

# Méthodes pour l'automatisation de la dosimetrie pour les traitements radiothérapiques.

Methods for automatization of the dosimetry for radiotherapy treatments.

#### Thèse de doctorat de l'université Paris-Saclay

Spécialité de doctorat: ... École doctorale n° 573 Interfaces : matériaux, systèmes, usages, ED INTERFACE Graduate School: Sciences de l'Ingénierie et des Systèmes, SIS

Thèse préparée dans les unités de recherche Radiothérapie (Institut Régionale du Cancer de Montpellier), Advanced Research (TheraPanacea), et MICS, Mathématiques et Informatique pour la Complexité et les Systèmes (Université Paris-Saclay, CentraleSupélec), sous la direction de Nikos Paragios, Professeur, et la co-direction de Paul-Henry Cournède, Professeur

Thèse soutenue à Paris-Saclay, le JJ mois AAAA, par

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#### Composition du jury

Membres du jury avec voix délibérative

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Rapporteur & Examinateur / trice

Examinateur ou Examinatrice

Examinateur ou Examinatrice



Titre: Méthodes pour l'automatisation de la dosimetrie pour les traitements radiothérapiques.

Mots clés: Mathématiques, Intelligence Artificielle, Radiothérapie

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Title: Methods for automatization of the dosimetry for radiotherapy treatments.

Keywords: Mathematics, Artificial Intelligence, Radiotherapy

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# Acknowledgments

A PhD is more than just hard work; it thrives on mentorship, collaboration, and unwavering support. [...]

# List of Contributions

- Teaching: Consistency and Reproducibility of Grades in Higher Education: A Case Study in Deep Learning replace icon
- ArXiV: Radiotherapy Dosimetry: A Review on Open-Source Optimizer
- ESTRO: A Novel Framework for Multi-Objective Optimization and Robust Plan Selection Using Graph Theory
- SFPM: Dose Volume Histograms Guided Deep Dose Predictions
- AIME: Radiotherapy Dose Optimization via Clinical Knowledge Based Reinforcement Learning (full paper coming soon)
- ASTRO: Clinically Dependent Fully Automatic Treatment Planning System
- SFRO: Attention Mechanism on Dose-Volume Histograms for Deep Dose Predictions

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# Introduction

#### Abstract

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1.1.1	luction to Cancer
1.1.1	Cells proliferating
	•
	DNA messed up
	variety of cancer
	some safe
	some not safe
1.1.2	who is concerned?
1.1.3	risk factors
	environment impacts the probability of getting cancer
	living habits as well
	genetic impacts as well (e.g. "cancer gene")
1.1.4	possible treatments
	surgery
	RT
	chemotherapy
	combination
1.1.5	10 cancer markers
	cell proliferation
	reprogram cellular metabolism
	stop cell growth arrest
	evade apoptosis
	escape immune system
	ability to undergo a sufficient number of successive cell cycles of growth and division to
	generate macroscopic tumors
	create new blog vessels to get nutriments
	allow cell escape and metastasis formation
	change cellular response phenotypic via plasticity
	senescence
	cancer can be considered as a living thing on its own
	beyond the cellular level, impacting tissues
110	
1.1.6	4 cancer conditions
	mutation
	epigenetic reprogramming
	inflammatory context
	disruption of microbiota
1.1.7	phases of cancer
	initiation
	promotion
	$tumorigenesis + neoangiogenesis \dots $
	evolution (local, regional, metastasis)
1.1.8	cancer classification:
	tumor, node, mestastasis
	stages classification:
	stages classification:  stage 0 which corresponds to a so-called in situ tumor  stage 1 which corresponds to a single, small tumor

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	image generation	
	text generation	
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	healthcare	22

### 1.1 Introduction to Cancer

#### 1.1.1 what is cancer?

Cells proliferating

DNA messed up

variety of cancer

some safe

 $(e.g.:\ mole/freckle)$ 

some not safe

worse make the human die

#### 1.1.2 who is concerned?

more and more ppl

#### 1.1.3 risk factors

environment impacts the probability of getting cancer

(e.g.: UV exposure)

living habits as well

(e.g.: smooking)

genetic impacts as well (e.g. "cancer gene")

#### 1.1.4 possible treatments

surgery

RT

chemotherapy

combination

#### 1.1.5 10 cancer markers

cell proliferation

reprogram cellular metabolism

stop cell growth arrest

evade apoptosis

escape immune system

ability to undergo a sufficient number of successive cell cycles of growth and division to generate macroscopic tumors

create new blog vessels to get nutriments

allow cell escape and metastasis formation

change cellular response phenotypic via plasticity

senescence

cancer can be considered as a living thing on its own

beyond the cellular level, impacting tissues

#### 1.1.6 4 cancer conditions

mutation

epigenetic reprogramming

inflammatory context

disruption of microbiota

#### 1.1.7 phases of cancer

initiation

promotion

tumorigenesis + neoangiogenesis

evolution (local, regional, metastasis)

#### 1.1.8 cancer classification:

tumor, node, mestastasis

stages classification:

stage 0 which corresponds to a so-called in situ tumor

stage 1 which corresponds to a single, small tumor

stage 2 which corresponds to a larger local volume

stage 3 which corresponds to invasion of the lymph nodes or surrounding tissues

stage 4 which corresponds to a wider extension in the body in the form of metastases

#### 1.1.9 cancer causes

various reasons why

environment

inherited mutations

mistake in DNA copy

#### 1.1.10 personalized treatments

revolution

rapid advances

help of mathematics

help of AI

## 1.2 Introduction to Mathematical Optimization

#### 1.2.1 optimization def

selection of a best element, with regard to some criteria

#### 1.2.2 in math: more precisely

optimization problem consists of maximizing or minimizing a real function by systematically choosing input

brute force

 ${\bf heuristics}$ 

1.2.3	notion of allowed set			
1.2.4	discrete vs continuous optim			
1.2.5	many real-world and theoretical problems may be modeled in continuous general framework			
1.2.6	$\max(\mathbf{f}) <=> \min(\mathbf{f}) \text{ hence only min}$			
1.2.7	notion of local vs global min			
1.2.8	feasibility			
1.2.9	existance			
1.2.10	optim algos			
1st ord	${ m er}$			
gradien	gradient descent			
line search				
quasi-n	ewton methods			
2nd ord	2nd order			
newton	newton's method			
0th ord	er			

- 1.2.11 least squares
- 1.2.12 multi-objective optimization
- 1.3 Introduction to Artificial Intelligence
- 1.3.1 quick def
- 1.3.2 general idea
- 1.3.3 common architectures

 $\mathbf{FC}$ 

MLP

CNN

RNN

transformers

- 1.3.4 Classic AI vs Learning AI
- 1.3.5 Machine learning vs Artificial Intelligence vs Deep Learning
- 1.3.6 applications
- 1.3.7 learning types

supervised

un-supervised

self-supervised

reinforcement / semi-supervised

1.3.8 tasks

classical tasks

regression

classification

partitioning

dimension reduction

generative AI

images => training is difficult

text

## 1.3.9 recent progress

computer vision

playing games

(a way to assess intelligence)

image generation

text generation

healthcare

# Radiotherapy

#### Abstract

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2.1	Patien	t Path
	2.1.1	Detection / diagnostic
	2.1.2	RT Prescription
	2.1.3	CT scan
	2.1.4	Contouring
	2.1.5	Treatment Planning
	2.1.6	Irradiation Sessions
	2.1.7	Follow-up
2.2	Machir	nes
	2.2.1	Molds / 3D-RT
	2.2.2	MLC-LINAC
	2.2.3	Tomotherapy
	2.2.4	CyberKnife
	2.2.5	Brachytherapy
2.3	Irradia	tions techniques
	2.3.1	IMRT
		Step and Shoot
		Sliding Window
	2.3.2	VMAT
2.4		nent Planning Systems
2.1	2.4.1	Manufacturer
	2.1.1	Eclipse (Varian)
		ONE   Planning (Elekta)
		Precision (Accuray)
	2.4.2	Non-manufacturer
	2.4.2	RayStation (RaySearch)
		matRad (German Cancer Research Center - DKFZ)
		AutoPlan (TheraPanacea - coming soon)
2.5	Dogim	etry steps
۷. ن	DOSIIII	Challenges
	2.5.1	BOO
	$\frac{2.5.1}{2.5.2}$	
0.0	2.5.3	LF
2.6	Simula	${ m tion}$

#### 2.1 Patient Path

- 2.1.1 Detection / diagnostic
- 2.1.2 RT Prescription
- 2.1.3 CT scan
- 2.1.4 Contouring
- 2.1.5 Treatment Planning
- 2.1.6 Irradiation Sessions
- 2.1.7 Follow-up
- 2.2 Machines
- 2.2.1 Molds / 3D-RT
- 2.2.2 MLC-LINAC
- 2.2.3 Tomotherapy
- 2.2.4 CyberKnife
- 2.2.5 Brachytherapy

#### 2.3 Irradiations techniques

#### 2.3.1 IMRT

Step and Shoot

Sliding Window

2.3.2 VMAT

## 2.4 Treatment Planning Systems

#### 2.4.1 Manufacturer

Eclipse (Varian)

ONE | Planning (Elekta)

Precision (Accuray)

#### 2.4.2 Non-manufacturer

RayStation (RaySearch)

matRad (German Cancer Research Center - DKFZ)

AutoPlan (TheraPanacea - coming soon)

## 2.5 Dosimetry steps

Challenges

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- 2.5.1 BOO
- 2.5.2 FMO
- 2.5.3 LF
- 2.6 Simulation