

SIRIC Montpellier Cancer

Professor David AZRIA

6th December 2022



SIRIC Montpellier Cancer audition team



**Nathalie
BONNEFOY**
Director of IRCM



Sophie POTY
Doctor in chemistry
Expert in RPT

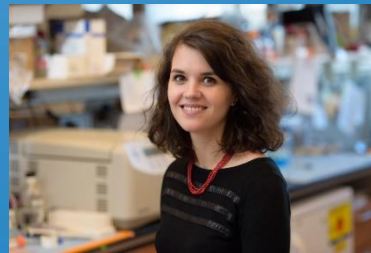


**Laurent
LE CAM**
Team Leader at IRCM
Deputy director IRCM

**David
AZRIA**
Head of Radiation
Oncology Dpt
Scientific Director ICM



**Vanessa
GUILLAUMON**
SIRIC General Manager



**Florence
COUSSON-GELIE**
Scientific Director of
Prevention Center (ICM)
Co-Director of Epsilon
Laboratory

Presentation outline

- **Context and environment of the application**
- **Project 2023-2027**
 - Partner institutions
 - The new strategic action plan
 - Governance and management
 - Scientific strategy
 - Health democracy policy
 - Provisional budget
- **Conclusion**

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Montpellier: Territory of research and innovation

□ National leading academic research community

- TOP 5 in BioHealth
- Shanghai Ranking : TOP 200 World, TOP 4 France outside Paris

□ Recognised for its scientific excellence



- Labelled by the French State for “Feed-Protect-Cure” Programme

□ Strong innovative capacity



- Incubator Business Innovation Center: World Top5 UBI Global
- One of the 5 French « University Innovation Cluster »

□ Active political support



- Med Vallée project of Montpellier Metropolis dedicated to Biohealth



Faculties of Medicine of Montpellier



SIRIC has been key for structuring cancer research in Montpellier since 2013



SIRIC
MONTPELLIER CANCER
Site de Recherche Intégrée sur le Cancer

Translational research capacity

Novel cutting-edge resources and core facilities

Collab. clinicians & researchers

×4 clinicians in labs
ERC 2022 Starting Grant (MD)

Tech transfer & valorisation

4 new start-up companies
NovaGray, DiaDx, MabQi, Diag2Tech

Training in cancer research

« **Cancer Biology** » Master II program
Erasmus Mondus

Interdisciplinary research

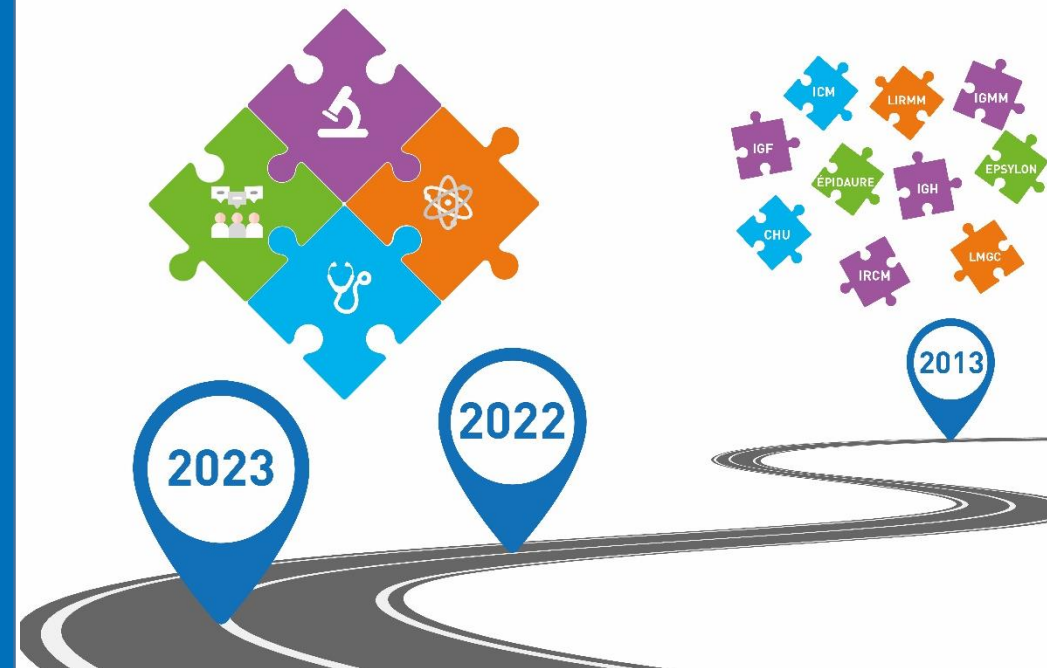
New strategic partnerships with the Chemistry and MIPS communities

Strong leverage effect

Up to ×10
from SIRIC seed funding



Prof. Marc YCHOU
SIRIC Director 2013-2022
Current ICM General Director



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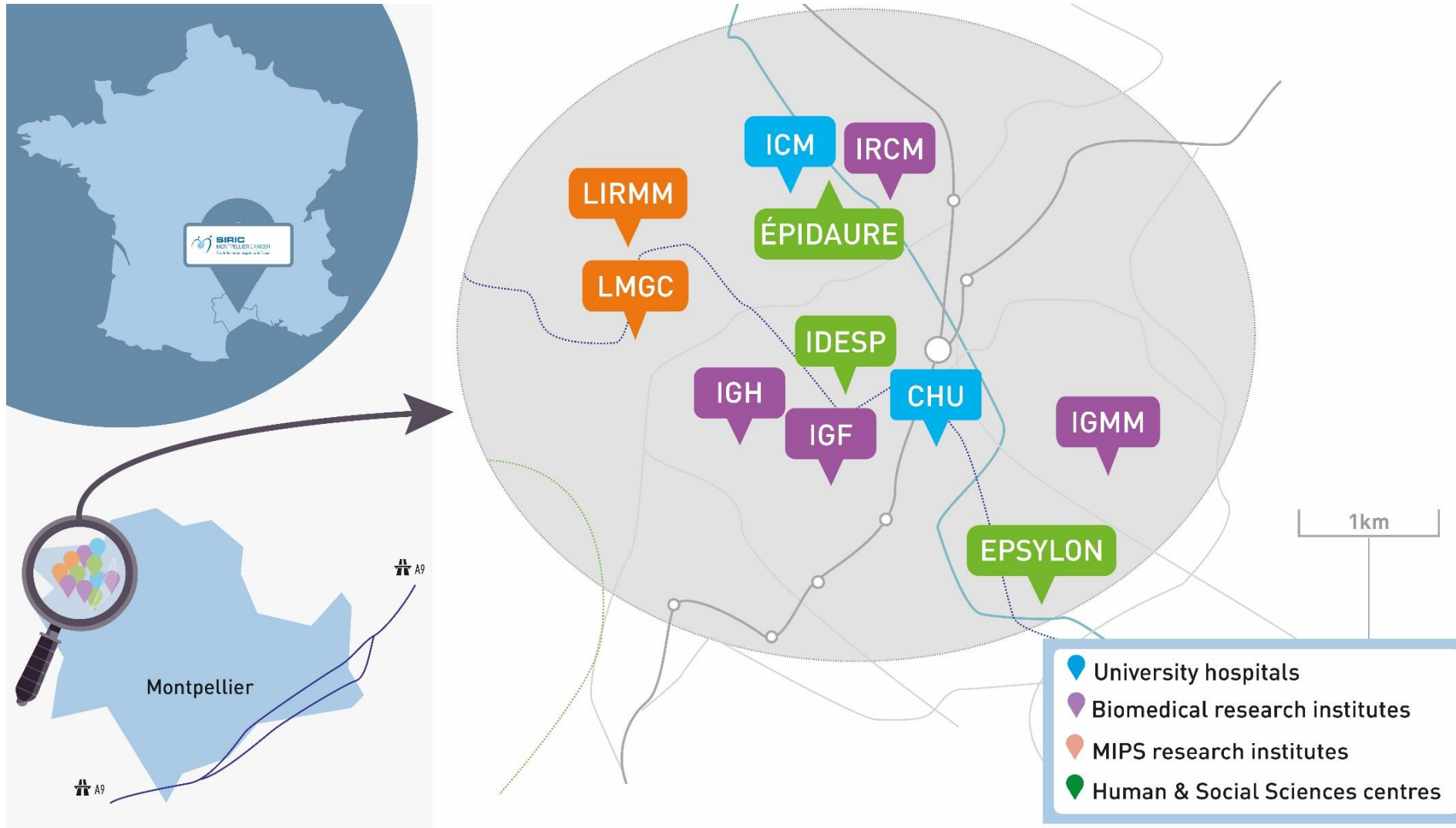
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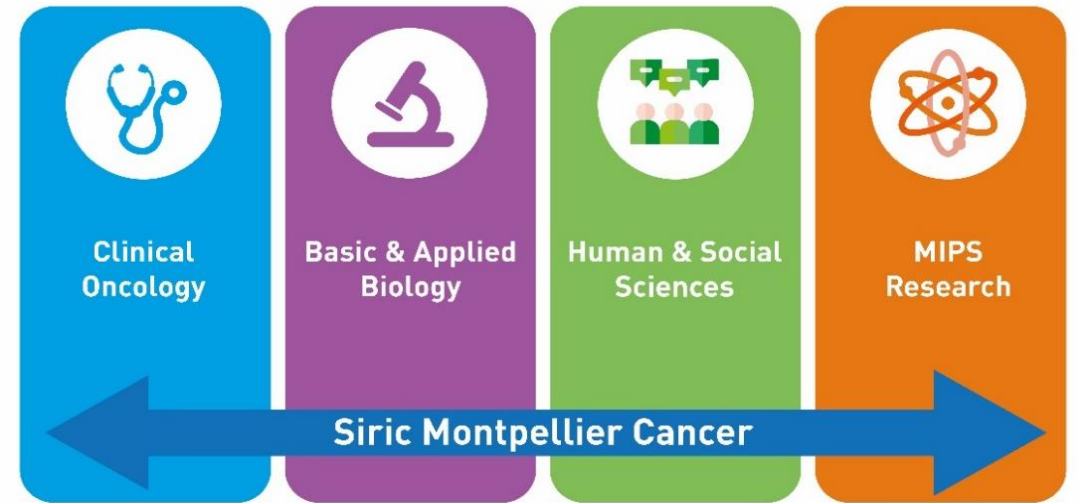
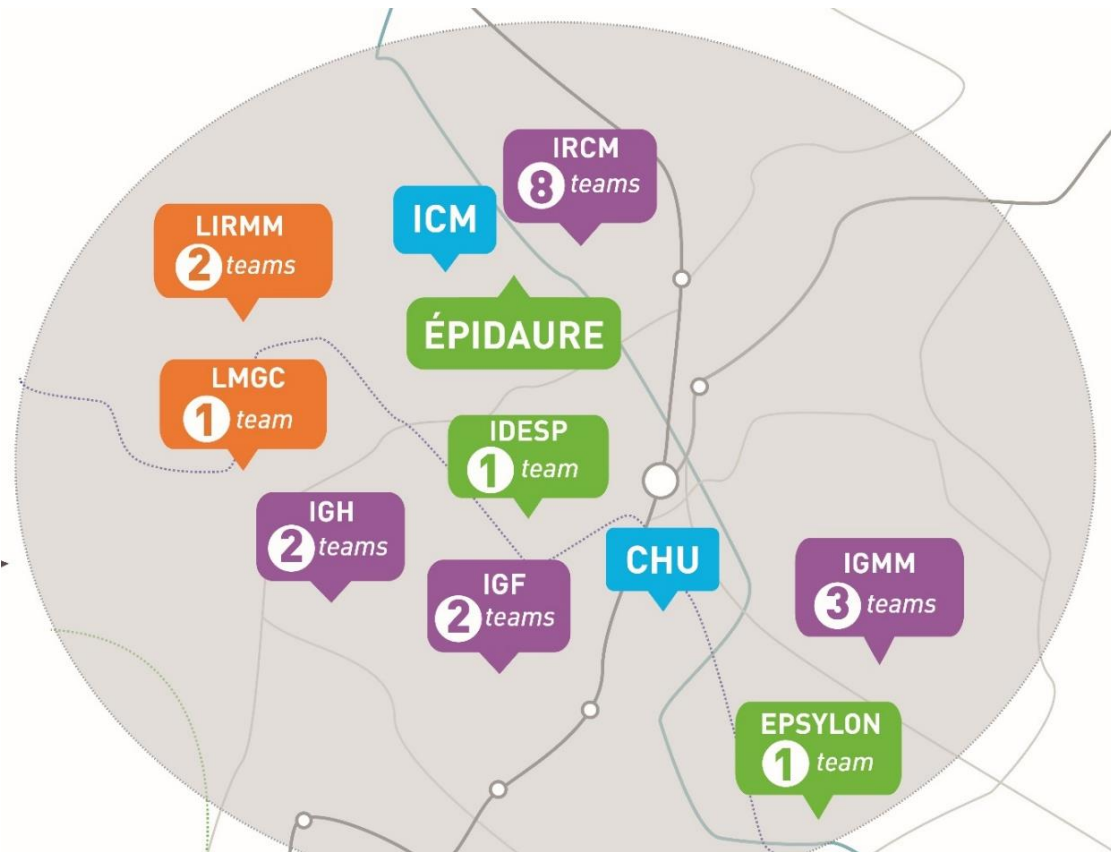
SIRIC Montpellier Cancer

Critical mass of key oncology leaders within a 1-km radius



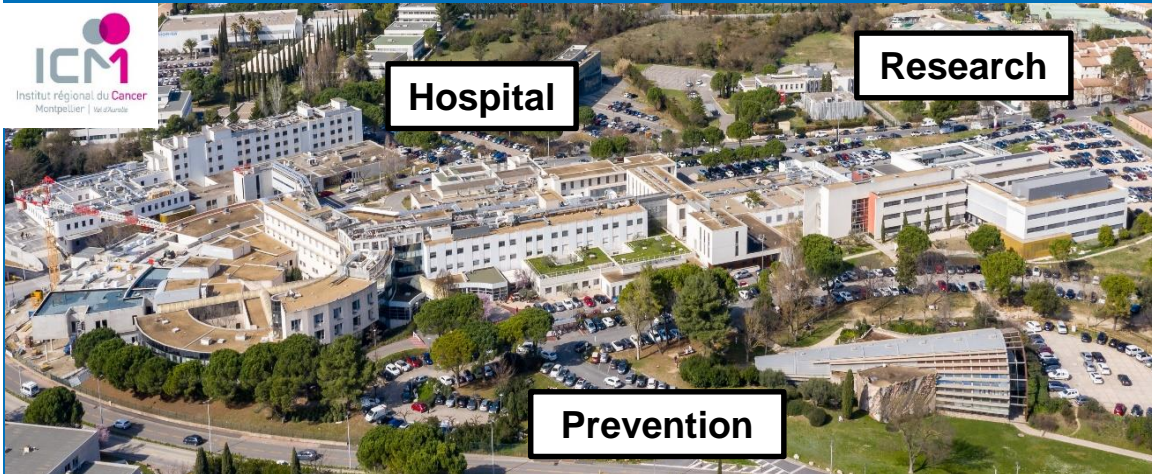
SIRIC Montpellier Cancer

Integration of medical, scientific and societal research in oncology

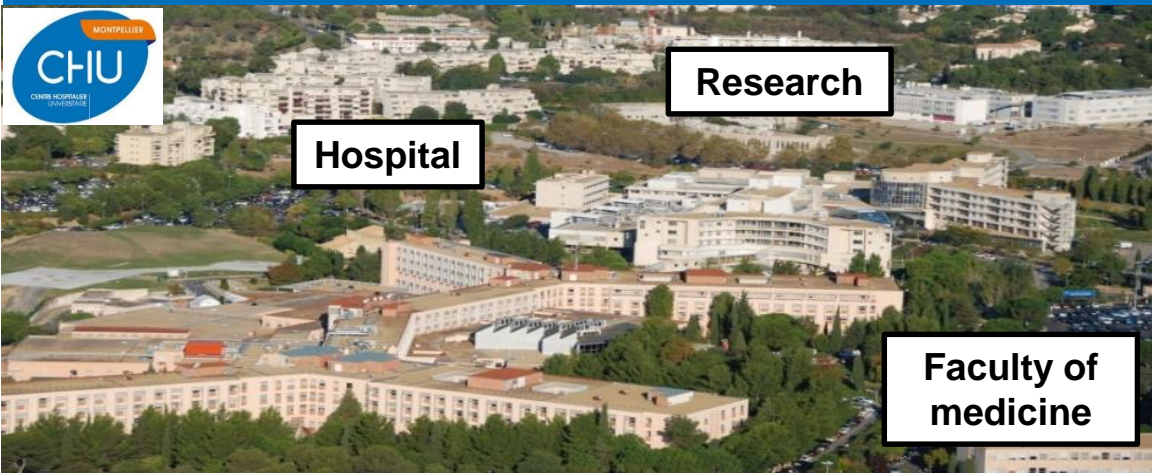


20 constitutive research teams
25 clinical teams and support infrastructures

Montpellier Cancer Institute



Montpellier University Hospital Centre



Montpellier clinical oncology

Multidisciplinary clinical expertise

Two major university hospitals

- ❑ 200 physicians in oncology
- ❑ Active file of 50,000 patients/year
- ❑ > 2,000 patients enrolled in clinical trials annually
- ❑ A portfolio of 250 clinical studies / year

High-quality clinical research

- Clinical and Translational Research certified ISO 9001
- Biometry certified ISO 9001
- ICM/CHU Early-Phase Clinical Trial Centre certified by INCa
- Unique French Datacentre for UNICANCER Federation



A strong synergy of expertise in cancer biology

IRCM and 3 multi-thematic research institutes including top-notch research groups involved in cancer biology

Institute of Cancer Research of Montpellier



- Focus exclusively on cancer research

Institute of Molecular Genetics of Montpellier



- Basic research on liver biology, T-cell differentiation and immunotherapy

Institute for Functional Genomics



- Expertise in epigenetics

Institute of Human Genetics



- Expertise in genome dynamics and epigenetic controls



A unique cluster of expertise in Human and Social Sciences applied to cancer

Highly specialised in interventional research

ICM Prevention & Health
Education centre « Epidaure »



ICM Supportive Care
Department



Laboratory of Psychology



Desbrest Institute of
Epidemiology & Public Health



Two major research institutes in MIPS

newly integrated into the SIRIC Montpellier Cancer



**Laboratory of Computer Science, Robotics and
Microelectronics of Montpellier**



Laboratory of Mechanics and Civil Engineering



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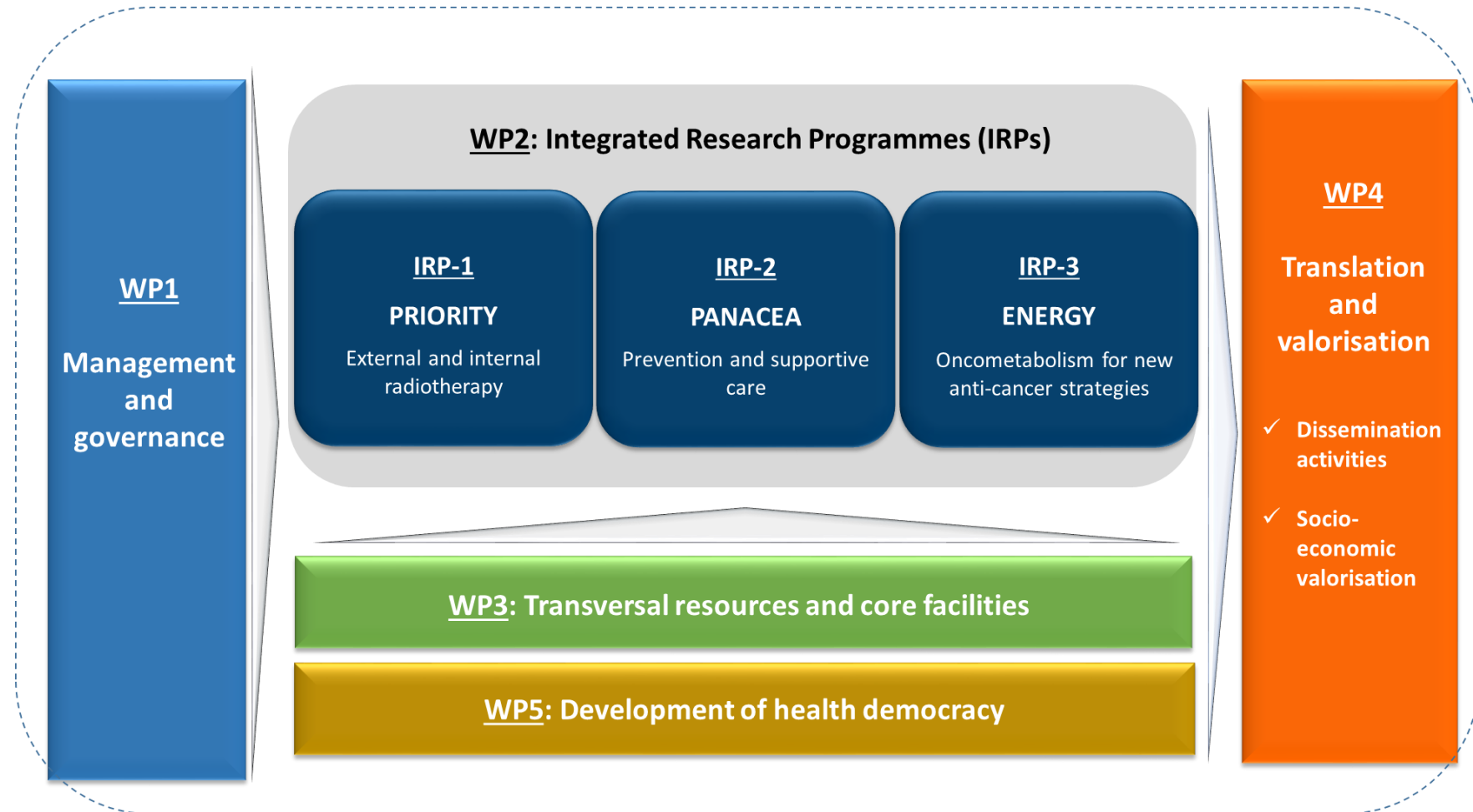
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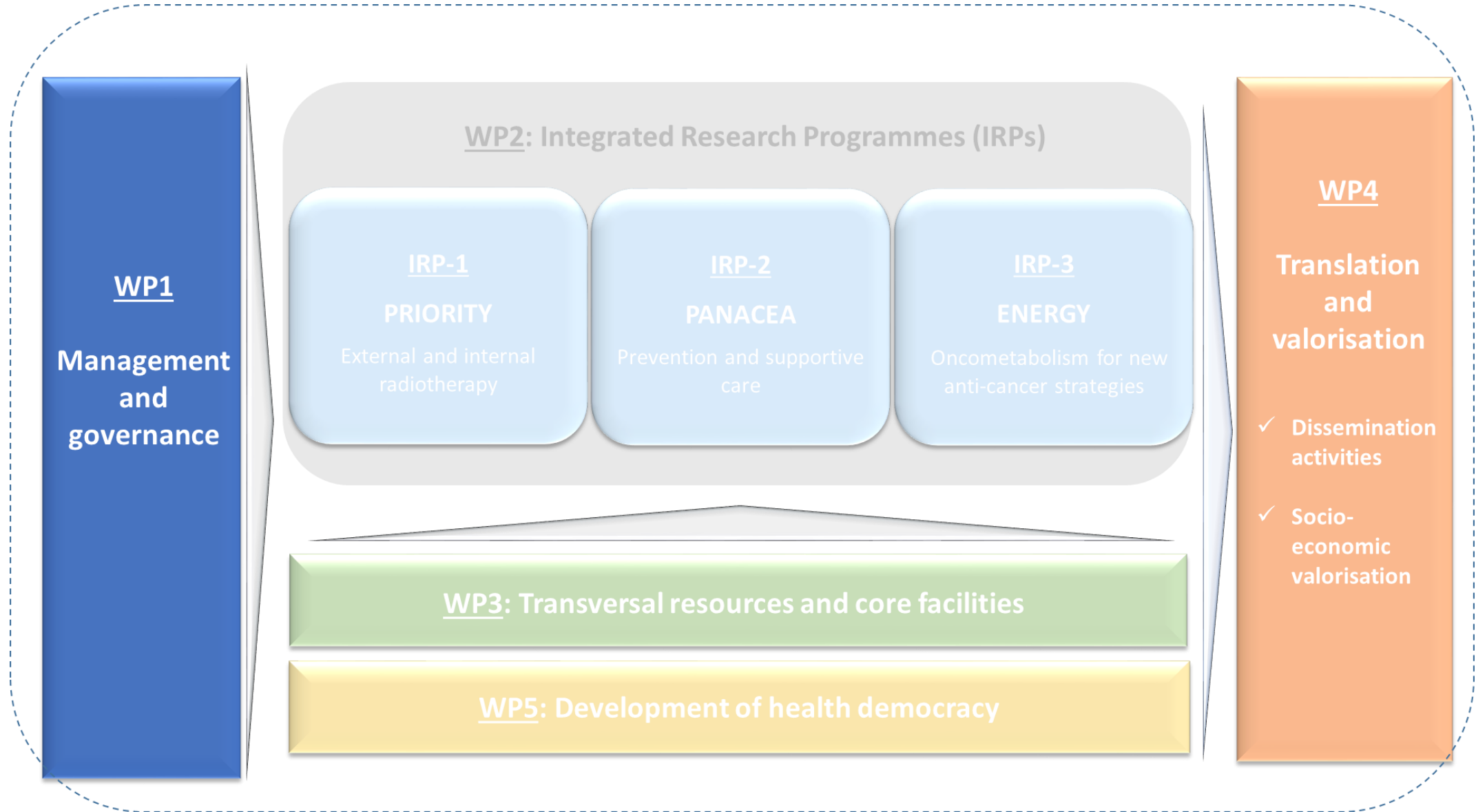
SIRIC 3 in the new 5-year strategic action plan

« Practice-changing » SIRIC

To change the patient's care pathway by a multilevel approach

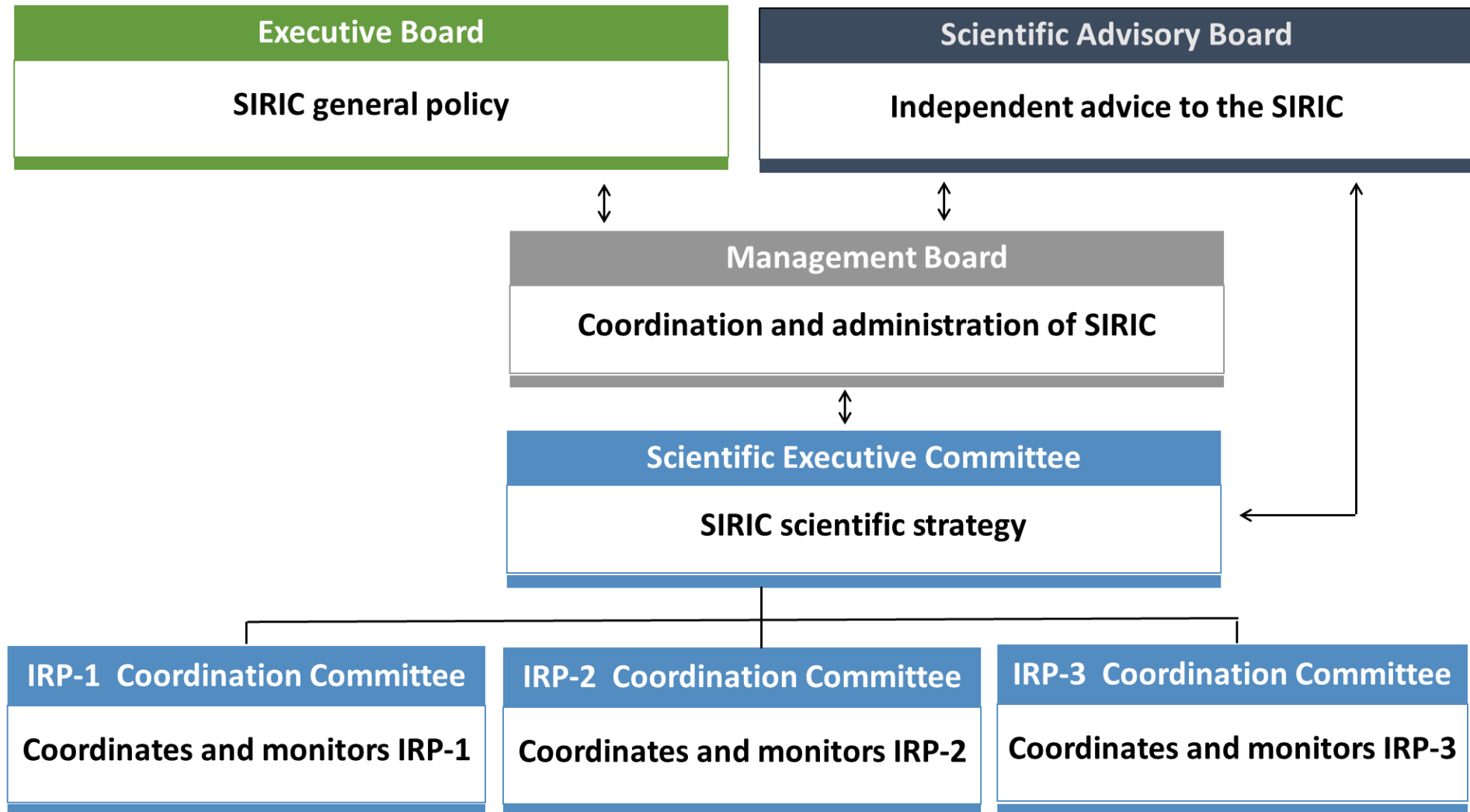


Strategic action plan



WP1 A similar and robust governance and managerial organisation

SIRIC Montpellier Cancer organisational chart



Scientific Advisory Board

Connections with international top-notch research institutes

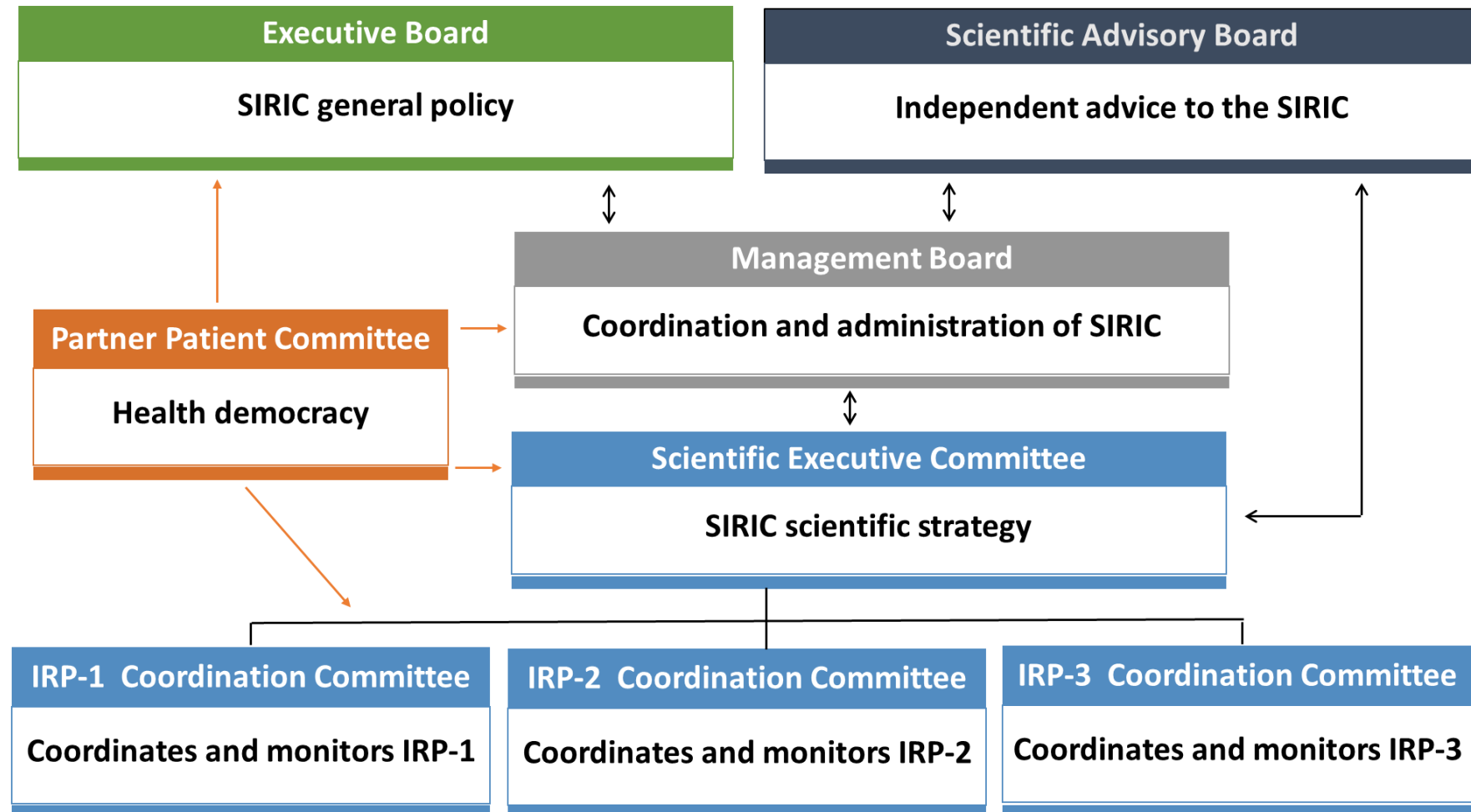


8 internationally leading scientists

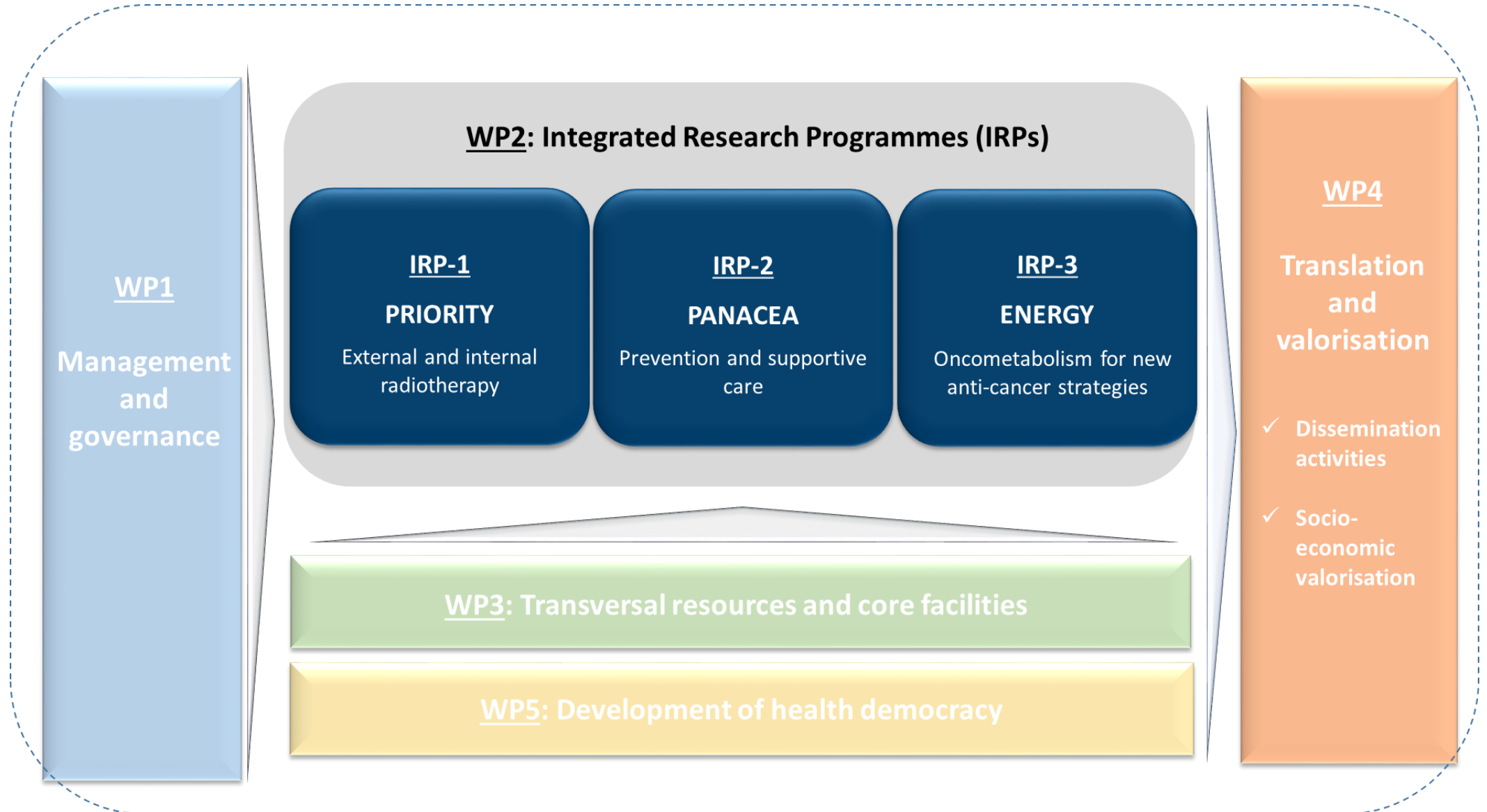
- **Vickie Baracos**, Palliative Care Medicine, University of Alberta (Canada)
- **Christian Frezza**, Metabolomics in Aging, University of Cologne (Deutschland)
- **Marie Johnston**, Health Psychology, University of Aberdeen (Scotland)
- **Florian Lordick**, Medical Oncology, University Cancer Center Leipzig (Deutschland)
- **Esat Mahmut Özşahin**, Radiation Oncology, University Hospital Lausanne (Switzerland)
- **Kevin Prise**, Radiation Biology, Belfast Queen's University (Ireland)
- **Hebert Alberto Vargas**, Medical Imaging, Memorial Sloan Kettering Cancer Center, New York (US)
- **Mariia Yuneva**, Oncogenes and tumor metabolism, Francis Crick Institute, London (UK)

WP1 A similar and robust governance and managerial organisation

SIRIC Montpellier Cancer organisational chart



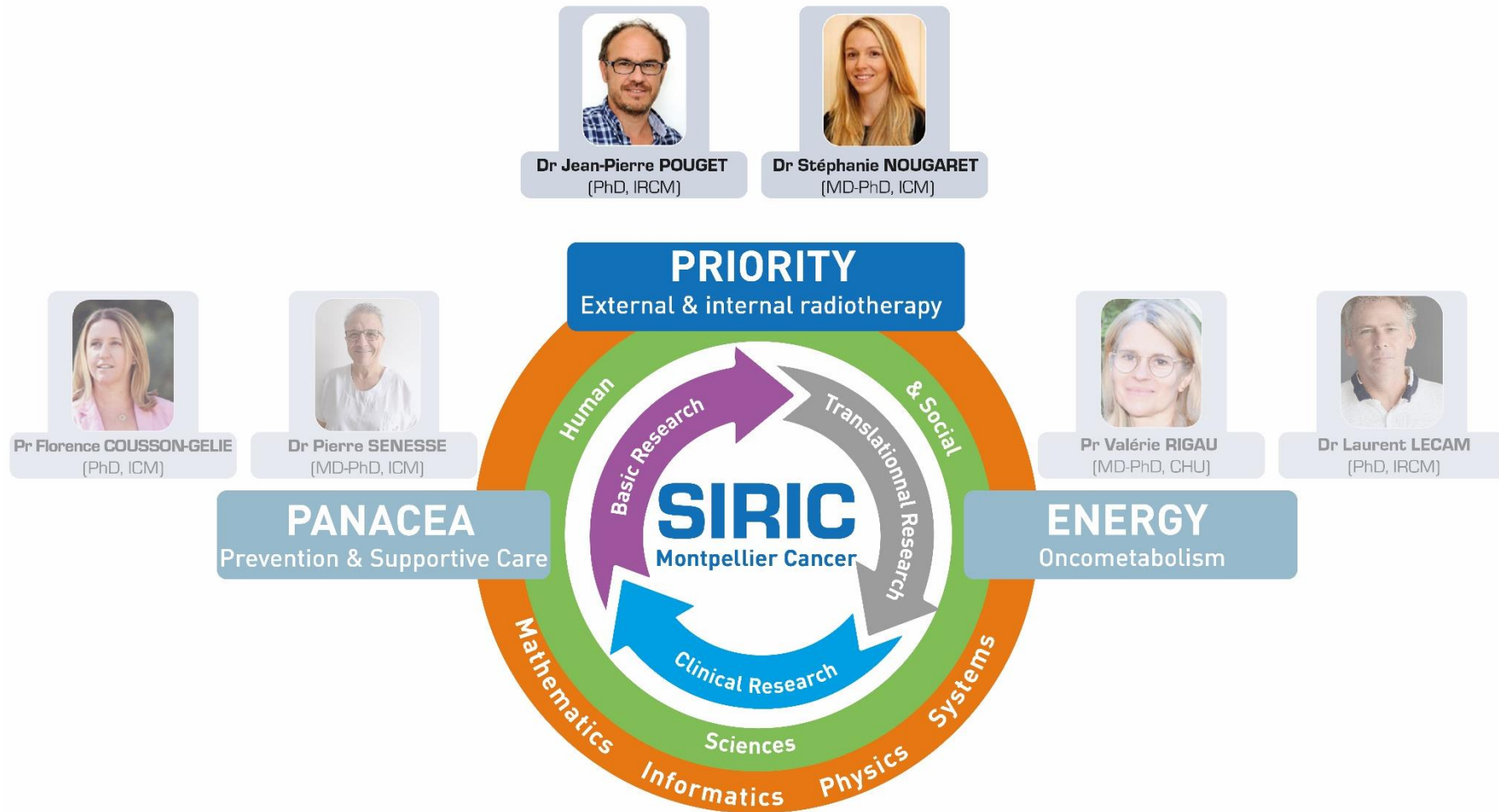
Strategic action plan



We selected 3 IRPs to change the patient's care pathway by a multilevel approach



We selected 3 IRPs to change the patient's care pathway by a multilevel approach



IRP-1

Integrating imaging, biology and dosimetry to optimize RIT and EBRT

Unique research integrating network of leaders in ionising radiation treatments



Unique INSERM group in France including radiation sciences, imaging and cancer biology

Radiobiology, radionuclide therapy (alphatherapy), external RT, radiophysics, dosimetry, immunology, TME, imaging



Multidisciplinary clinical expertise

Innovative radiation therapy, reference centres for PDAC and HCC, world leading expertise in MRI, clinico-biological databases of PDAC and HCC



A network of experts in MIPS

Computational science, AI, thermodynamics of materials



Cutting-edge core facilities



Preclinical models



Experimental radiotherapy



Genomics proteomics



Imaging Mass Cytometry



Small animal imaging

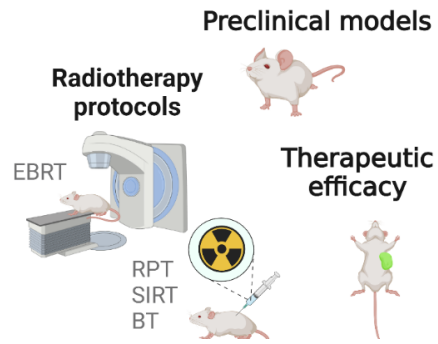


Experimental histology



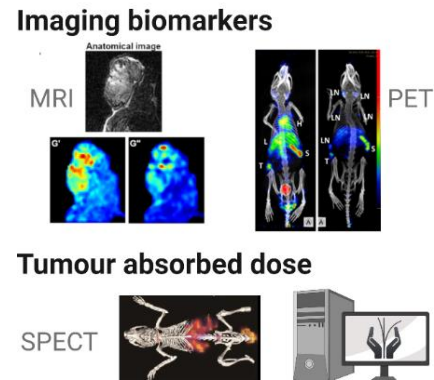
Four interconnected research axes

1 - Optimise irradiation of PDAC and HCC using preclinical models



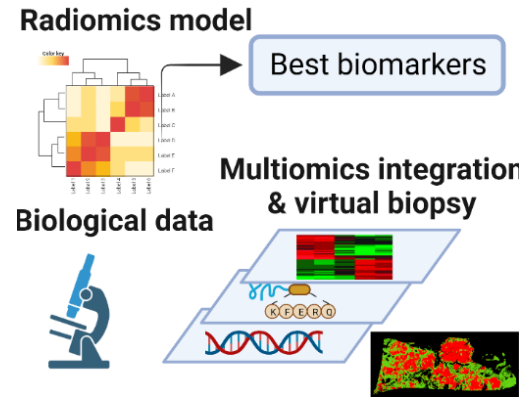
- **Tasks 1-2:** Establish **preclinical models and irradiation protocols** of PDAC and HCC
- **Task 3:** Evaluate **therapeutic efficacy and toxicity of EBRT and Alphatherapy**

2 - Acquire multimodal and multiparametric images during irradiation



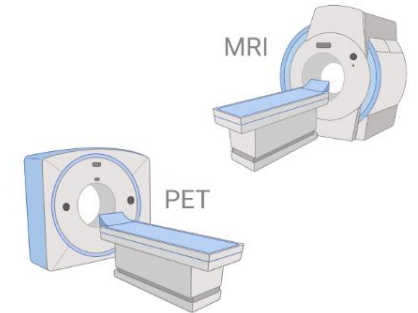
- **Tasks 1-8:** Measure a set of **tumour features by imaging**
- **Task 9:** Assess the **tumour absorbed dose** during irradiations

3 - Develop and implement a predictive model



- **Task 1:** Establish the **predictive radiomics model**
- **Tasks 2-3:** Extract **imaging & biological data**
- **Tasks 4-5:** Multi-omic and multi-imaging integration for **virtual biopsies**

4 - Transfer the predictive model into the clinic



- **Task 1:** Implement the MRI parameters into clinical machines
- **Task 2:** Implement the predictive model in **clinical MRI**
- **Task 3:** Develop new PET radioligands for monitoring tumour response to irradiations

We selected 3 IRPs to change the patient's care pathway by a multilevel approach



Changing preventive and supportive care strategies to reduce at-risk behaviours and treatment complications

From cancer primary prevention to post-treatment rehabilitation



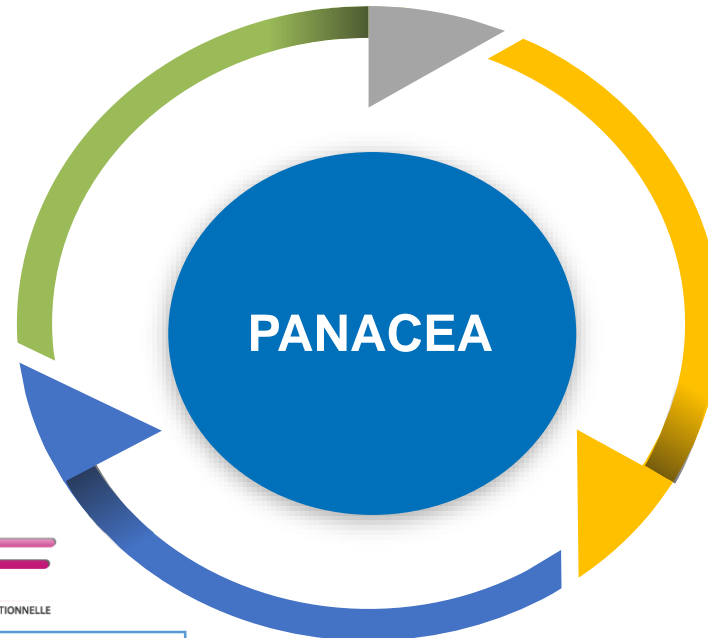
A unique cluster of HSS researchers

Primary prevention, health education, onco-psychology, behavioural sciences, sport sciences, public health



Multidisciplinary clinical expertise

Supportive care, clinical nutrition, onco-psychology, adapted physical activity, addictology, medical cancer genetics



Cancer biology

Circulating biomarkers, oncometabolism, epigenetics

1 - Adapt and disseminate theory-based interventions of cancer prevention



Task-1: Towards the general population

- In school children to reduce sun exposure
- In adults at work to promote physical activity



Task-2: Towards a cancer genetic-risk population

- Transfer theoretical tools of cancer prevention to target populations



2 - Introduce early supportive interventions to improve clinical benefit in cancer patients



Task-1: In GI cancers

- Early nutrition interventions
- Biomarkers of cancer cachexia

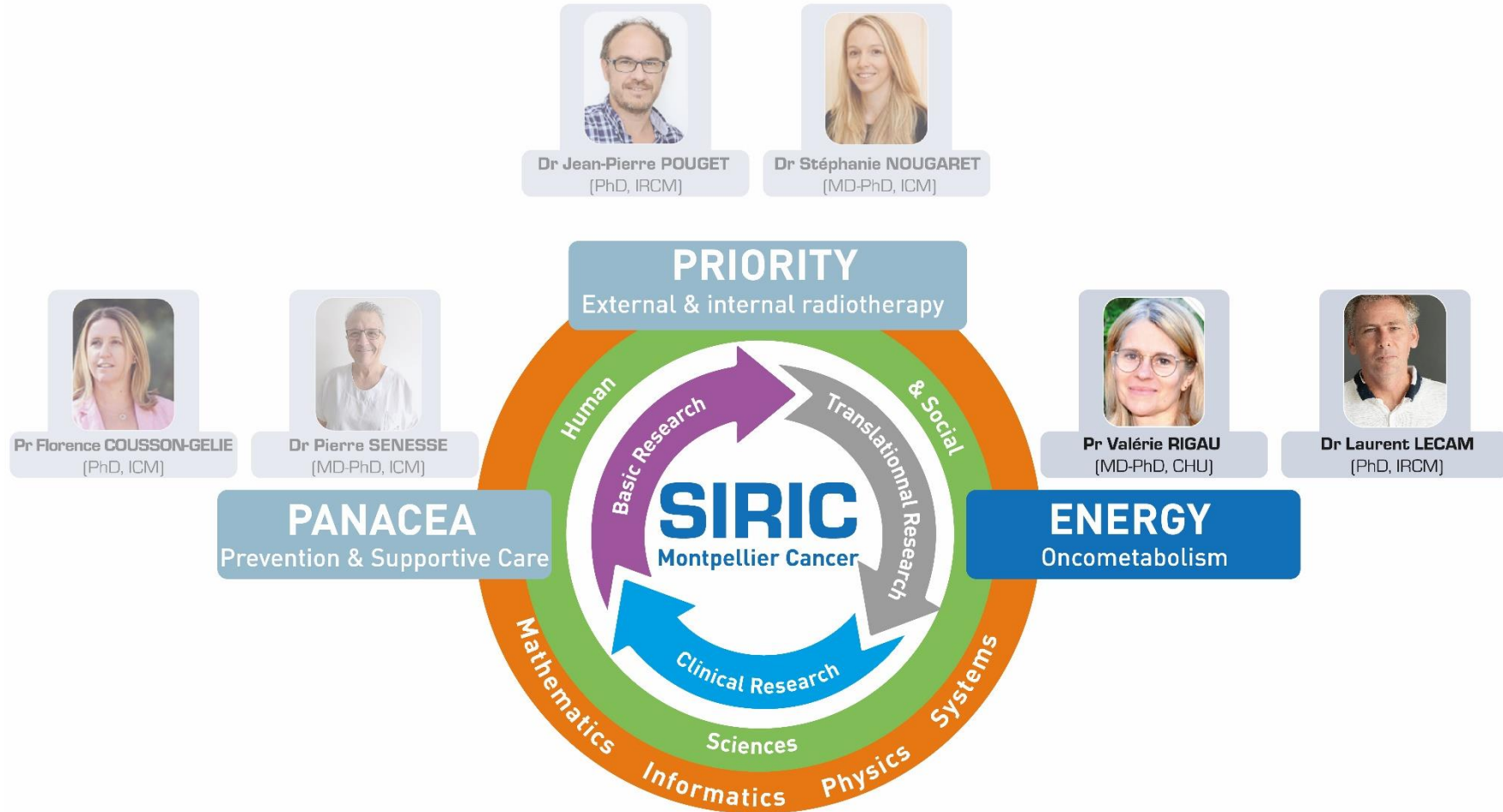
Task-2: In brain tumors

- To evaluate the effect of a mixed neuropsychological rehabilitation intervention in DLGG patients

Task-3: A digital application in oral therapies

- To help health professionals to prevent deleterious interactions between oral cancer therapies and complementary medicines

We selected 3 IRPs to change the patient's care pathway by a multilevel approach



Deciphering tumour metabolic cross-talks to design new anti-cancer therapies

Teams with international reputation in disciplines at the interface of metabolism and cancer biology



World-class research in cancer biology

Oncometabolism, epigenetics, epitranscriptomics, tumour microenvironment, immunity



Advanced computational sciences

Bioinformatics, machine learning, data modeling



Multidisciplinary clinical expertise

- Cutaneous melanomas, liposarcomas, gliomas, GI cancers, hemopathies
- Clinico-biological databases

Cutting-edge core facilities



Preclinical models



Metabolomics



Epitranscriptomics



Imaging Mass Cytometry



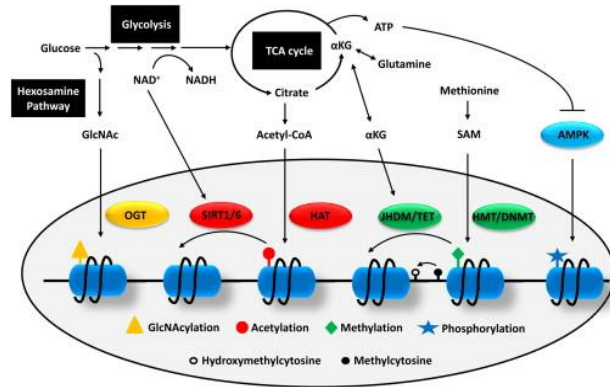
Antibody engineering



Organoids

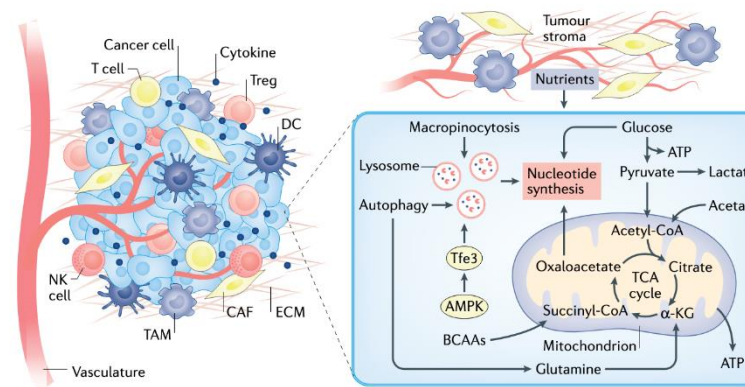
Three interconnected research axes

1 – Characterize molecular mechanisms linking metabolic changes to the epigenome and epitranscriptome



- **Task 1:** Identify new biomarkers of therapeutic responses and drug resistance in cancers with defined metabolic alterations
- **Task 2:** Identify new therapeutic strategies targeting metabolic and epigenetic changes

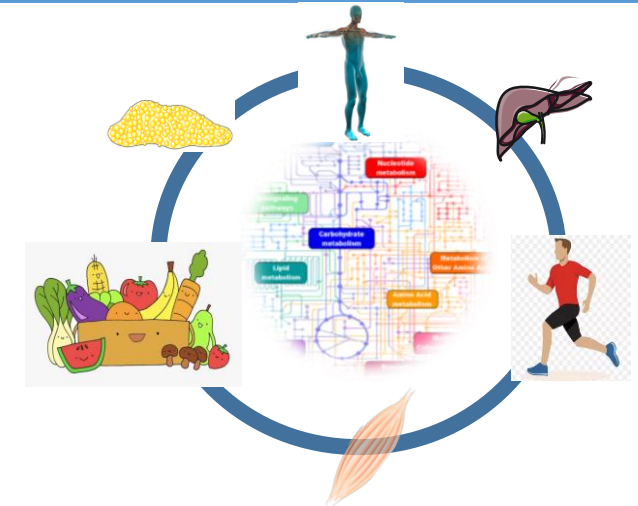
2 – Investigate how metabolic changes contribute to the cellular heterogeneity in the tumour ecosystem and to drug response



From Martinez-Reyel et al. Nat. Rev Cancer 2021

- **Task 1:** Evaluate the links between metabolic heterogeneity and therapeutic responses
- **Task 2:** Evaluate the impact of metabolic changes on the efficacy of immunotherapies

3 - Understand the metabolic crosstalks between tumour cells and their host

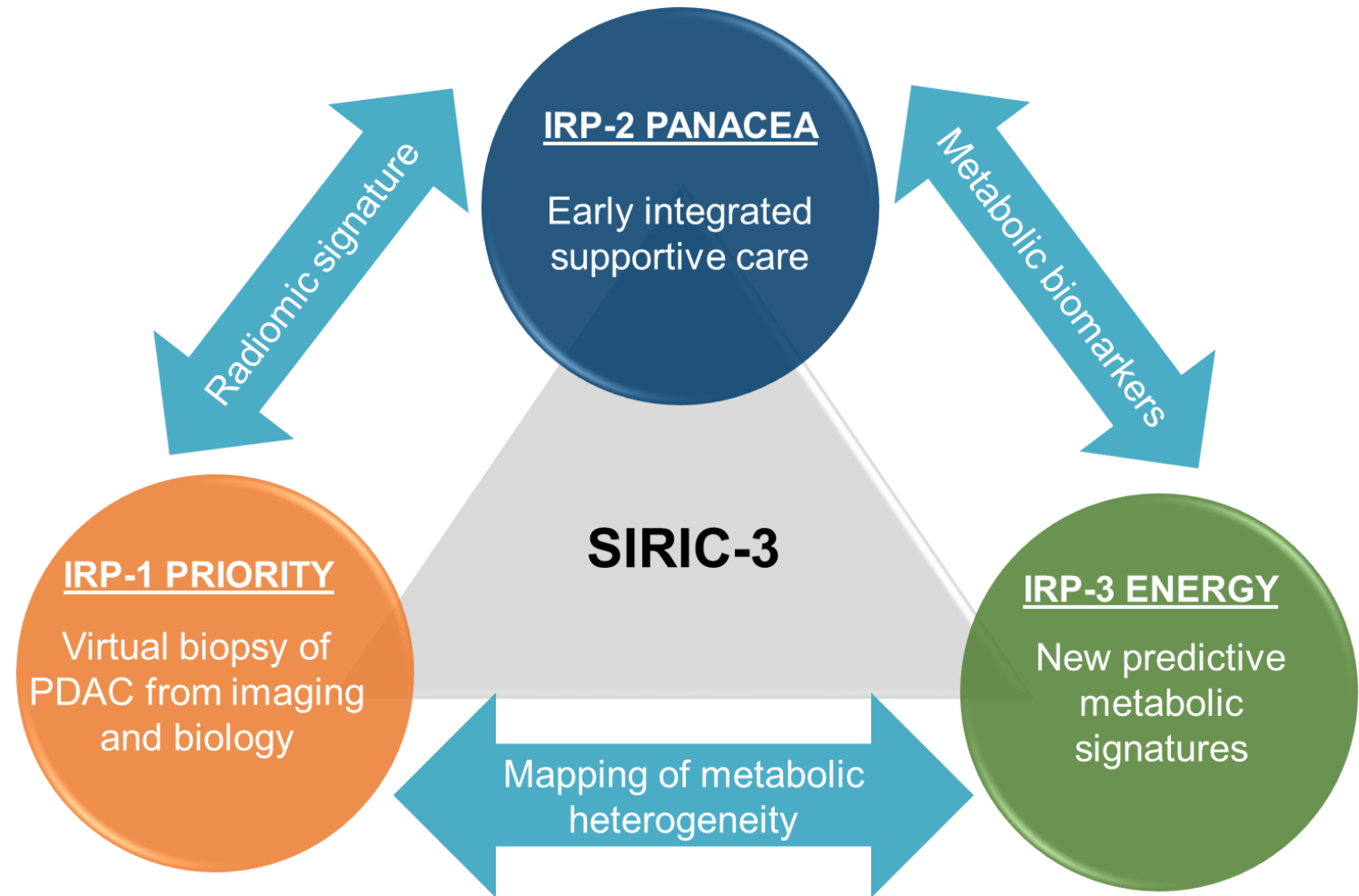


- **Task 1:** Investigate new mechanisms by which cancer cells reprogram the metabolism of distant organs
- **Task 2:** Explore the effects of specific diets and adapted exercising on the metabolic reprogramming of cancer cells

Example of cross-programmes synergies



Biobank with clinical data



Change the patient's care pathway of pancreas cancer by a multilevel approach

Strategic action plan

WP1
Management
and
governance

WP2: Integrated Research Programmes (IRPs)

IRP-1
PRIORITY
External and internal
radiotherapy

IRP-2
PANACEA
Prevention and supportive
care

IRP-3
ENERGY
Oncometabolism for new
anti-cancer strategies

WP3: Transversal resources and core facilities

WP5: Development of health democracy

WP4
Translation
and
valorisation

- ✓ Dissemination activities
- ✓ Socio-economic valorisation

WP3 Optimise transversal resources and core facilities for IRPs



An existing tool-box for preclinical & translational research



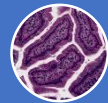
Preclinical models



Biological resources



Preclinical imaging



Experimental Histology



Experimental radiotherapy



Liquid biopsy



Imaging Mass Cytometry



Antibody engineering



Priority actions as part of SIRIC-3

Spatial molecular imaging

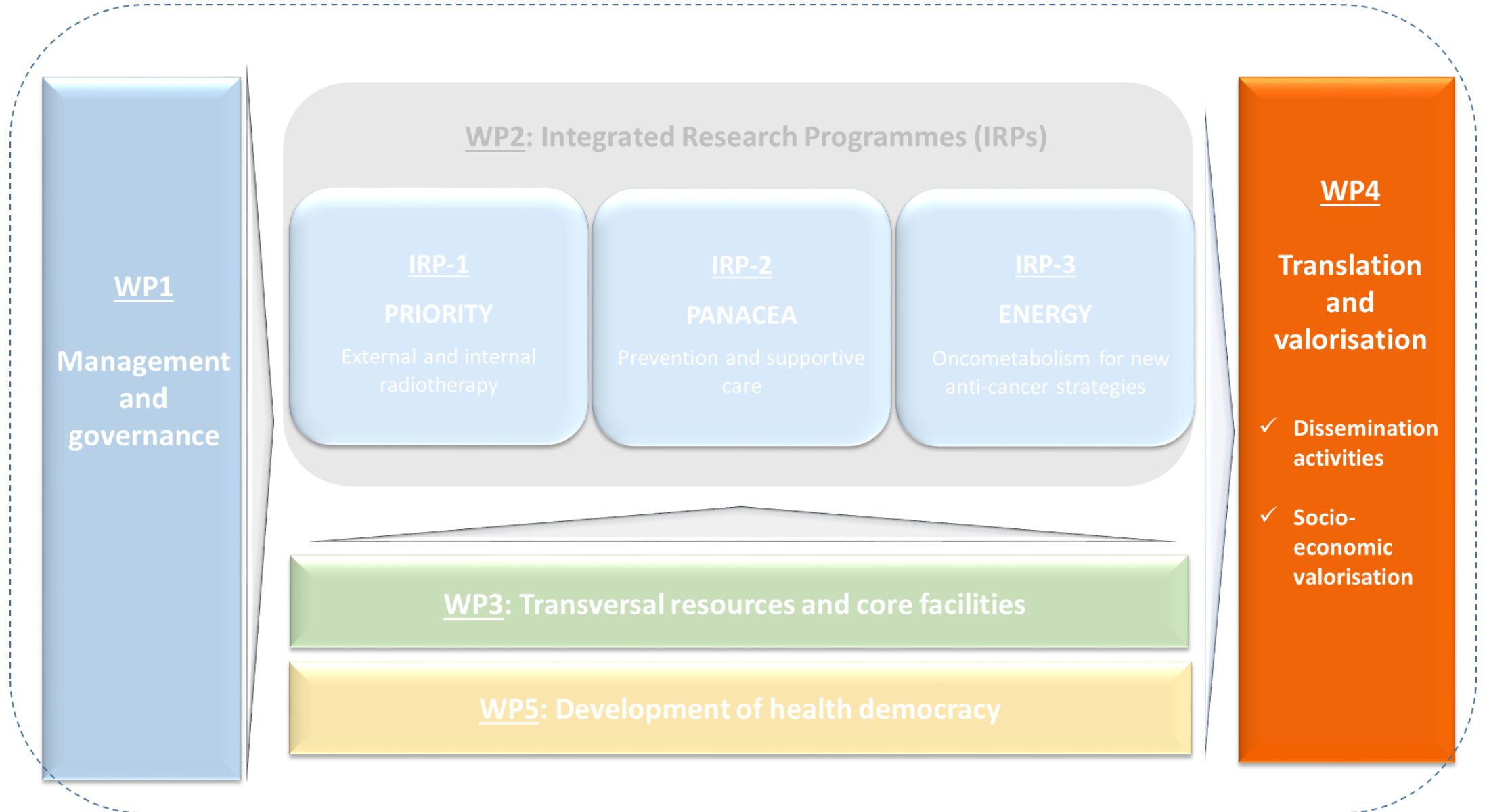
Experimental 3D-guided radiotherapy

α -particles emitters therapy with radiomics

AI-based radiomics platform for MRI



Strategic action plan



WP4

Promote knowledge and practices dissemination

□ For professionals

Via academic networks

- Onco-Occitanie, expert networks



Promoting High-level scientific events

- Montpellier Oncology Innovation Symposium (6-7 Jul-2023)
- 10th International Symposium on Auger Processes (6-8 Sept-2023)
- National Congress on Cancer Prevention (Sept-2023)
- Annual SIRIC Meetings



Improving training and education

- **Masters** : cancer biology, health psychology, Radio-TRANSNET
- **University Diplomas**: translational research, supportive care, nutrition in oncology



□ For patients and the general public

- **Epidaure**: a unique tool for prevention and health education (Via Occitanie Agency of Health based in Montpellier)
- **Federating a network of patient associations**
- **Creating educational videos, booklets, events**

<http://montpellier-cancer.com>



WP4

Boost tech transfer and industrial partnerships

Relies on a powerful innovation ecosystem in oncology

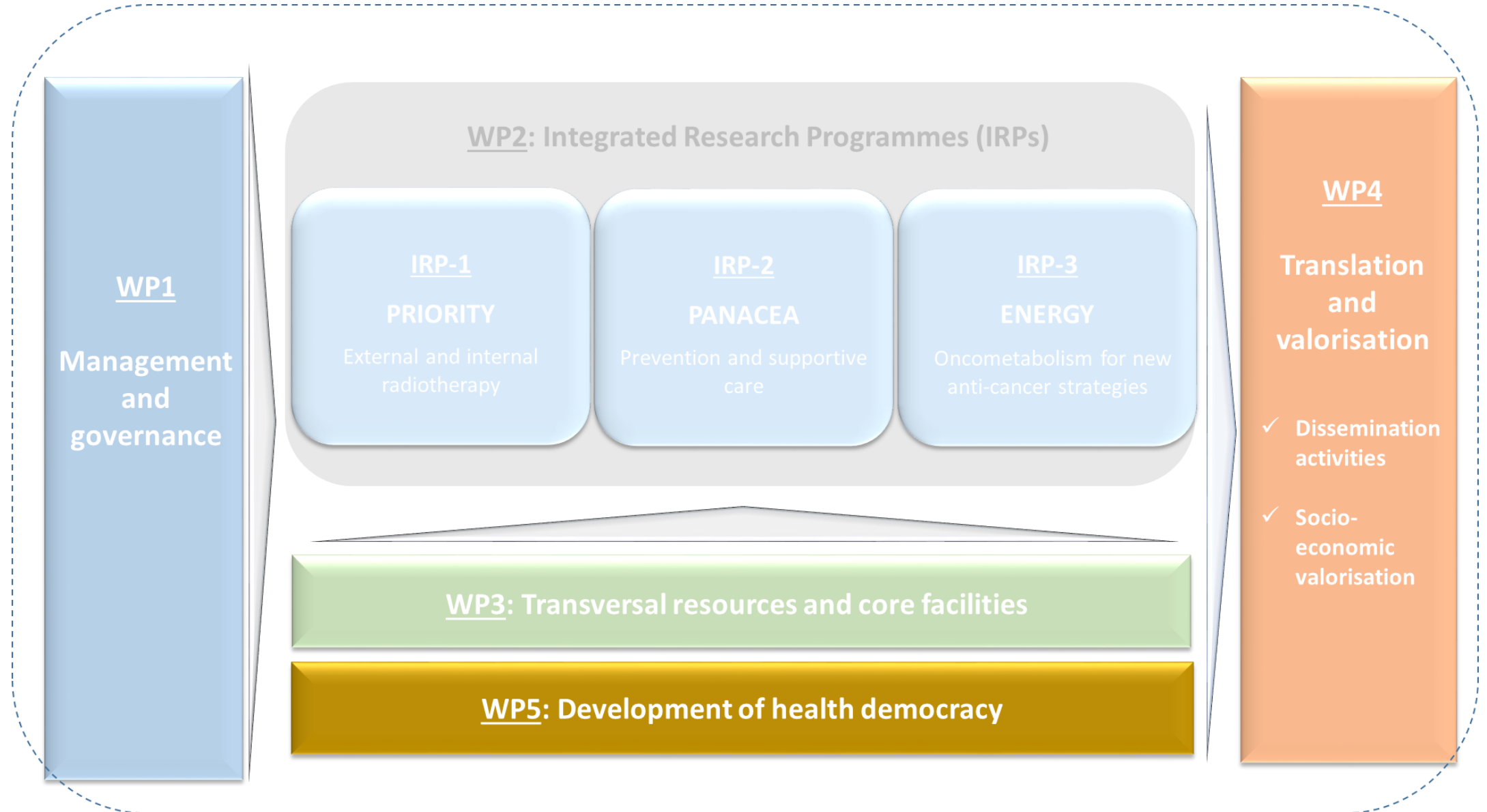


- A broad spectrum of industry partnerships
- A strong commitment to host companies on campus
- Network of efficient technology transfer tools
- Oncology Innovation Transfer Centre (CTIO, by 2024)

A fertile ground for the dissemination and exploitation of SIRIC research



Strategic action plan



WP5

Pursue health democracy development

2 partner patients to coordinate SIRIC health democracy policy



Maguy Del Rio

Coordinator patient at ICM and
new member (nov 2022) of the
INCa national committee



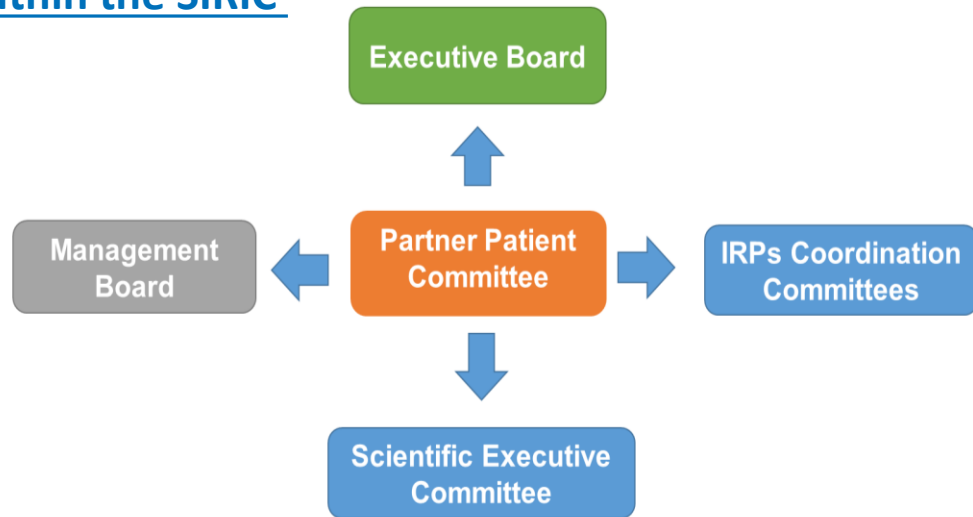
Cyril Sarrauste de Menthère

“Mon Réseau Cancer colorectal” Association
Executive Board of the “Digestive Cancers Europe”
Association



Guarantee the integration of the « patient experience » into SIRIC projects

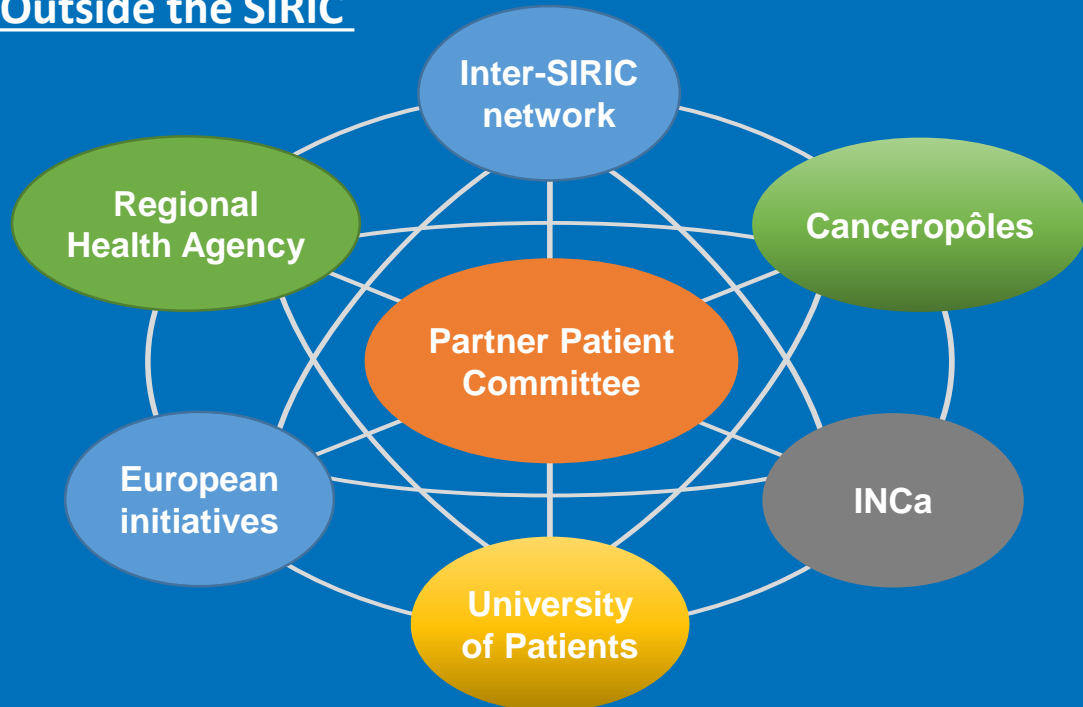
Within the SIRIC



Targeted actions

- Set-up a **Partner Patient Committee**
 - ❖ At all levels of SIRIC actions
 - ❖ Linked to regional, national and international initiatives
- Develop the **patient-researcher partnership** through IRPs
- Involve patients in **knowledge dissemination strategy**
- Lead an **inter-SIRIC network of partner patients**

Outside the SIRIC



Inter-SIRIC Colorectal Cancer Meeting (March 2022, Montpellier)

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Partner institutions

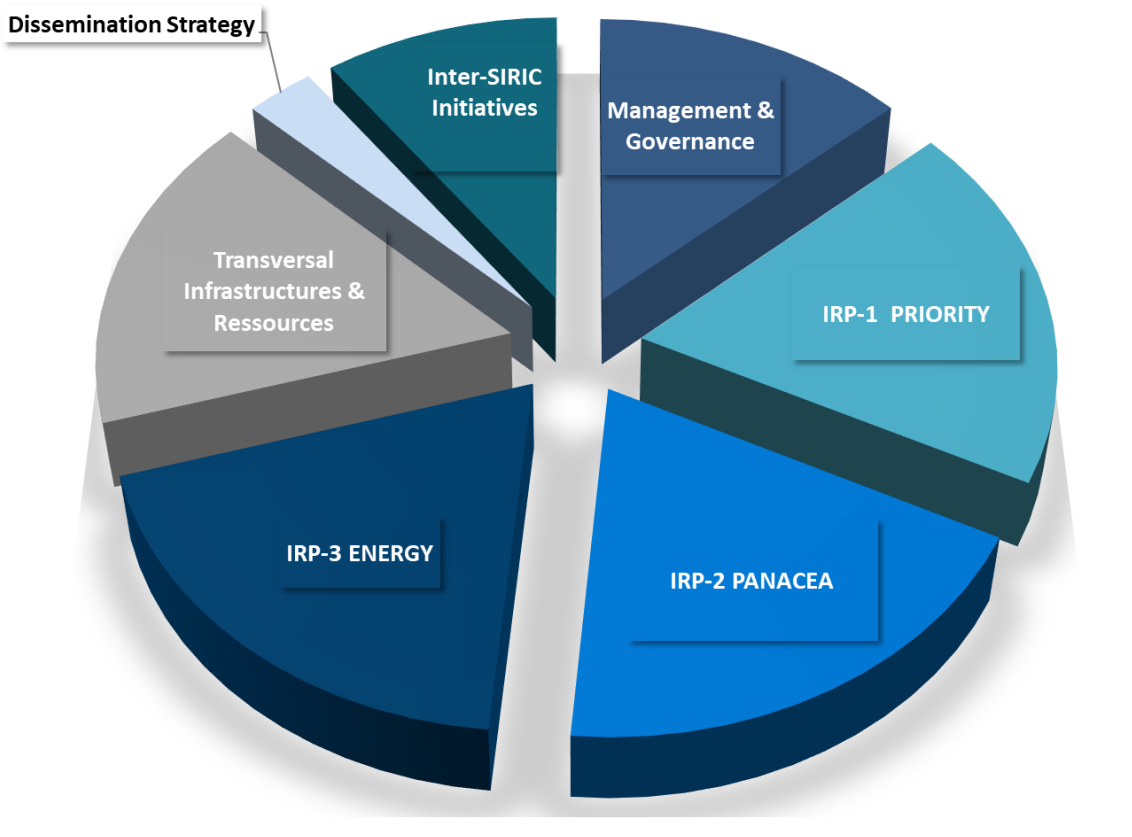
The new strategic action plan

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Provisional budget 2023-2027



- Grant centrally managed by ICM Finances Dept.
- **Total budget 6M€**
 - 3420k€ (57%) for the 3 Programmes
 - 1000k€ (17%) for transversal resources
 - 175k€ (3%) for dissemination activities
 - 600k€ (10%) for inter-SIRIC initiatives
 - 805k€ (13%) for management and general administration

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« Practice-changing » SIRIC

To change the patient's care pathway by a multilevel approach

Leverage a unique cluster of excellence in internal and external RT to provide **game-changing management of targeted irradiation treatments**

Establish a **highly competitive research network** targeting **oncometabolism as a source of new therapeutic strategies**



Capitalize on **strong expertise in HSS interventional research** to **demonstrate efficacy of prevention and early supportive care**

