

France HADRON:

benefits, challenges and future direction of a national collaboration

Jacques BALOSSO, MD, PhD

University Joseph Fourier, Grenoble 1

Jean Louis HABRAND, MD, PhD

University of Caen, Centre François Baclesse and IGR Coordinators of France HADRON



















What is France HADRON?

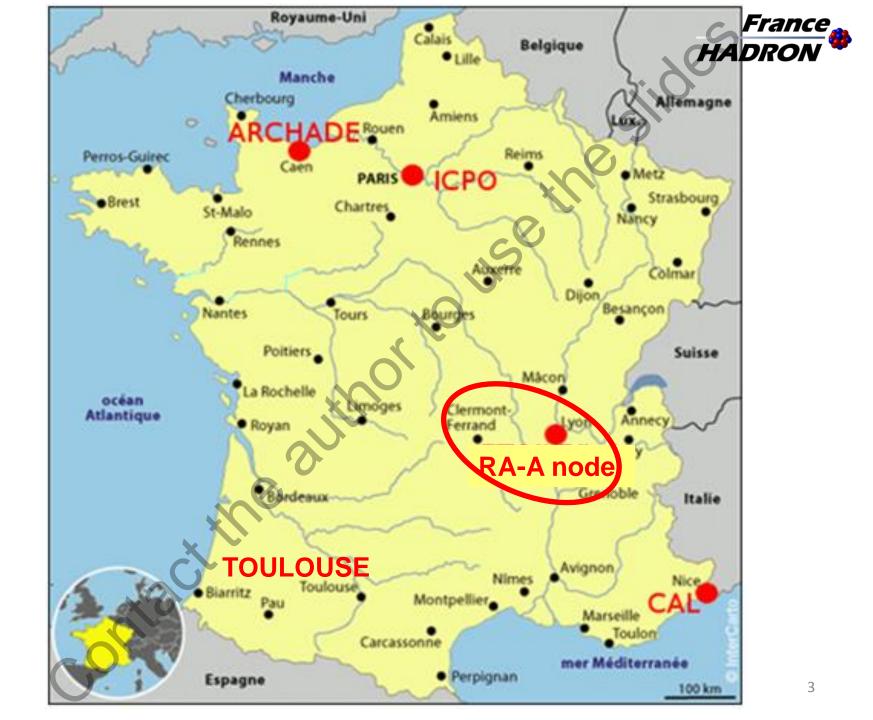
France HADRON has three dimensions:

1) It is a network of institutions engaged in hadrontherapy research and development with identified teams, laboratories and scientific capabilities and/or facilities:

These institutions are located in 5 sites in France:

- Lyon-Clermont-Frd (IPNL, LPCC, and late ETOILE project)
- Caen (LPCC, ARCHADE project, GANIL is coll)
- The 2 protontherapy centers at Nice (CAL)
- and Orsay (ICPO)
- Toulouse (CCR and Pericles project)

With the partnership of CNRS/IN2P3 (IPNL & LPC) and of IRSN,





What is *France HADRON*? (2)

France HADRON has three dimensions:

- 2) It is <u>a scientific collaboration</u> gathering all the scientific teams engaged in hadrontherapy research and development in France.
 - More than 25 teams are thus identified and participating to the scientific project of FrHA
 - These teams are located in many places, not only in the nodes of the network:
 - Paris, Strasbourg, Lyon, Marseille, Grenoble, Clermont-Fr, Toulouse, Orsay, Nice, Caen, Nantes...

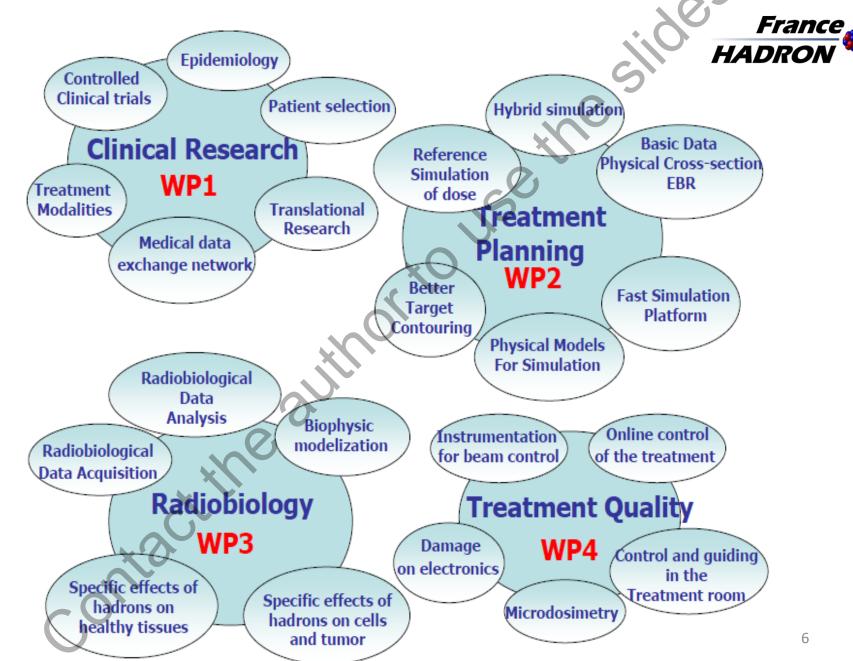
This collaboration is organized in 4 working packages



France HADRON - 25 teams

- Multidisciplinary: medicine, physics, biology, computer, etc.
- Teams involved :
 - Lyon Clermont-Fd: teams of C. Rodriguez-Lafrasse, M. Beuve, D. Dauvergne, G Montarou, N. Foray, D. Sarrut, B. Shariat, B. Ribba, P. Pommier.
 - Nice: teams of JM. Hannoun-Levi, P. Mandrillon.
 - Orsay: teams of A. Fourquet / R. Dendale, A. Mazal, J. Hall / F. Pouzoulet.
 - <u>Caen</u>: teams of D Cussol PJ. Colin, M. Bernaudin, K. Boumediène, JL. Lefaix, MH. Moscatello, JL. Habrand.
 - Toulouse: teams of E. Moyal, P. Celsis / A. Laprie / M. Delannes, R. Ferrand
 - Marseille: team of Ch. Morel
 - Strasbourg: Team of D. Brasse/ Rousseau

France HADRON - scientific collaboration





France HADRON - WP leaders

- Organized into 4 working packages:
 - WP1 How to identify and assess the medical value of hadron therapy (clinical research); Jean Louis HABRAND (Caen-Paris)
 - WP2 How to improve treatment plans (measurements, modeling and computer simulation); Daniel CUSSOL (Caen)
 - WP3 How to better understand the effect of treatment (radiation biology, radiotoxicology); Claire RODRIGUEZ-LAFRASSE (Lyon)
 - WP4 How to improve the quality control of treatment (instrumentation);
 Denis DAUVERGNE (Lyon).



What is France HADRON? (3)

France HADRON has three dimensions:

- 3) It is also an <u>economical program</u> as one of the ≈ 12 <u>national infrastructures for health and biotechnology</u> development supported by the so called "grand emprunt" launched in 2010 by the French government.
 - The goal of this program is to sustain investments for future economical and societal returns.
 - France HADRON has thus been credited of a global amount of 15M€ for seven years.
 - Beyond this date it will have to find its own incomes to continue.



France HADRON received 15 M€ HAD for 2013-2019 principal support

The provisional distribution is the following:

- 5495,60 k€ for the general activities, the <u>beam</u> access cost and the clinical research support
- 5900,00 k€ for equipments to make possible new and easier beam access: 5 M€ for proton in Nice and Orsay and 0,9 M€ for carbon at GANIL
- 3604,40 k€ are kept as a reserve for future developments presently in discussion.



From where comes France HADON? A bit of history (1)

The France HADRON collaboration is based on a large heritage:

- The sites of Caen (GANIL), Nice (Médicyc) and Orsay (ICPO) are existing for more 20 years...
- A Regional program for hadrontherapy research (PRRH) has been initiated in 2002 in Lyon just after the announcement of the carbon ion project in 1997, known as ETOILE project
- The CNRS launched the Mi2b GDR, end of 2004
- Caen presented its own project, Asclépios, in 2005
- A first attempt of national program in 2007 did not succeed (opposition of the INCa...)
- However the 3rd "plan cancer" (2014-2019) proposes a "national regulation for expensive facilities as for example protontherapy facilities 10

France

A bit of history (2)

- **Projects** for new centers or upgrade of existing centers are existing for both carbon ions and protons: at Lyon (former **ETOILE**), Caen (**Archade**), Orsay (2010), Nice (2014-16) and more: Toulouse, Nantes, Lyon, ... for **protons**.
- Since 2001 the French teams have effective links with the CERN, GSI, CNAO, MedAustron and the NIRS. Moreover, they have actively participated to every European consortia and research programs as: Enlight (2001-...), Enlight++, ULICE (2009-14), ENVISION, Partner (2010-13), Intervision.
- Teams from other towns than those having project of centers are joining and actively participating, mainly through CNRS teams: Clermont-Ferrand, Marseille, Strasbourg, Grenoble... totalizing more than twenty research teams all over France.
- Hadrontherapy applications and researches are thus active in France for more than twenty years

A bit of history (3) Difficulties to overdraw in 2010 HADR

- The spreading (and sometime competition) of the material and teams
- The deep lack of access to the beams, which are either:
 - In medical centers with few research infrastructure (Nice, Orsay)
 - Or in research centers which agenda is not specifically devoted to hadrontherapy (GANIL)
- A weak industrial and governmental support in France although a very high know how in research institution (IN2P3, CEA) and some well known small corporations (Pantechnic, Sigmaphi)
- The lack of referent carbon ion center in France and advanced protontherapy center
- And the lack of controlled randomized comparative clinical studies in the domain of hadrontherapy although 27 years of neutrontherapy and 22 years of protontherapy in France (circa 10000 patients, 50% having ocular tumours)



France HADRON – objectives are to overdraw these difficulties

- To <u>federate</u> research teams and organize research at a national level,
- To fund beam time and beam line access for research.
- To <u>open</u> new research beam lines for protons and carbon ions, to increase available beam time
- To <u>optimize</u> technical means and procedure for hadrontherapy,
- To network in the frame of the European programs
- To have a positive input on economy



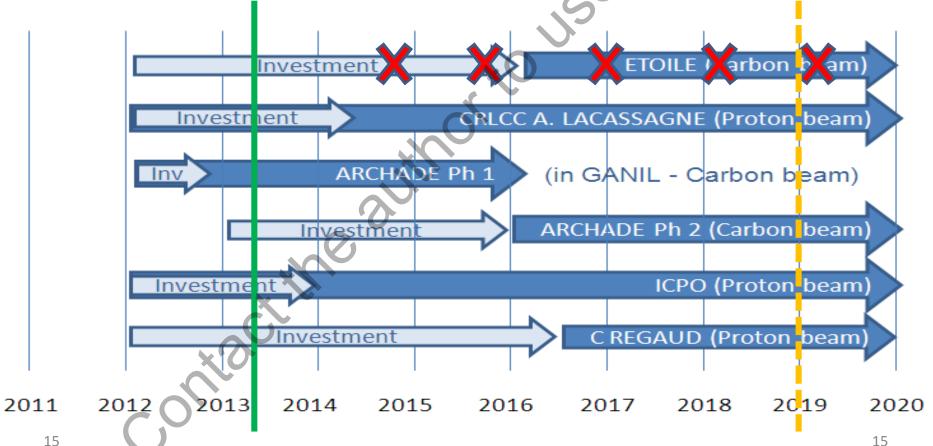
National Agency for research (ANR) and CNRS for management (DR19 in Caen)

France HADRON **Coordinators: Jacques Balosso** & JL Habrand (3 years mandate) Assistance (2 persons) Steering Committee (SC/CD) (President Gérard Montarou, 3 years Scientific mandate) **Executive Management Committee / Advisory** - IN2P3 representative **Experience Committee (MC/CG) Board (SAB)** - ITMO TS representative (monthly meeting) - ITMO Cancer representative (President Régis Férrand, 2 years (annual meeting) Partners and Node mandate) representatives - Node representatives Scientific representative of - WP leaders territorial Councils Regional representatives of Minister of science (DRRT)



France

The financial starting point is 1st of March 2013, the financial agreement has been signed mid September 2013, the launching meeting took place in **Meudon (near Paris) the 14th of October 2013**



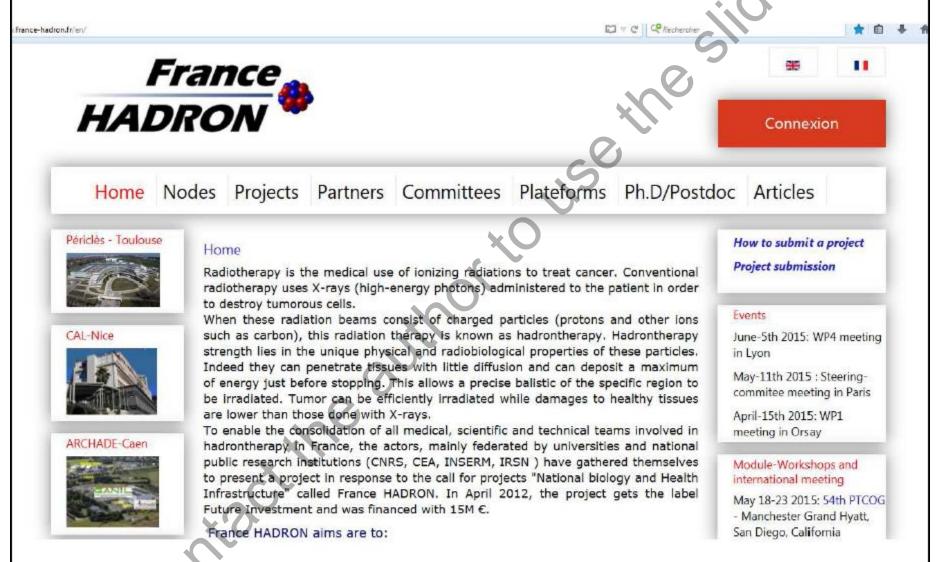
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FrHA early experience (1)



- 300 hours of beam access have been provided
- 5 internal meetings for scientific coordination have been carried out
- Monthly remote of presential meeting of MC
- 2 meetings/y of the SC
- 2 Scientific advisory board meetings in 2011 and 2015
- A consortium agreement has been signed by 23 parties
- Scientific production:
 - Since 2013: 47 full international papers, 1 patent
 - About 50 participations to meetings (oral or poster)
 - Most of the publications are still issued from **Lyon** group \approx 50%
- FrHA has a web site: www.france-hadron.fr

The Web site: www.france-hadron.fr



FrHA early positive experience (2) HAD



- Thanks to FrHA the hadrontherapy research domain is now:
 - unified and coordinated
 - has gained more visibility
 - and should have continuity... hopefully.
- WP leaders have a very important role and are gaining visibility and authority to manage their part of the project
- The success of R&D in new imaging instrumentation and in high LET radiobiology in Lyon
- Rising demands internal and external to participate...
- Good and active participation to the WP meetings

FrHA early experience (3)

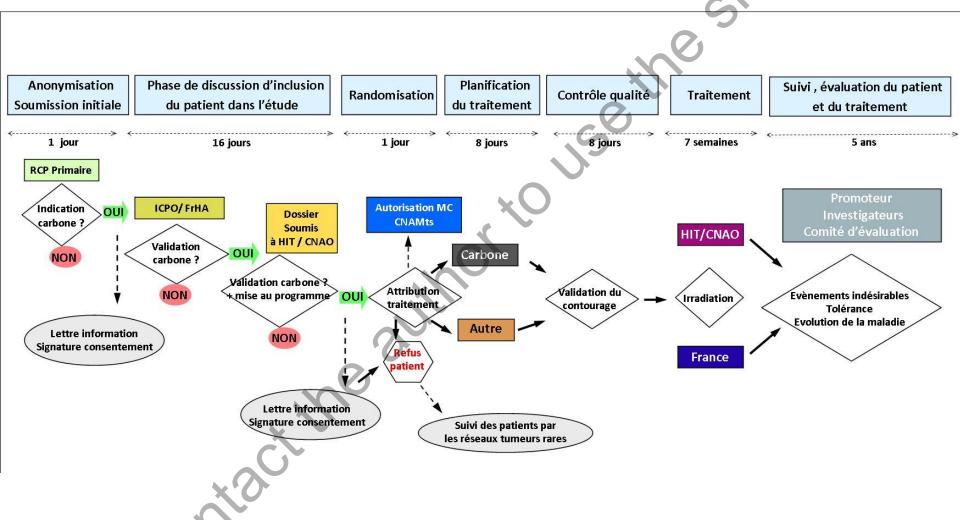


- Challenges are many!
 - Turning into cooperation former competition
 - Coping with oriented and medically applicable research goals
 - Running a multi site multi institutional multi disciplinary –
 multi personalities ... project
 - Balancing protons and carbon ion researches and institutional interests
 - Attracting more MD interest AND participation...
 - Attracting less eventually futile demands "just to test hadrons"...or so.
 - Producing good science and publishable works from new comers although long lasting wait for beam and repeated experiments...
 - How to integrate new scientific teams and new facilities?

Future directions

- France **
 HADRON
- For clinical activities two project are going to boost participation:
 - The multicenter international clinical trial France HADRON –
 CNAO ULICE: it will involve all the large radiation oncology department of France
 - The ProtonShare network to access comparison of treatment plan for decision making in protontherapy
- For R&D FrHA has to make relevant choices for the complementary investments to strengthen its best positions: carbon ions scientific facility and hadron QA imaging for treatment ...
- More implication in medical training activities
- To have **yearly scientific assessment** (SAB) about different topics each year
- To participate actively to European programs (as did ETOILE) in the frame of ENLIGHT and Horizon 2020 and maybe international one in collaboration with NAPTA.

Workflow of the FrHA - CNAO - ULICE trial



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ProtonShare



Refering centers

Particle therapy **Centre**

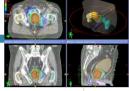


Automatically Comparison of the treatment plans

DICOM data Images, Contours, **Doses Medical data** Keyboard or .pdf







Planification for protons

Acceptation of the treatment

Treatment total of partial? Follow-up? Registration of side effects



Report of the **Expert**

Review of the medical record







Automatical analysis of the side effects



Extensive medical and dosimetric data hase







Action in Share Place

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Thank-you

Sontact the author