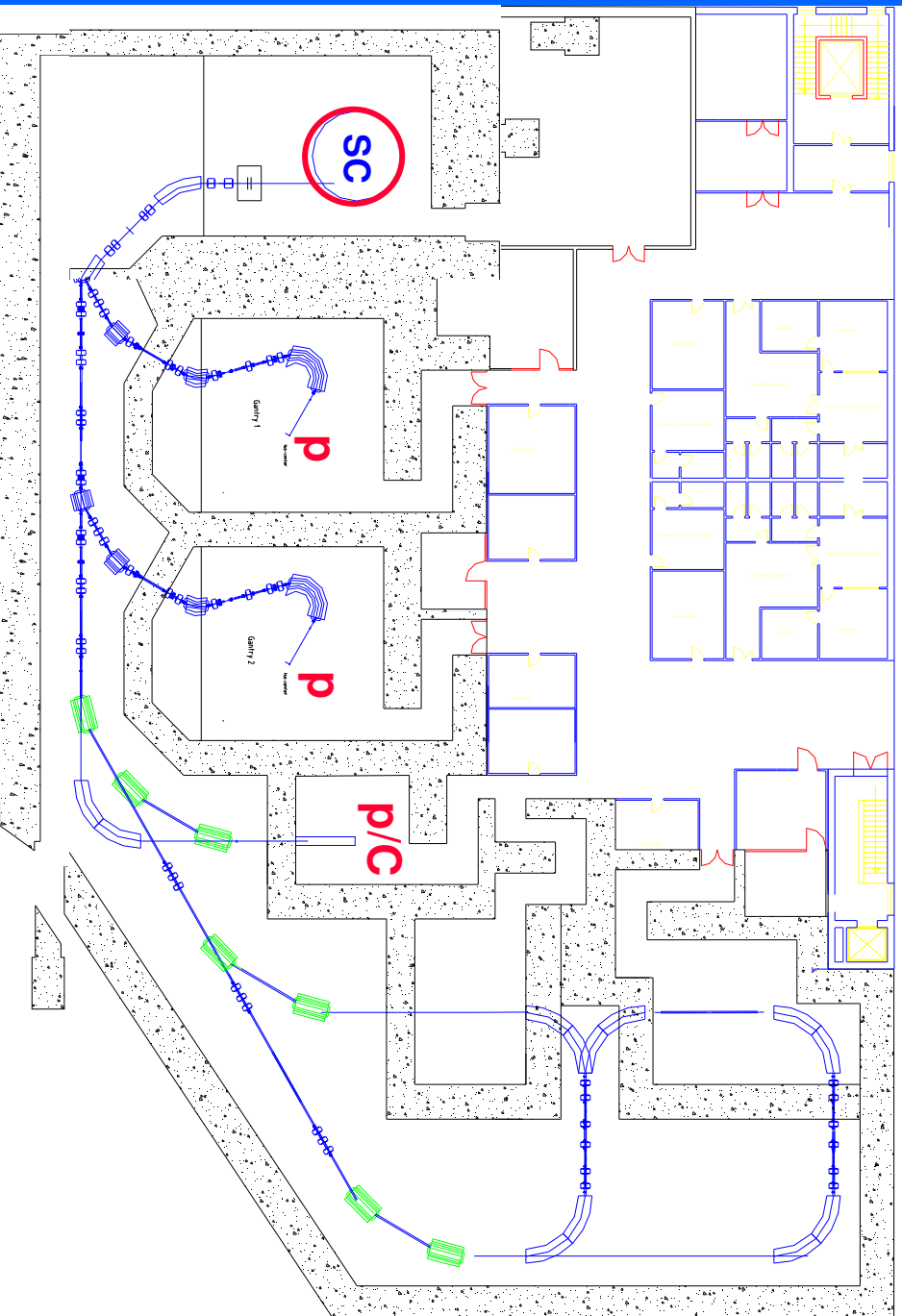
3600 MeV



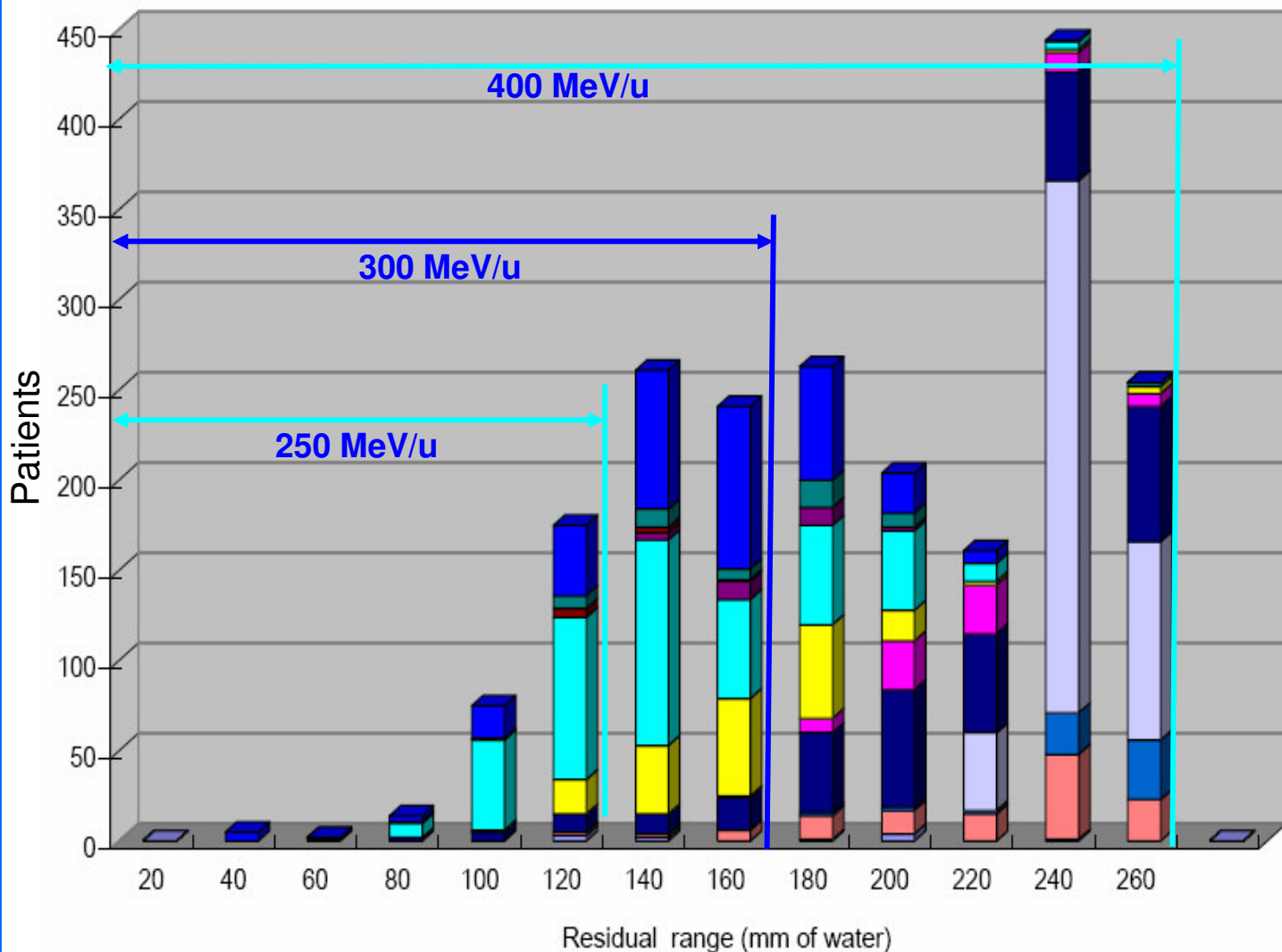
SCENT = Superconducting Cyclotron for Exotic Nuclei and Therapy

IBA (Belgium) offers today the INFN 300 MeV/u SC cyclotron

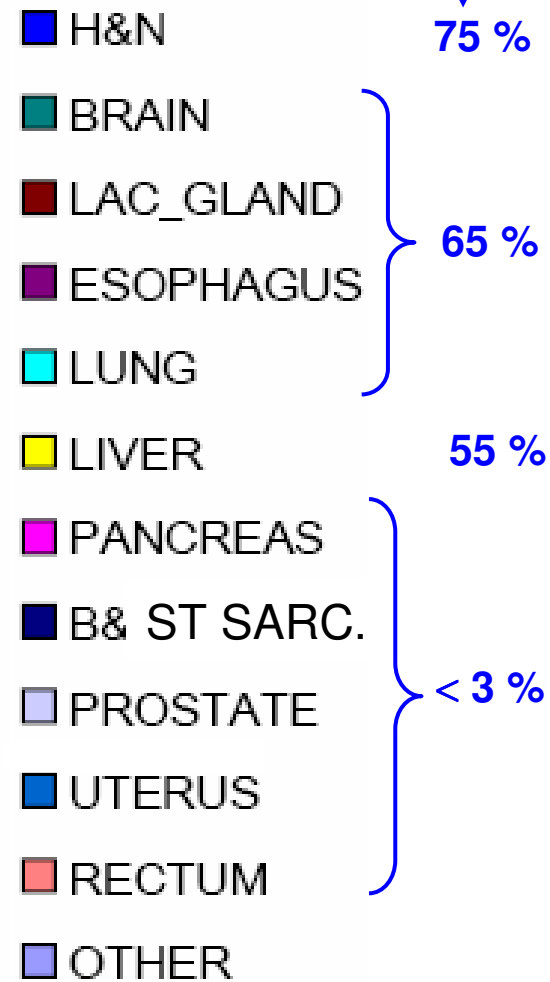
# LNS project for Cannizzaro Hospital in Catania



# HIMAC statistics of the maximum range used per patient

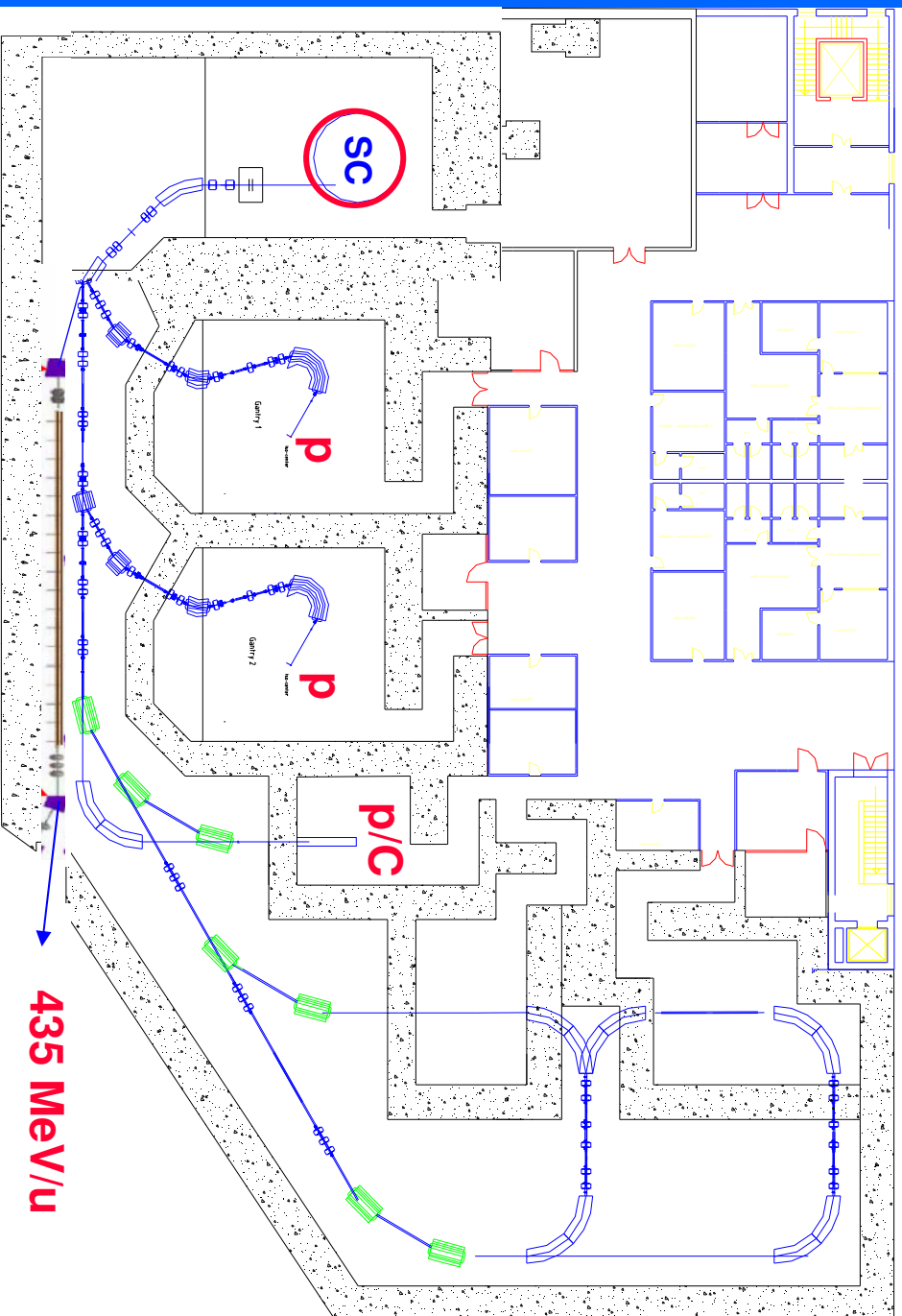


% treated with 300 MeV/u



# Upgrading with CABOTO = CARbon BOoster for Therapy in Oncology

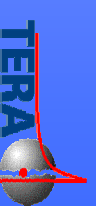
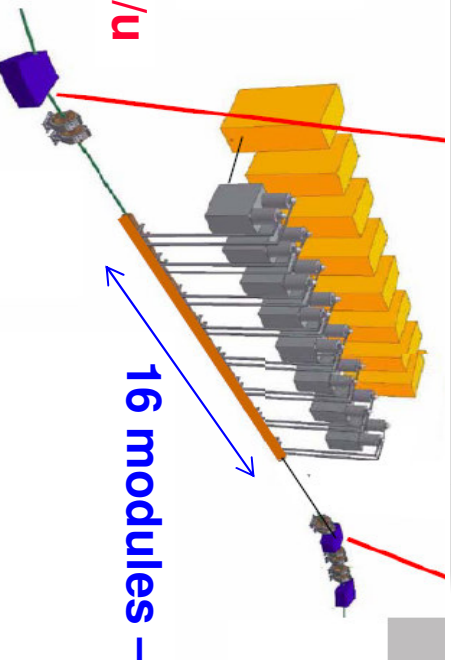
The C<sup>+6</sup> source  
has to be  
developed

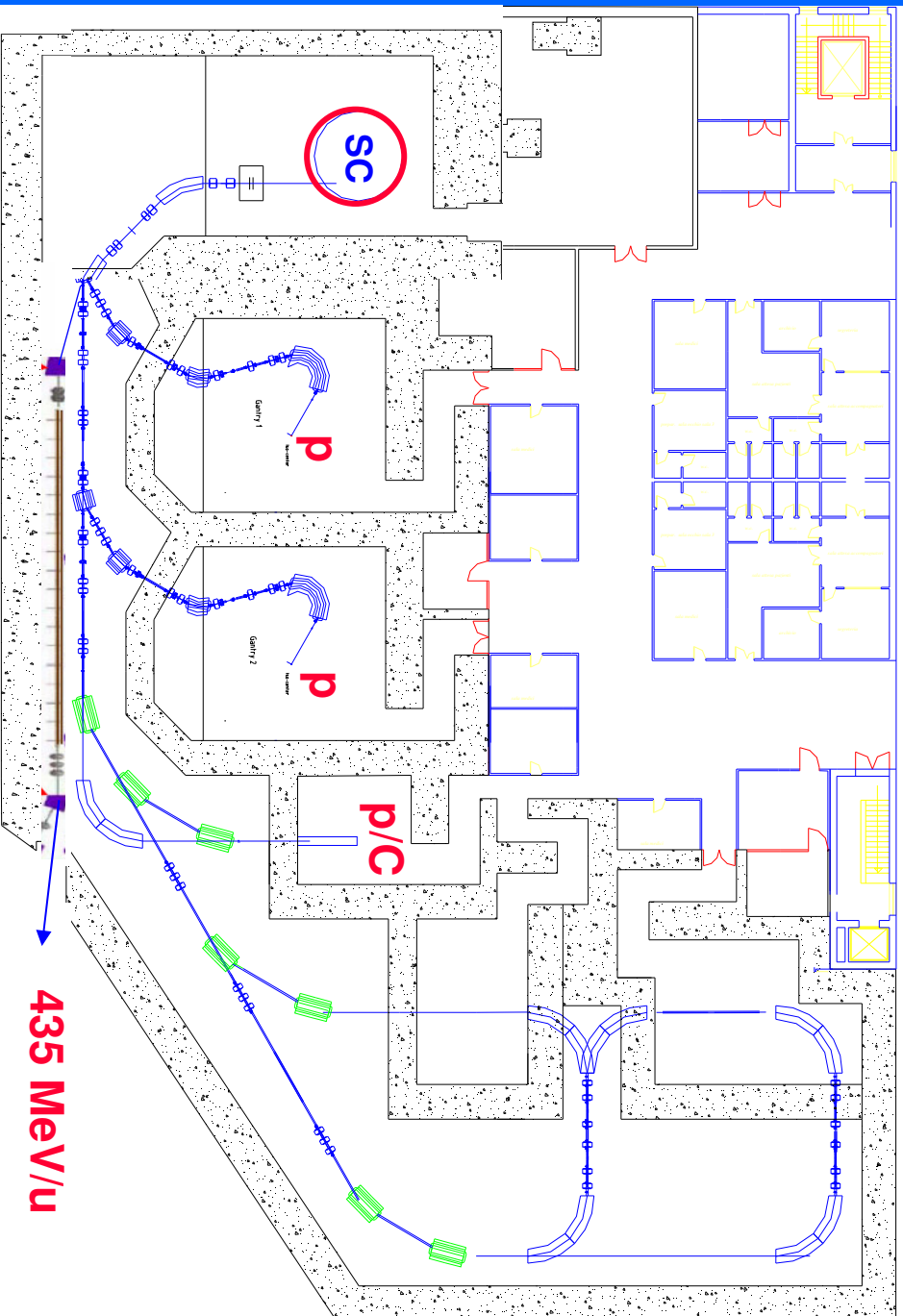


435 MeV/u  
carbon ions

300 MeV/u

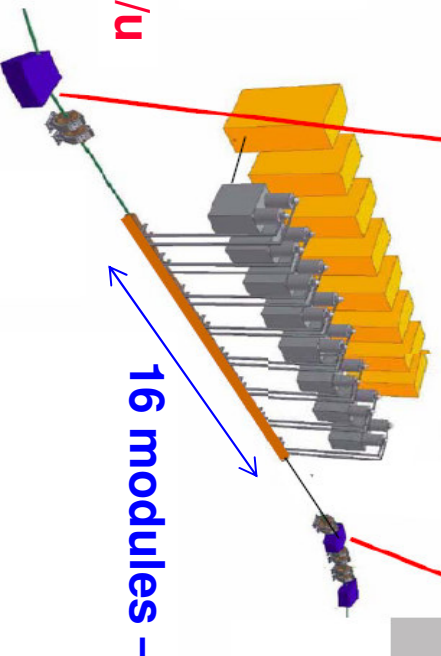
16 modules – 22 m





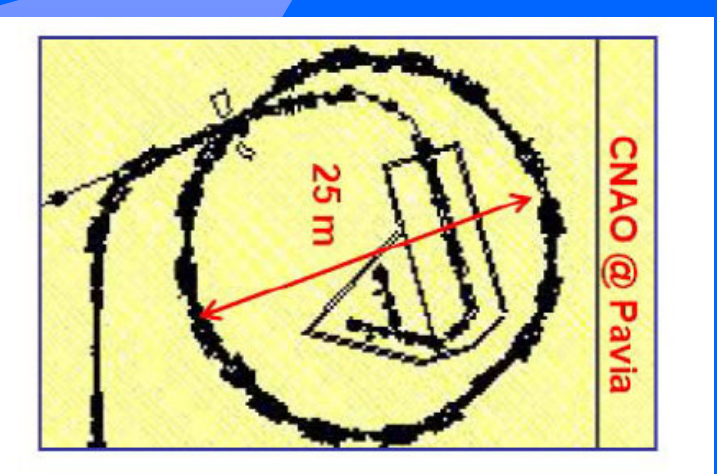
**300 MeV/u**

**16 modules – 22 m**



**435 MeV/u**  
**carbon ions**

## Comparison with CNAO synchrotron



## A module of CABOTO

Power input

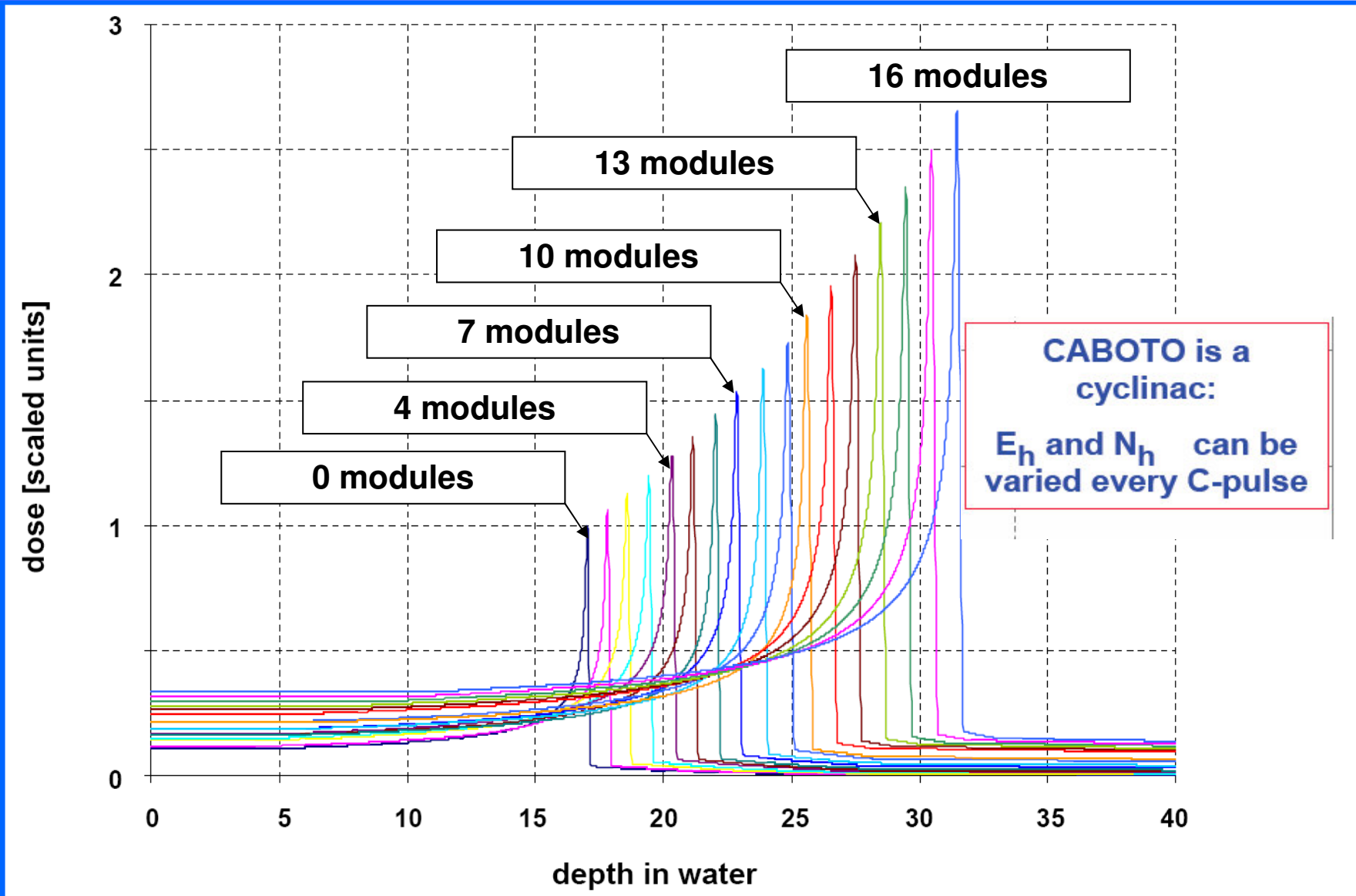
Biperiodic chain of  
accelerating and coupling cells

120 cm

Bridge coupler for the housing  
of a PMQ

End cell for the housing  
of a PMQ

# Ion Bragg curves by adjusting the 16 powers: 300-435 MeV/u



## *The protontherapy Centre in Trento*

## ***Characteristics of the ATreP Centre***

**Centro ospedaliero finanziato dall'Assessorato e alla Ricerca ed Innovazione.**

**Capacità di trattamento di 600 pazienti/anno a regime.**

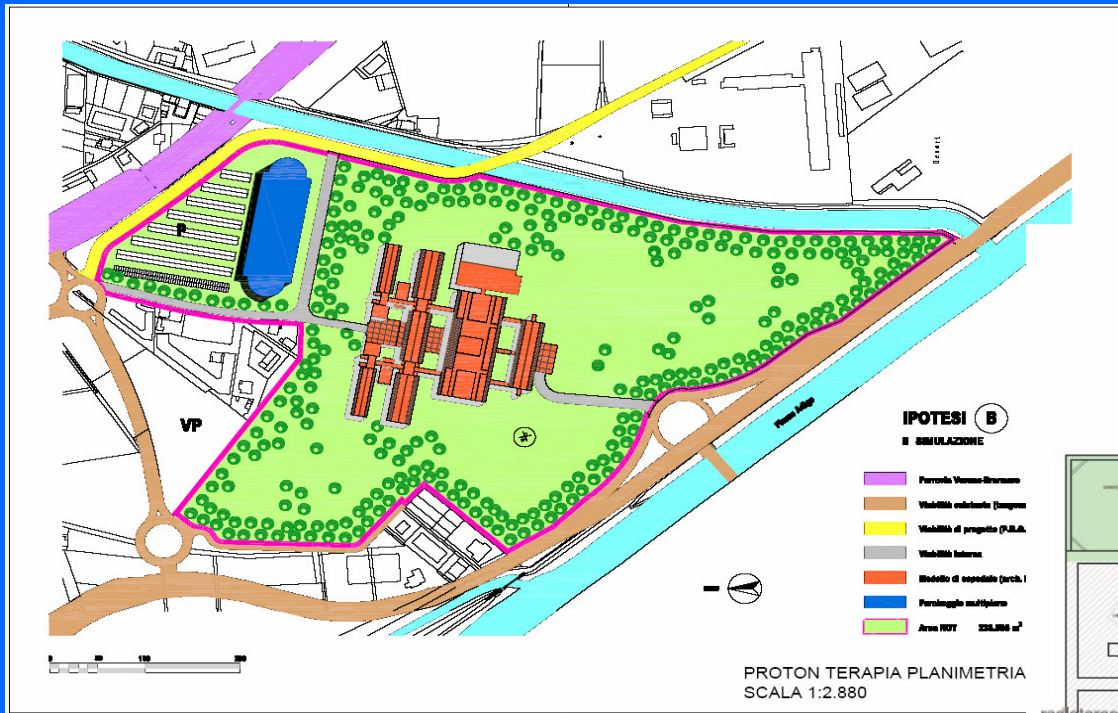
**Una testata isocentrica e una linea fissa, con la predisposizione per una seconda.**

**Spazzolamento attivo.**

**Preparazione del paziente fuori della sala di trattamento.**



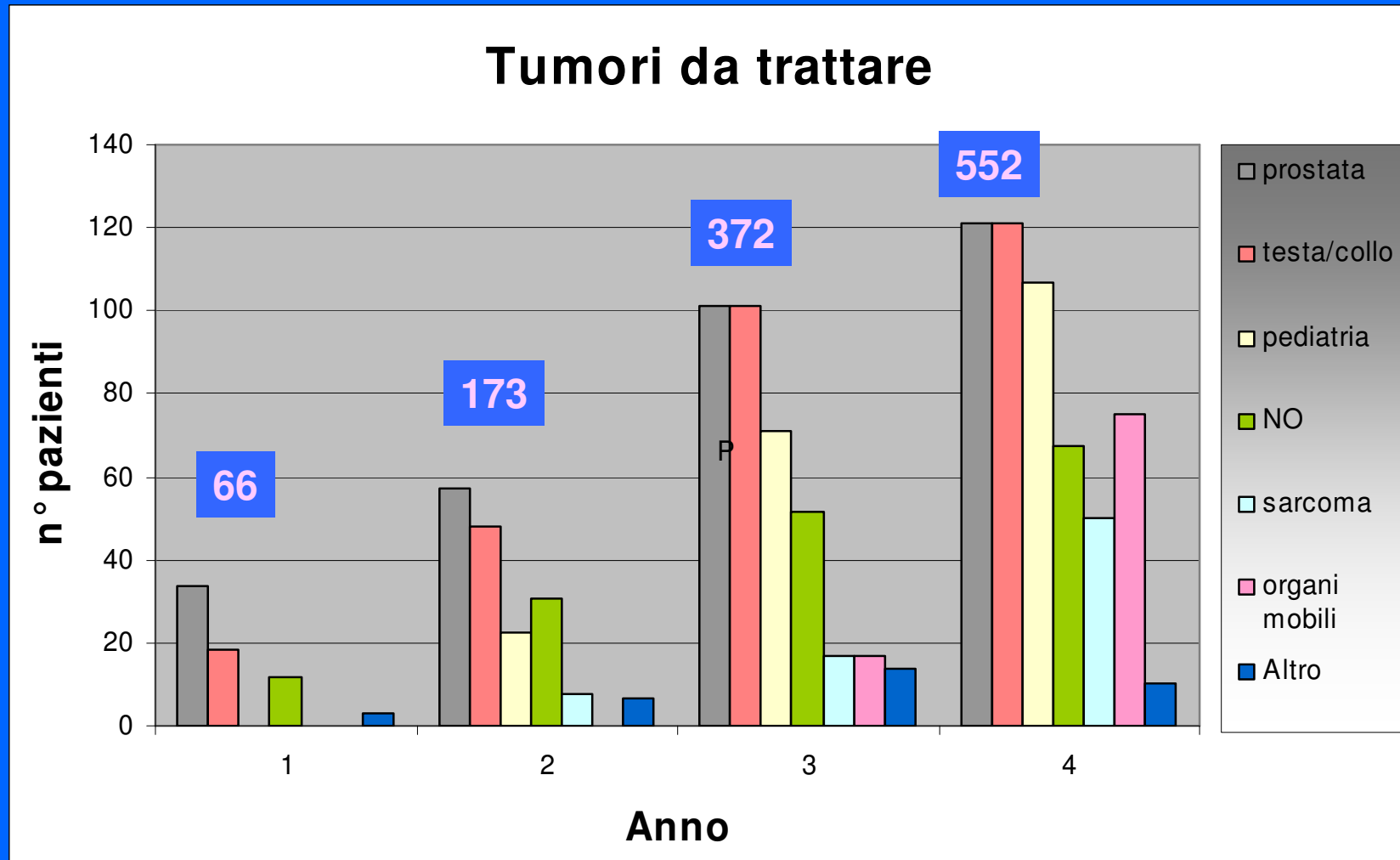
# Foreseen solution



Un solo fornitore responsabile dell'intero complesso

Contratto di manutenzione e probabilmente di operazione a lungo termine.

## Treated tumours in the first 4 years



***THE END***