



Organ preservation for rectal cancer.

Should we use HDR or

Contact x-ray *CXB* boost ?

J.P. GERARD



Centre Antoine Lacassagne



Nice



Hôpitaux de Lyon

Leiden, 20th November 2017

15 mn

DECLARATION D'INTERET - DISCLOSURE

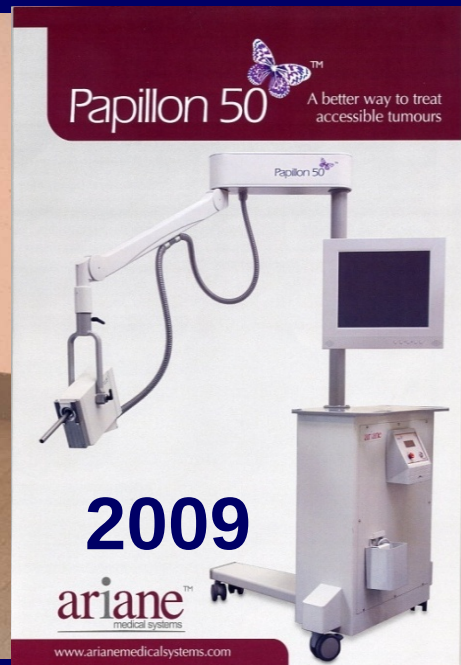
Ariane Medical Systems™ (UK)

Contact X Ray Brachytherapy 50 kV

"Papillon 50"

Philips
RT 50

1971



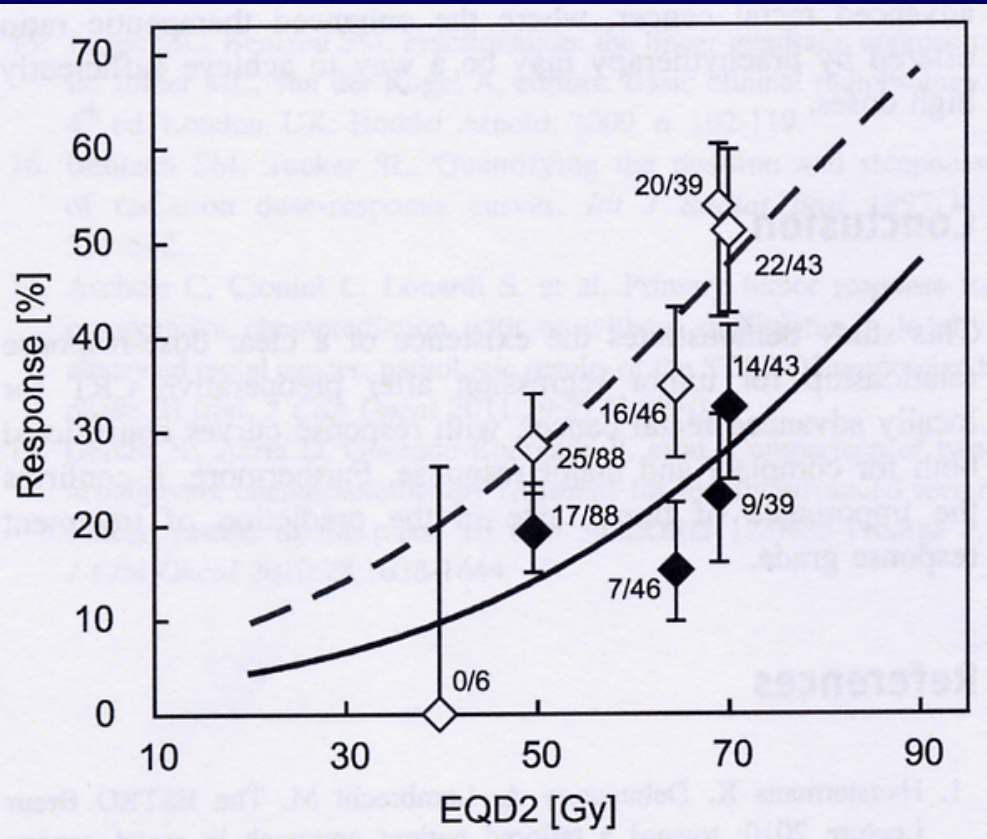
2009

J.P. GERARD

Medical Advisor

Rectal adenocarcinoma : radioresistant

Radiation Dose-Response Model (EQD 2)



T3 rectal ADK

for 50% pCR :

Dose = 92 Gy

Rectum Tolerance Dose

< 75 Gy

Rectum External Beam RT (EBRT) BOOST ??

EBRT : - stereotactic Body RT (SIB)

- cyberknife
- proton



ACCORD 12 phase III : 45 Gy vs 50 Gy **ypCR** : 13% - 19%

GRECCAR 12 « « : 50 Gy vs 60 Gy **R0** : 83% - 88%

But Toxicity is the ultimate Limit

With EBRT impossible to increase Dose > 80 Gy

Higher Tumor Dose
(no toxicity +++)

Endoluminal approach

Brachy RT *$(1/r^2)$*

FSD : Focus Surface Distance

Brachytherapy

1) **CXB**

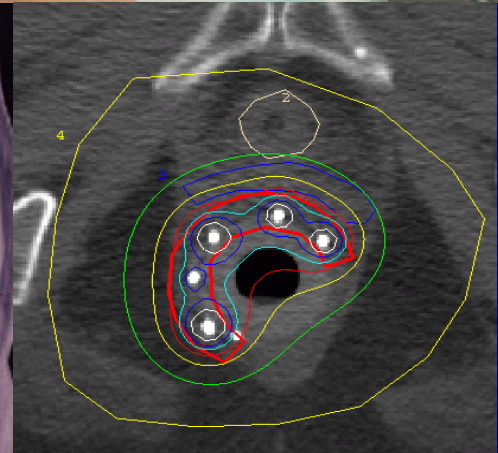
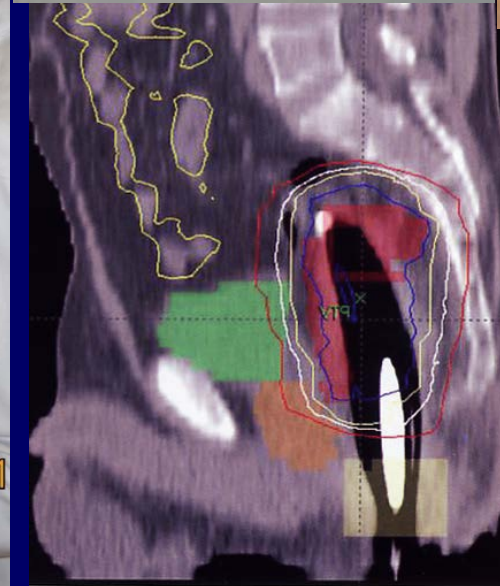
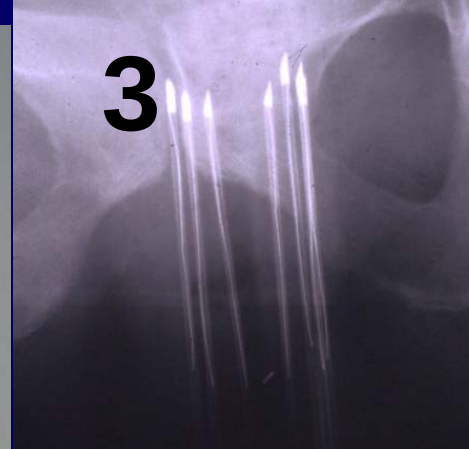
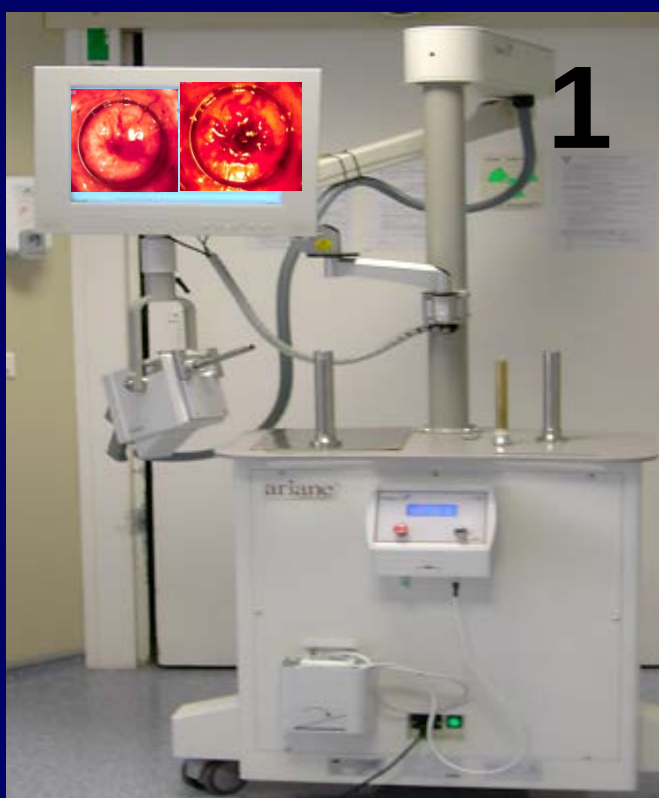
X50 kV

2) **Endoluminal**

HDR Ir192

3) **Interstitial**

HDR Ir192

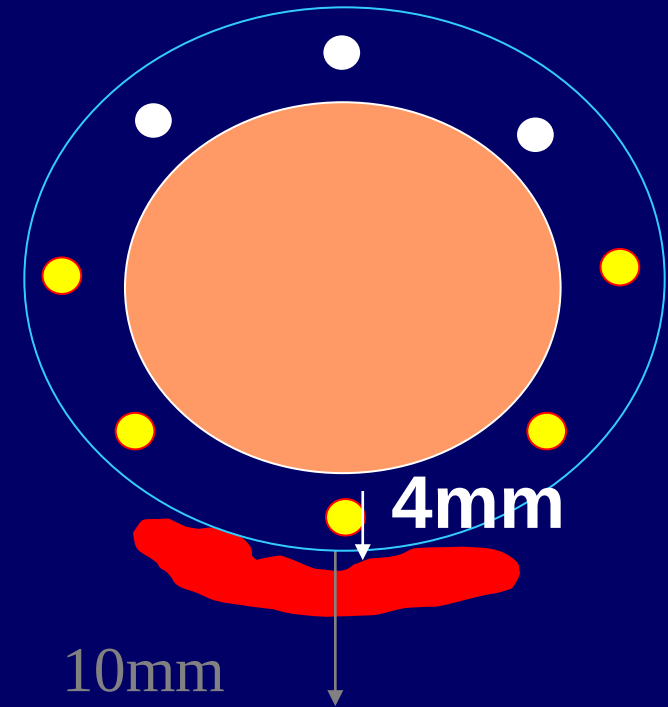


CXB > Iridium

- 1) Higher (*inhomogeneous*) dose in Tumor:**
- 2) Better accurate Targetting (*eye vs image guided*)**
- 3) Better protection of normal rectum**
- 4) Simple - radioprotection - Cheap**
- 5) Sustainable development (*no radioactive waste*)**

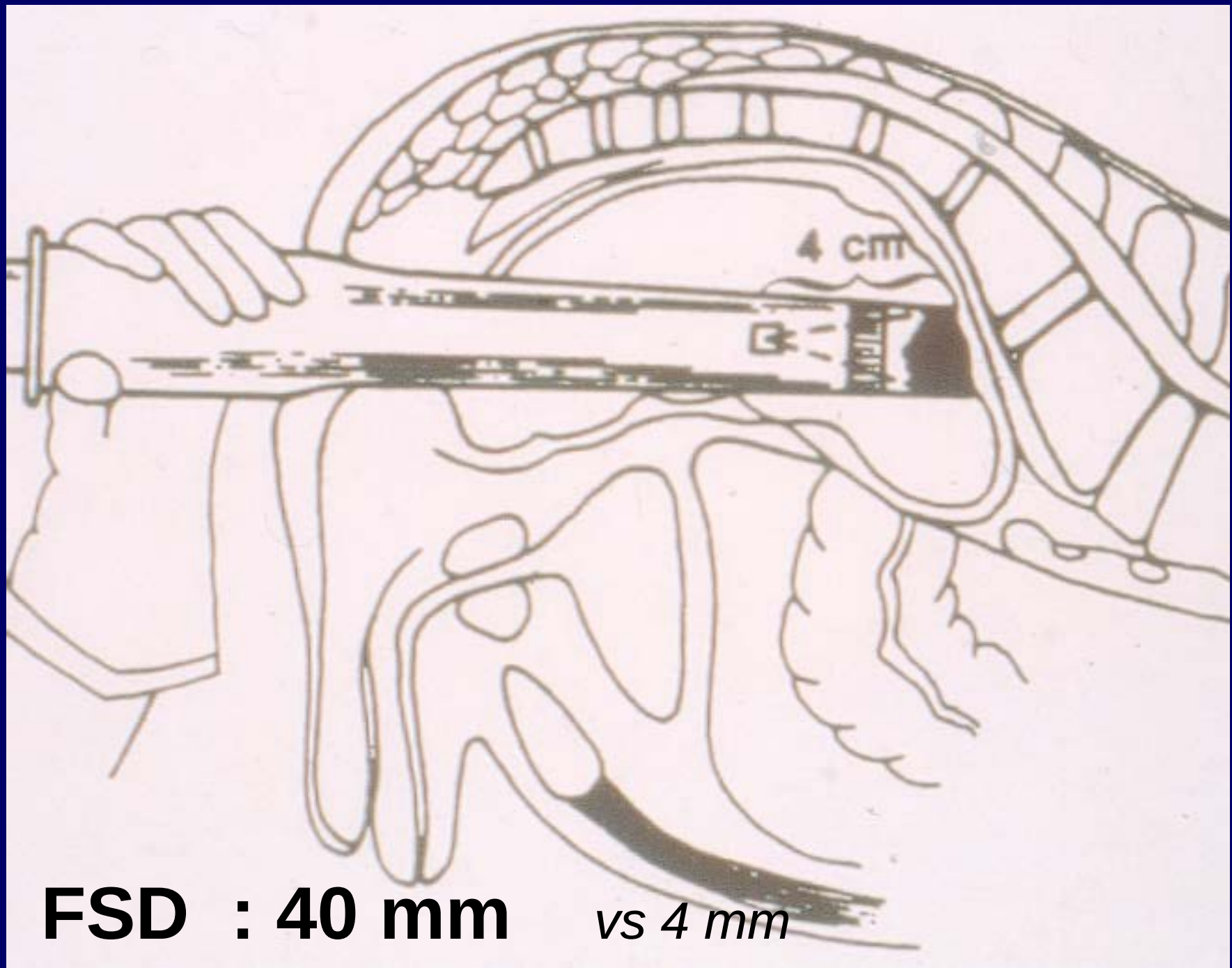
LIMITATIONS

Endoluminal Iridium 192 HDR



FSD : 4 mm

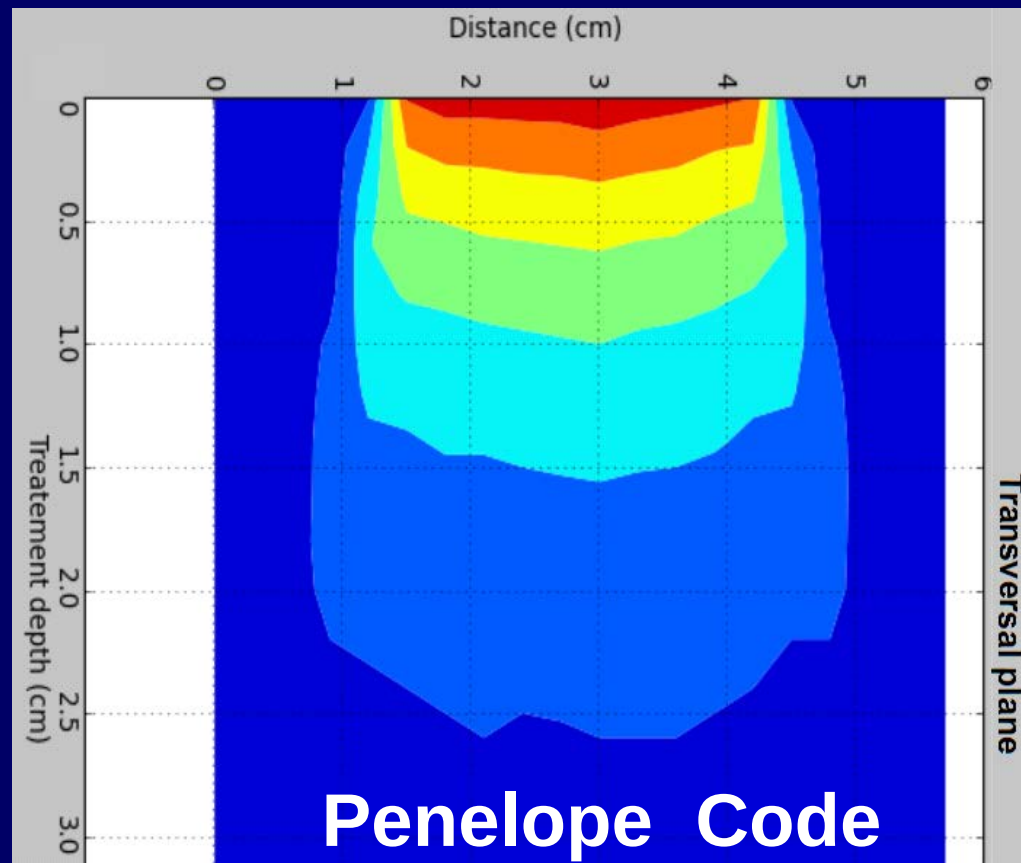
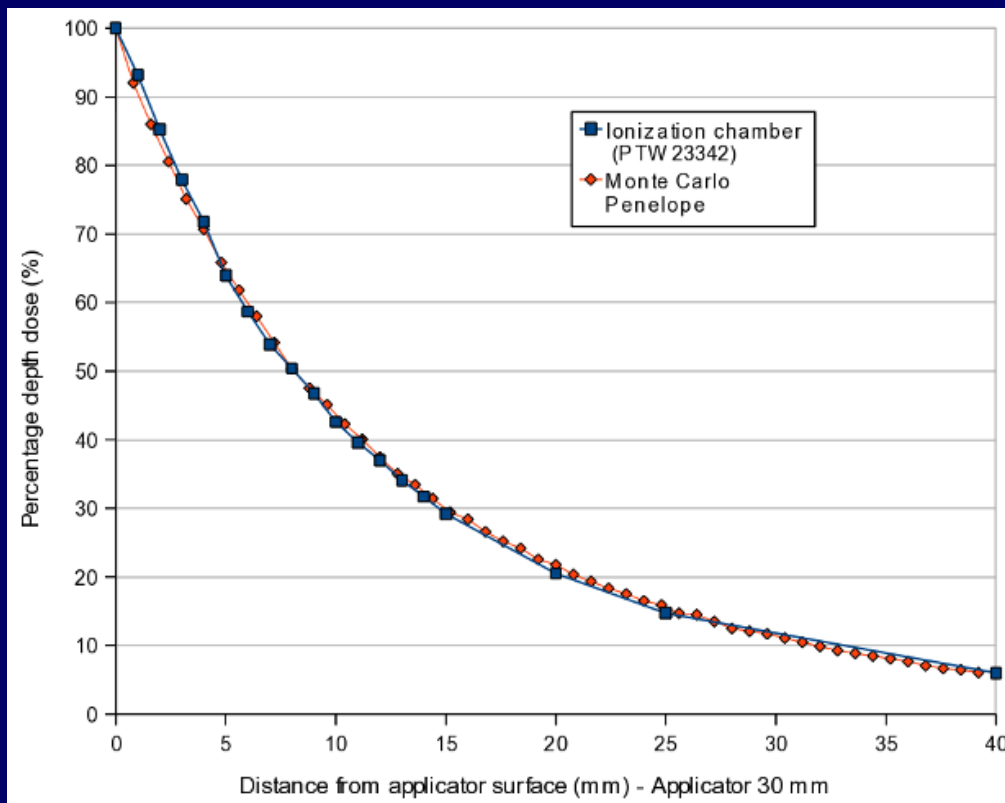
Courtesy Te Vuong



FSD : 40 mm *vs 4 mm*

Dosimetry : Monte Carlo

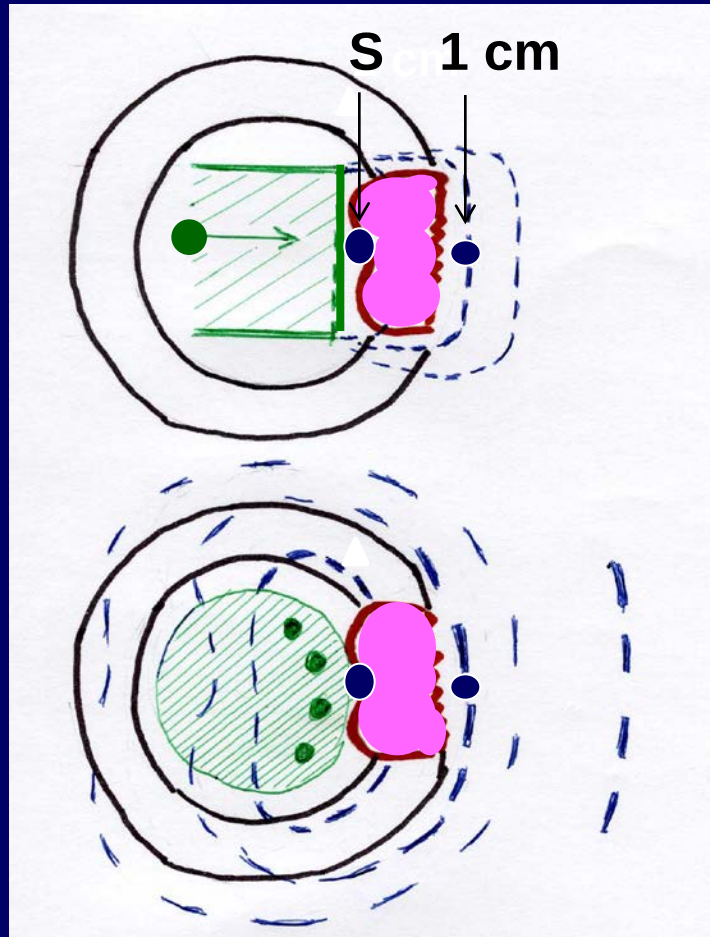
M C 2 PLAN™



Applic 3 cm $\varnothing$ - FSD : 4 cm

Dose 50% at 7 mm

Brachy Endoluminal Rxtherapy Rectum



Cont X Brachy

50 kV

FSD : 40 mm

Iridium 192

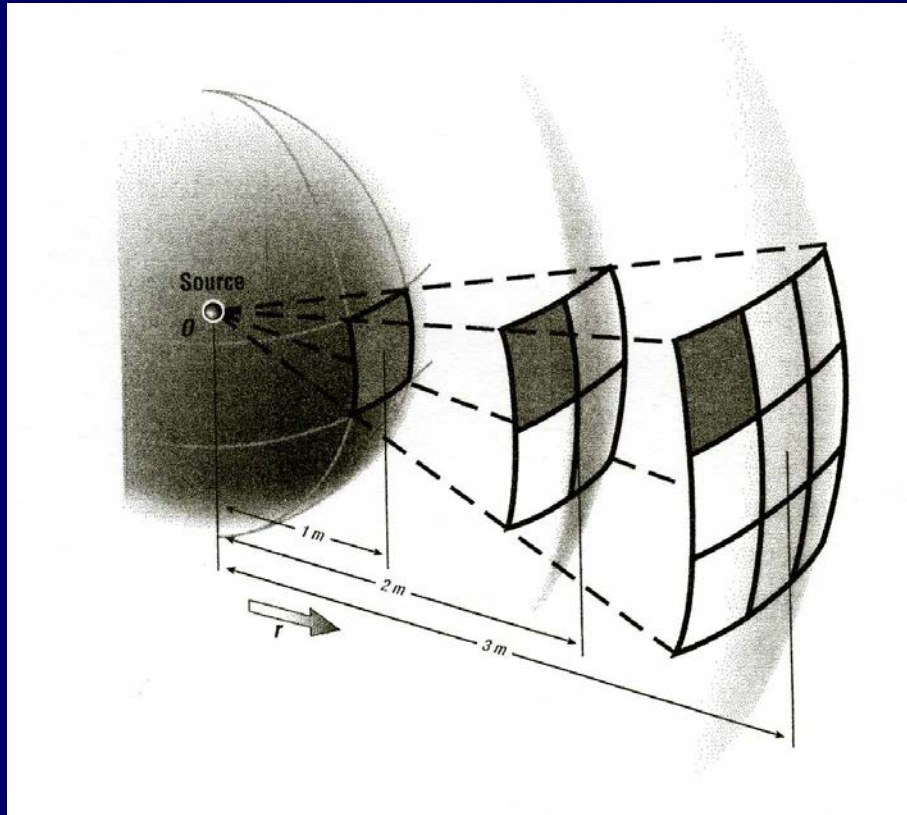
HDR 300 kV

FSD : 4 mm

Dose at **surface applicator (S)** vs Dose at **1 cm**

Inverse Square Law $S : 4 \pi r^2$

ISOTROPIC RADIATION distribution : sphere



1 cm $S : 1$ Dose **10 Gy**

2 cm $S : 4$ D 25% **2.5**

3 cm $S : 9$ D 11% **1.1**

Point Source

Rectum Brachytherapy - Dose Distribution

	Iridium 192 HDR 300 kV 0,4 cm	Contact X Brachy 50 kVp (25 kV) 4 cm
FSD		
Surface	10 Gy	10 Gy
T Depth 1cm	2.2 Gy	4.0 Gy

Protocol AAPM TG-43 U1 : $D = Sk.^{\wedge} \cdot (1/r^2) \cdot gp(r) \cdot \phi an (r)$
Catherine DEJEAN

When **FSD** is short :
0.5 to 6 cm (5 – 60 mm)

- Dose is **inhomogeneous**
- For penetration (% depth-Dose)
FSD (Focus- surface –Distance)
is much more important
than **Energy of beam**

CXB > Iridium

- 1) **Higher** (*inhomogeneous*) **dose in Tumor: Brachy RT** ($1/r^2$)
- 2) **Better accurate Targetting** (*eye vs image guided*)
- 3) **Better protection** of normal rectum
- 4) **Simple** - radioprotection - Cheap
- 5) **Sustainable development** (*no radioactive waste*)

LIMITATIONS

But ...



EXPERT
REVIEWS

Renaissance of contact x-ray therapy for treating rectal cancer

Expert Rev. Med. Devices 8(4), 483–492 (2011)

Jean-Pierre Gérard¹¹,
Arthur Sun Myint²,
Olivier Croce³,
Jacob Lindegaard⁴,
Anie Jensen⁴,
Robert Myerson⁵,
Jean-Michel
Hannoun-Lévi¹
and Serge Marcie¹

Contact x-ray therapy (CXRT) with 50 kV has proven to be an efficient radiation therapy technique to achieve local control and rectal preservation for early rectal adenocarcinoma. Despite these results, CXRT has not been used due to the shortage of the no longer manufactured Philips RT 50™ unit. Recently, a new CXRT machine (Papillon 50™) became available on the market. This machine delivers a beam of 50 kV with a dose rate close to 15 Gy/min and has a percentage depth dose of 50% at 6–7 mm. The applicator size varies from 2–3 cm in diameter. Due to the original design of the main tube, treatment delivery is quick and more comfortable for the patients. An online viewing system incorporated in the tube allows a good visualization of the tumor with improved accuracy of radiation delivery. An international collaborative trial (Contact Endoscopic Microsurgery [CONTEM]) was set up to accrue approximately 300 cases

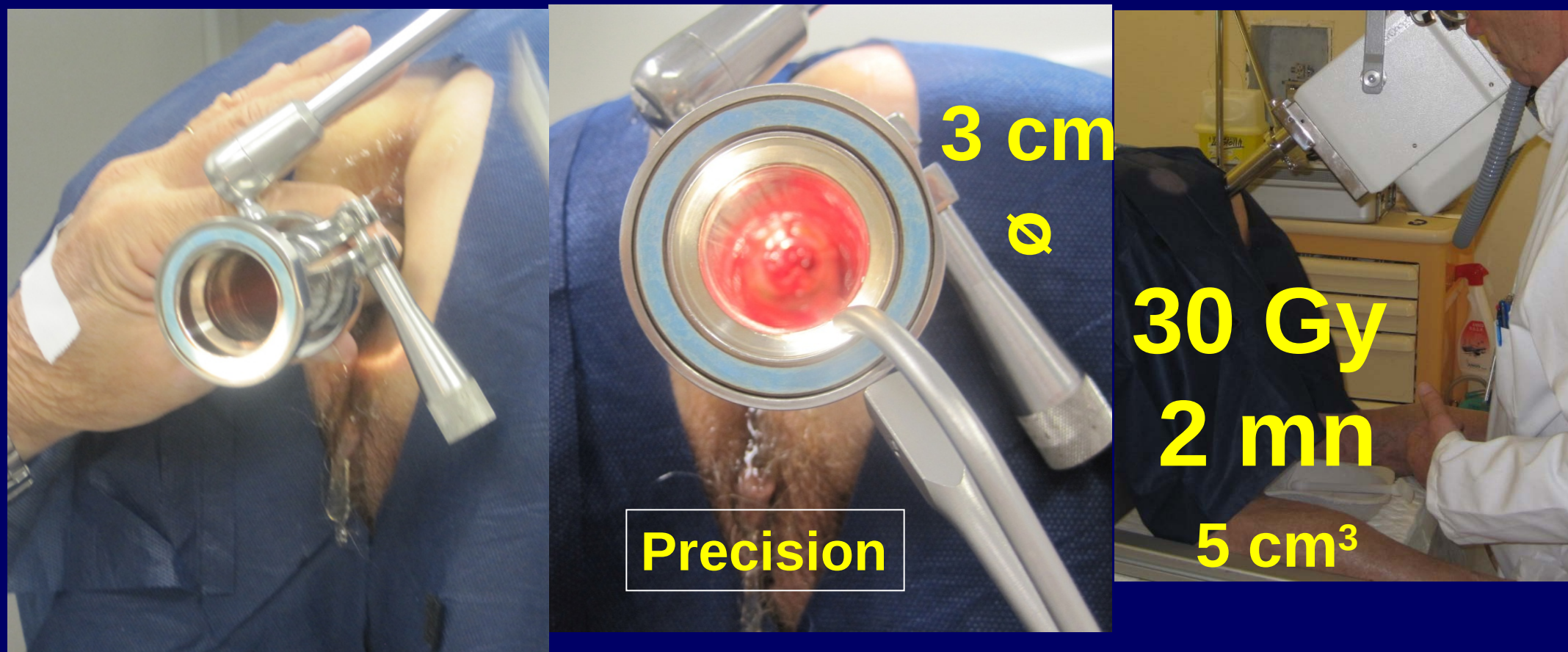
➤ **2001 Philips machine Stop**

2009 : Papillon 50™
UK- French design

Exp Rev Med Dev
2010; 34: 231
JP Gerard & al

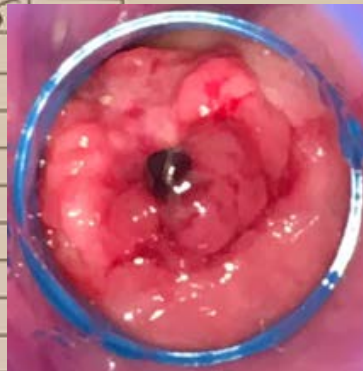


1) Rigid scope 2) Target- Fixation 3) Irradiation



Papillon 50 Transanal Endoscopic Brachytherapy
Non Toxic . Efficient . Simple (any age - ambulatory)
Clinical Expertise

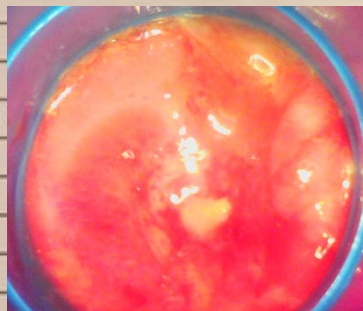
Dose dans les volumes				
	Date	Manips	3cm	2.5
Lu				
Ma				
Me	5/4	52k		
Je	20/7	Attkin	30 Gy	
Ve		Man		
		Salt		
		KB		
Lu				
Ma				
Me				
Je				
Ve				



F 47 y

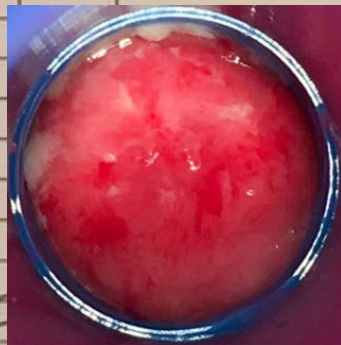
T2N0

Lu				
Ma				
Me	19/4	52k	10	20
Je		52k		
Ve		Man		
		Salt		



Lu				
Ma				
Me				
Je				
Ve				

Lu		KB		
Ma		52k		
Me	3/5	L-E	15	
Je		L-E		
Ve		Man		
		Salt		



Planned Preservation

Peau parfaite.

43 cm entre sans poulain
partite tte "chiquette"

Breps pour Netabolon (Therapie)
+ sang EDTA



Va m'emp. 20k/j m'alle
avec 4 sels/j l'empant
d'entre coxyx

Au TR aje v'illuse au
saut de j'ale. il est a
re. d'alle de 1.5 cm

Rep 90%

Tuberculose
Bacillus



va b'au 99 d'entre p'aleu.
reple r'aleu

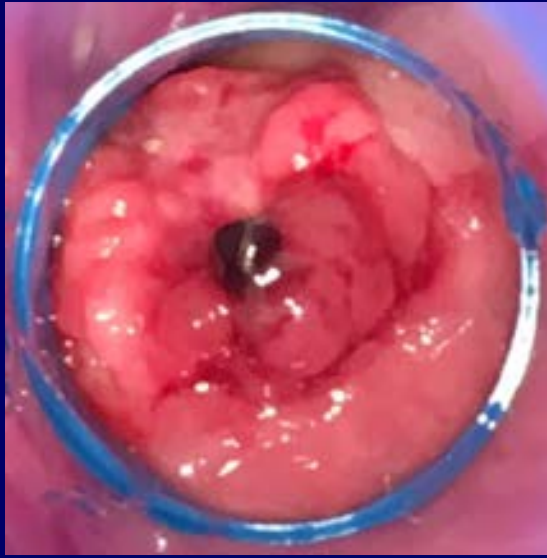
99/ans b'ous
Au TR b'at et s'ole j'alt p'ante
R'ite. p'ape un f'ou s'oupe
p'aleu. p'aleu b'ou

CCR++

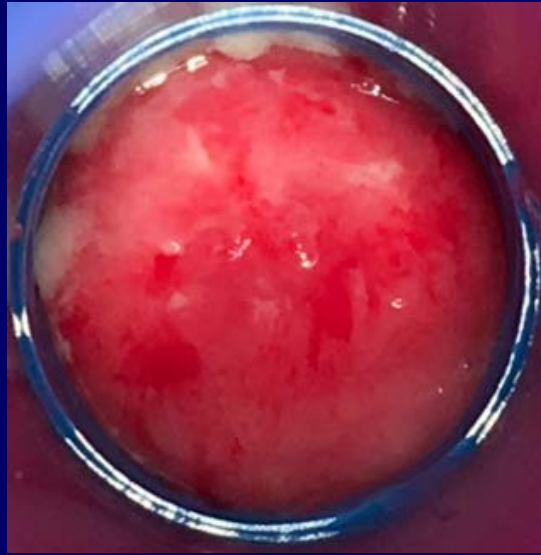
B'ou s'oupe 1.5 Gy j'ale
f'ou en



RCT v'au b'at 1.5 - 1.5 s'oupe



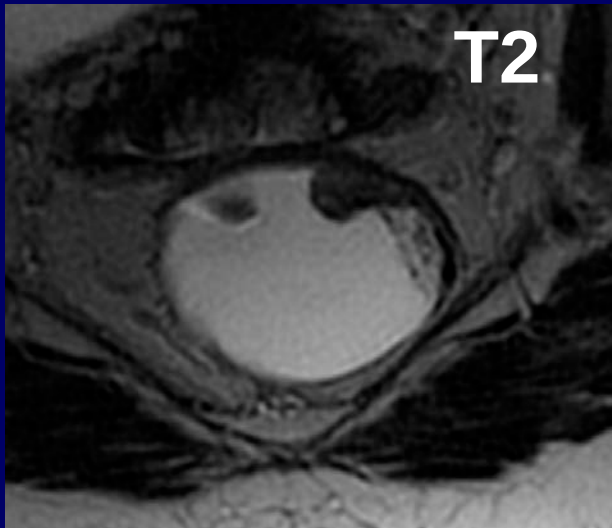
D1 - April 2017



D 28: cCR

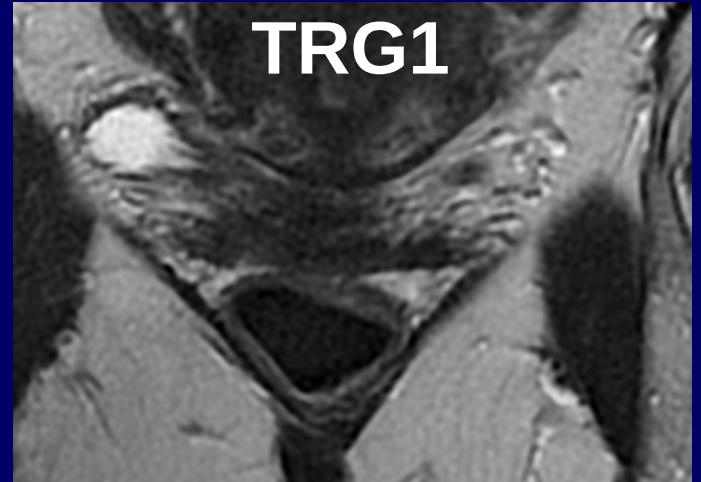


3 months



T2

**+
CAP 50**



TRG1

« Planned » Organ Preservation

CXB France

Nice 2002

JP Gérard
K Benezery
J Doyen

Mâcon 2010

N Barbet

Lyon-
Villeurbanne
2010

R Coquard
N Barbet



Epargne rectale - Papillon 50 - France

2003 – 2016 : 84 pts Contact: 90 Gy/ 3 fr + CAP 50
median age : 74 years Med Fol-up : 53 months

Inst	N° pt	T2	T3	cCR	Loc Rec	Org Pres	Sp S 4 y
Lyon V	16	13	3	16	2	14	83
Macon	11	7	4	10	1	10	86
Nice	57	32	25	53	3	57	81
Total	84	52	32	79 (94%)	6/79 (11%)	81 (94%)	82%

Frin AC & al Eur J Canc 2017;72:124-36

Rectal Preservation Papillon⁵⁰ Europe



2017 Europe
11 P 50 units (Amst.)



Liverpool – Clatterbridge
Papillon Suite: **Pr Sun Myint**

CXB + CRT organ preservation Europe

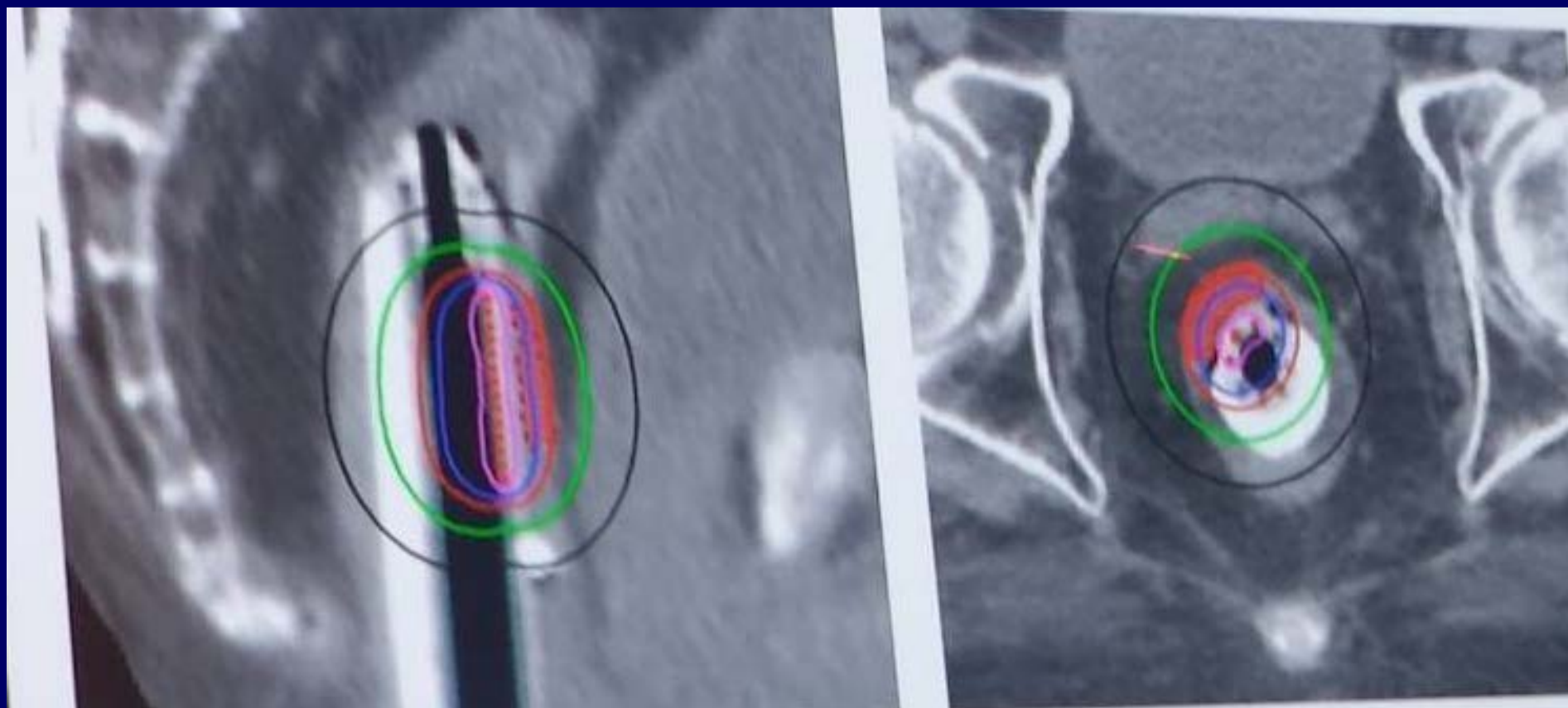
Inst	N° pt	T2	T3	cCR	Loc Rec	Org Pres
Clatterbridge 2002-2014	200	89 ₂₁	90	72%	11%	62%
Hull 2010-2014	43	26	17	90%	15%	89%
Nice – M.V. 2002-2016	84	52	32	94%	11%	94%
Total	327	188	139	85%	12%	81%

Sun Myin t BJR 2017 - Dhadda Clin Onc 2017 - Frin AC & al Eur J Canc 2017

CXB > Iridium

- 1) **Higher** (*inhomogeneous*) dose in Tumor: Brachy RT ($1/r^2$)
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- 3) **Better protection of normal rectum ++**
- 4) Simple - radioprotection - Cheap
- 5) Sustainable development (*no radioactive waste*)

LIMITATIONS

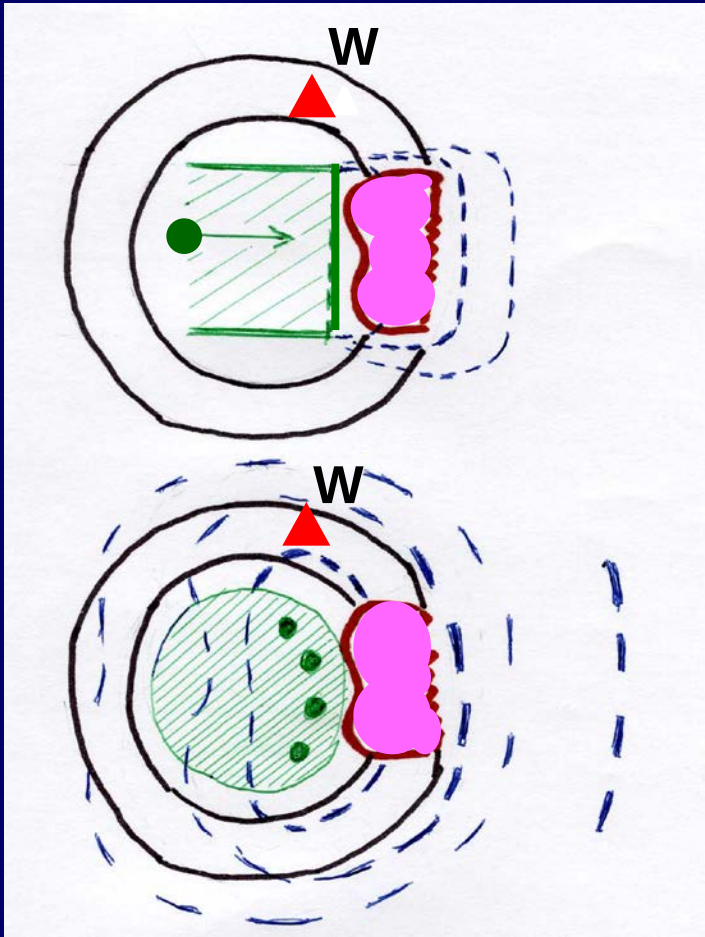


« Leyden » Hebert Study C Marijnen

Max Tolerated Dose : 7 Gy surface (X3) (even with balloon) 30 Gy

Rijkmans E Hebert study Int J Rad O B P 2017; 98: 908-17

Brachy Endoluminal Rxtherapy Rectum



Cont X Brachy

50 kV

FSD : 4 cm

Iridium 192

HDR 300 kV

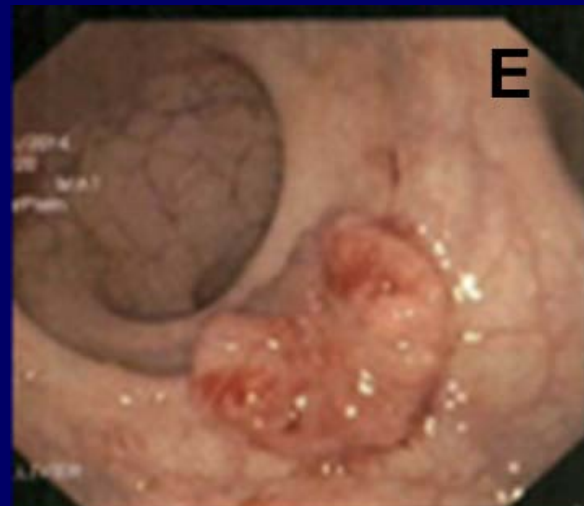
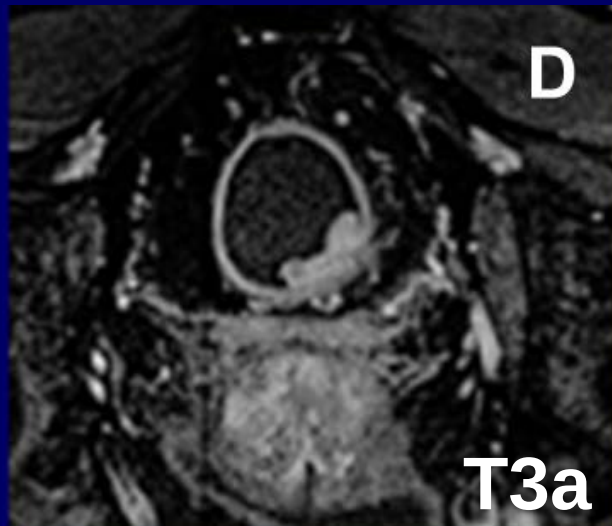
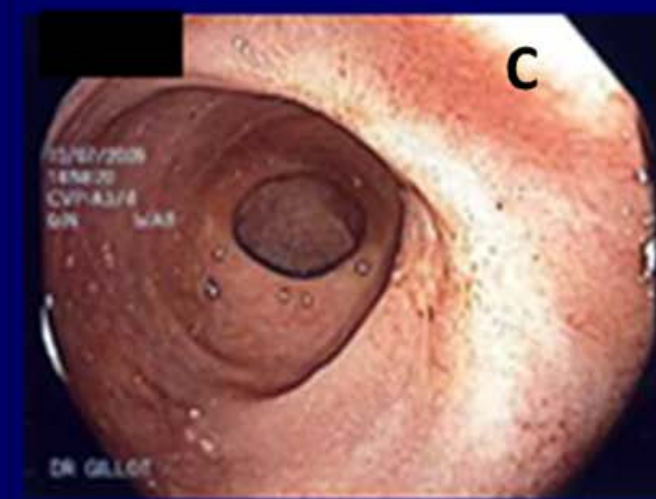
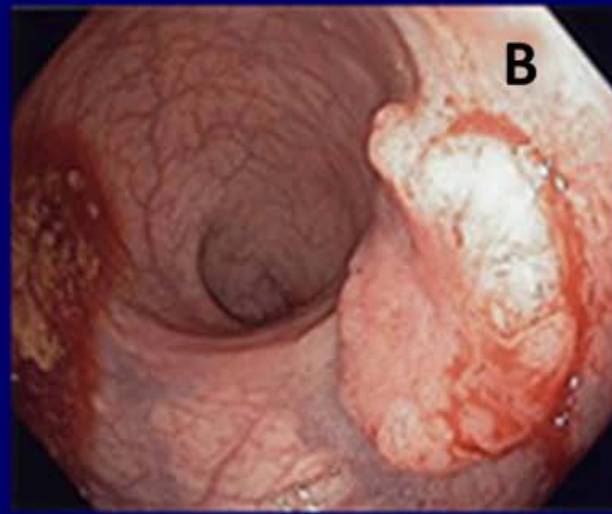
FSD : 0.4 cm

Dose at normal Rectal Wall : W

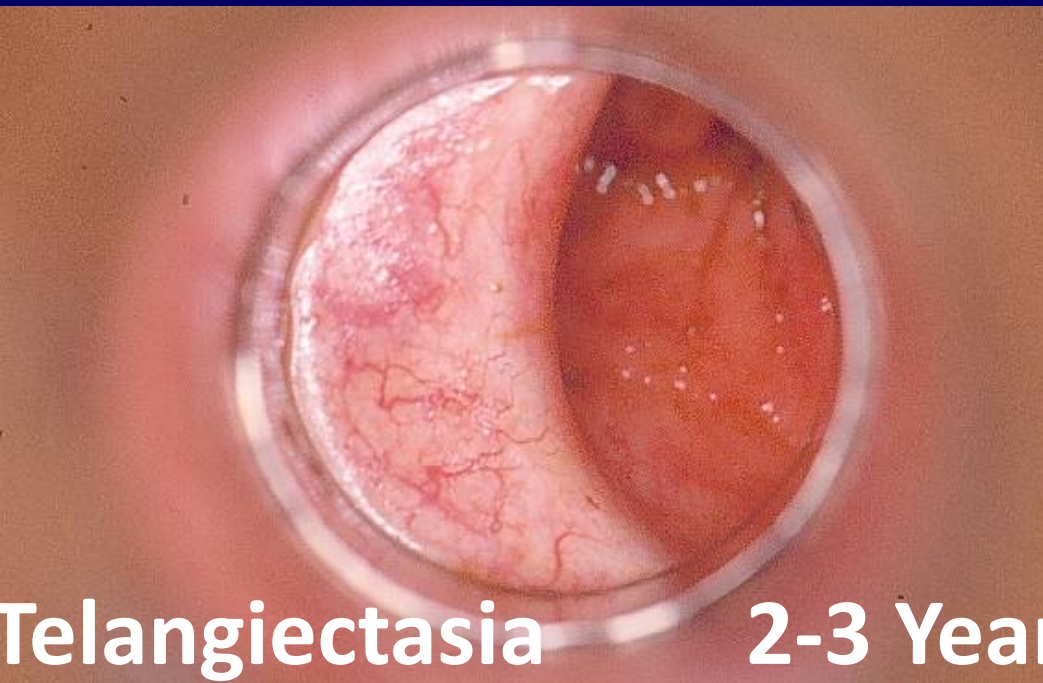
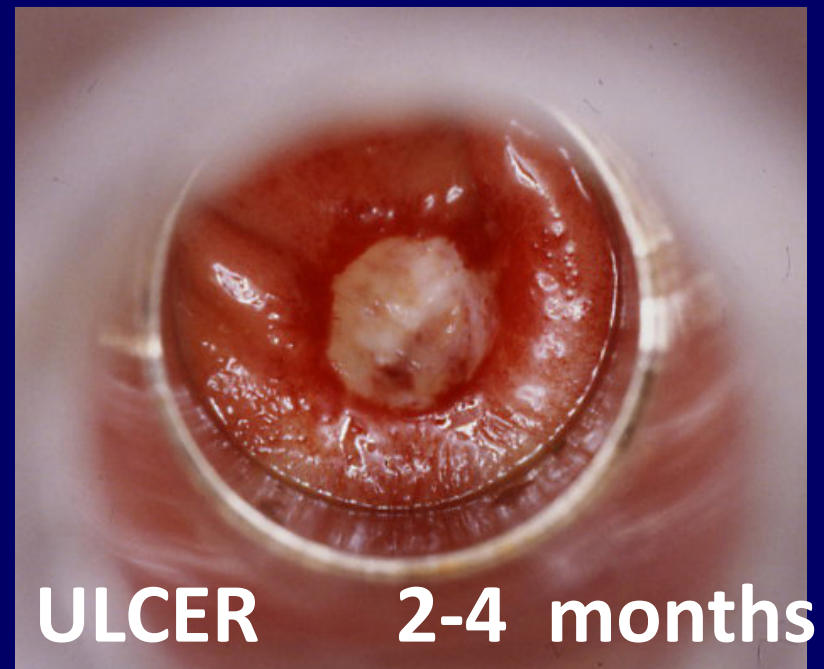
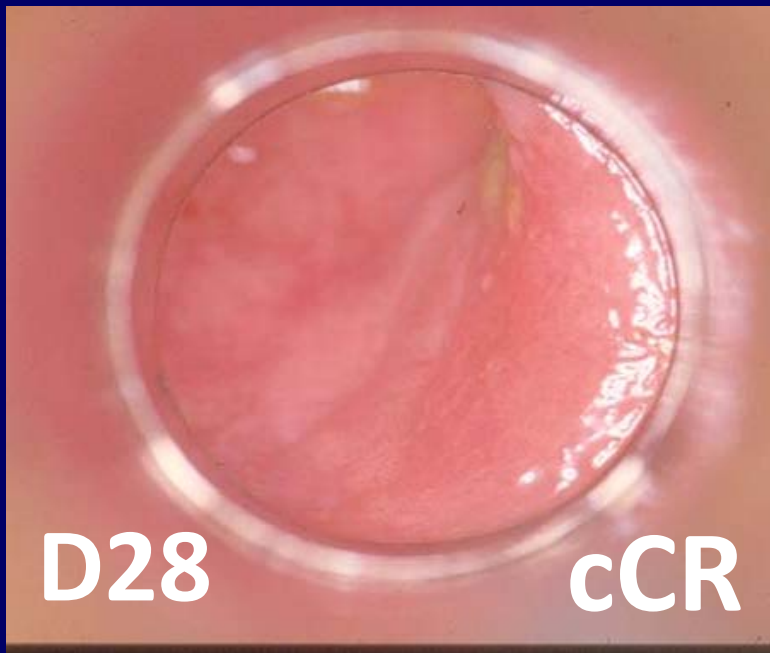
Rectum Brachytherapy - Dose Distribution

FSD	Iridium 192 HDR 300 kV 0,4 cm	Contact X Brachy 50 kVp 4 cm
Surface	10 Gy	10 Gy
1 cm lat.	4 Gy	< 0.1 Gy

protocol AAPM TG-43 U1 : $D = Sk \cdot \frac{1}{r^2} \cdot G(r) \cdot \phi_{an}(r)$



« Petit » T3a-b N0 ≤ 4 cm : CXB+ CRT



LARS SCORE	
During the past month, do you ever have occasion when you cannot control your flatus (wind)?	
No (never)	0
Yes, less than once per week	4
Yes, at least once per week	7
During the past month do you have any accidental leakage of liquid stool?	
No, Never	0
Yes, less than once per week	3
Yes, at least once per week	3
During the past month how often do you open your bowel?	
More than 7 times per day (24 hours)	4
4-7 times per day	2
1-3 times per day	0
Less than once per day	5
During the past month, do you ever have to open your bowel again within one hour of the last bowel opening?	
No, never	0
Yes, less than once per week	9
Yes, at least once a week	11
During the last month, do you ever have such a strong urge to open your bowel that you have to rush to the toilet?	
No, never	0
Yes, less than once a week	11
Yes, at least once a week	16

TOXICITY - BOWEL FUNCTION (Nice - 57 pts)

- **Rectal Bleeding** (1-3 years after RT)

G1-2 : 25% G3 : 9% (plasma-argon)

- **Bowel Function** (score)

- **MSKCC** : Good-excellent : 82.5% - Fair : 12.5%

Poor : 5%

- **LARS (26 pts)** : Good (0-20) : 15 Fair (21-29): 9 Poor > 30: 0

CXB > Iridium

- 1) **Higher** (*inhomogeneous*) dose in Tumor: Brachy RT ($1/r^2$)
- 2) Better accurate **Targetting** (*eye vs image guided*)
- 3) Better protection of **normal rectum : less toxicity**
- 4) **Simple - radioprotection** (*room shield*) - **Cheap**
- 5) **Sustainable development** (*no radioactive waste - Terrorism*)

Intraluminal Brachytherapy For Rectal Cancer

CXB > HDR

CXB > Iridium

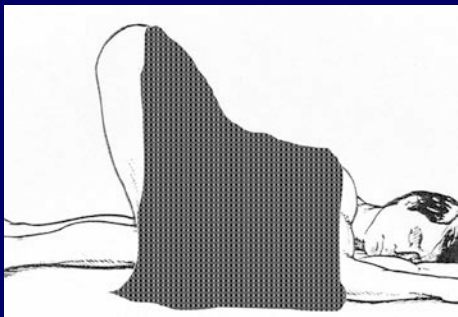
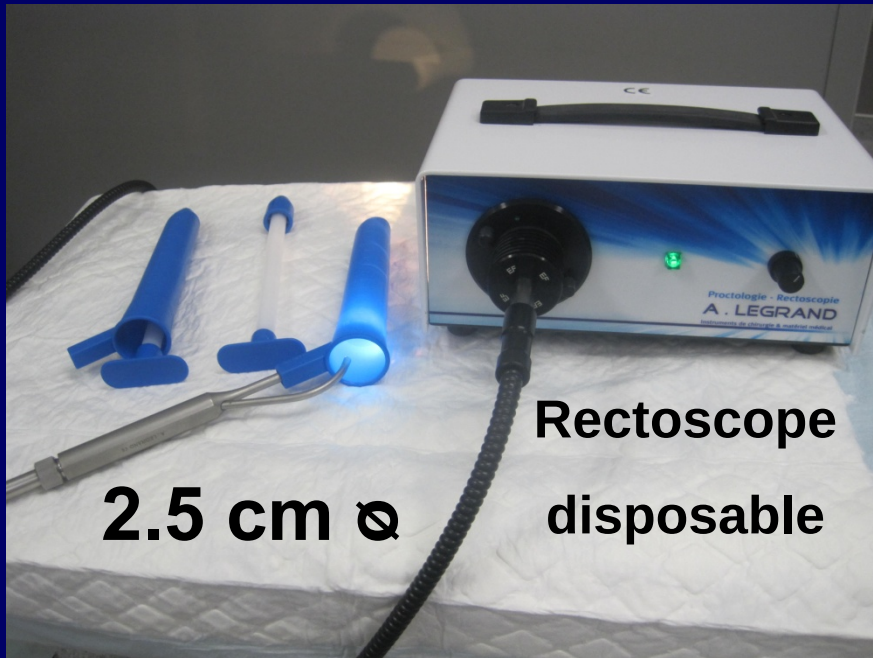
- 1) Higher dose in Tumor: Brachy RT ($1/r^2$)
- 2) Better accurate Targetting (*eye vs image guided*)
- 3) Better protection of normal rectum and OAR ++
- 4) Simple - radioprotection - Cheap
- 5) Sustainable development

***LIMITATIONS* +++**

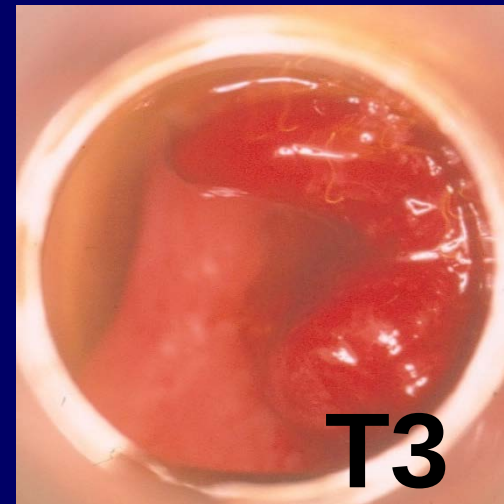
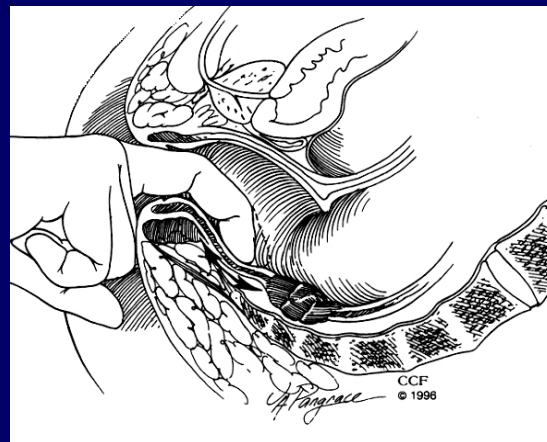
Limitations

- **Size Tumor** : $< 5 \text{ cm } \phi$ (*4 cm ?*)
- **Site** : Anal canal , Supra anal post , $> 10 \text{ cm AV}$
- **Operator dependent** : Rigid rectoscopy : **Training**
- **Still few strong evidence** : **OPERA**
- **Few machines but ...** **Papillon +**

Rigid RECTOSCOPY clinical practice



Knee-chest



CRT first line – Organ Preservation

Habr Gama (W Heald) : Watch & Wait **if cCR**

Sao Paulo : T2 vs T3 - Canada - US :

MSKCC : T2 CRT + loc. Exc ; Prospect

Europe : UK – NL- DK – France (GRECCAR) – Italy

IWWD : 1 009 pts

cCR : 15 - 50% Loc Rec : 25 % [10-35]

Organ preservation : 20-40%

Habr Gama ; Garcia Aguilar ; Jakobsen; Beets ; Rullier ; Renehan;

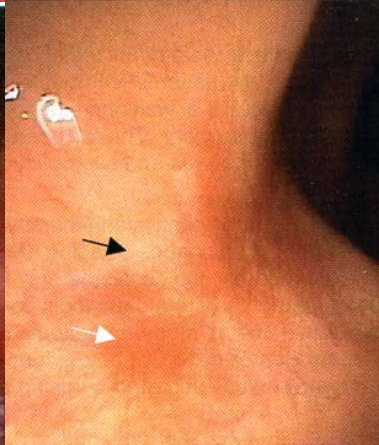
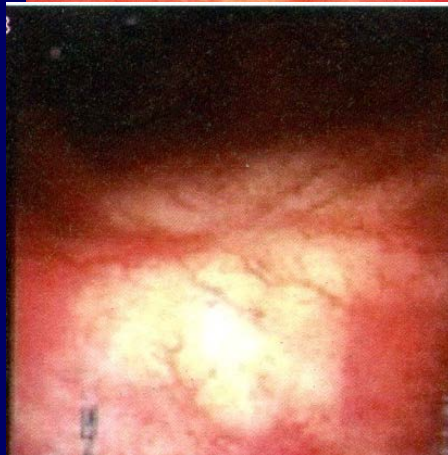
T2-3ab

CRT Alone

Habr Gama



Beets (Greccar)

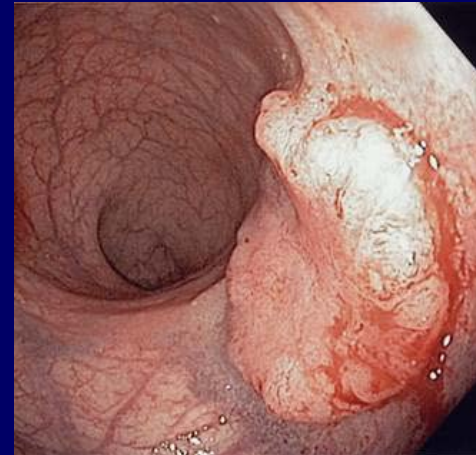


CRT alone

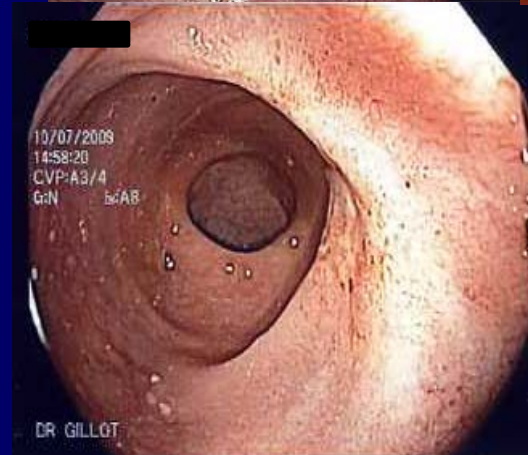
cCR 45 %
Loc Rec 30 %

CRT + boost CXB

Gérard



Sun Myint



CRT + Boost CXB

90 %
≤15 %

ORGAN PRESERVATION

7/6/2017

NAME	N°	PHASE	INCLUSION CRITERIA	DESIGN	END POINTS	N PATIENTS	STATUS	COMMENTS
STAR-TREC UNIVERSITY OF BIRGIMGHAM	NCT02945566	II	mriT1-3bN0 (<5 mm mesorectal invasion) or uT1-T3b Tumour < 4 cm	A -TME B - 50 Gy + Cap C - 5x5Gy	1 st - Organ saving rate at 12 months 2ndary – TME Rate, 3 year pelvic failure, mrTRP assessment by MRI rate, OS, SFS,	120 (411 600)	ONGOING	
OPERA CAL Pr Gérard	NCT02505750	III	A – EBRT + Boost 9Gy	A – Cap 45 Gy + Boost 9Gy B – Cap 45 Gy + CXB 90 Gy	1 st -Organ preservation rate without salvage surgery and absence of stoma at 3 year 2ndary – cCR, OS, DSF, TRG score	236	ONGOING	
GRECCAR 12 CHU Bordeaux E.Rullier	NCT02514278	III	cT2-T3, cN0-1 T < 4 cm, < 10 cm to AV	A – FOLFIRINOX (x4) + CAP 50 B- Cap 50	1 st -Organ preservation rate and absence of stoma at 1 year 2ndary – Compliance, tolerance, rpCR, histologic CR rate, curative surgery rate,	218	ONGOING	
MORPHEUS Jewish General Hospital Sir Mortimer B. Davis	NCT03051464	Pilot Study	mT2-T3, N0-N1 (<5 mm mesorectal invasion) T < 5 cm, < 10 cm to AV	A – Cap 45 + boost 9Gy B- Cap 45 + HDBRT 30 Gy x3	1 st -rectum preservation at 2 year 2ndary – Local recurrence, SFS, OS, QoL	40	ONGOING	

EVIDENCE OPERA : Phase III (Start 6-2015)

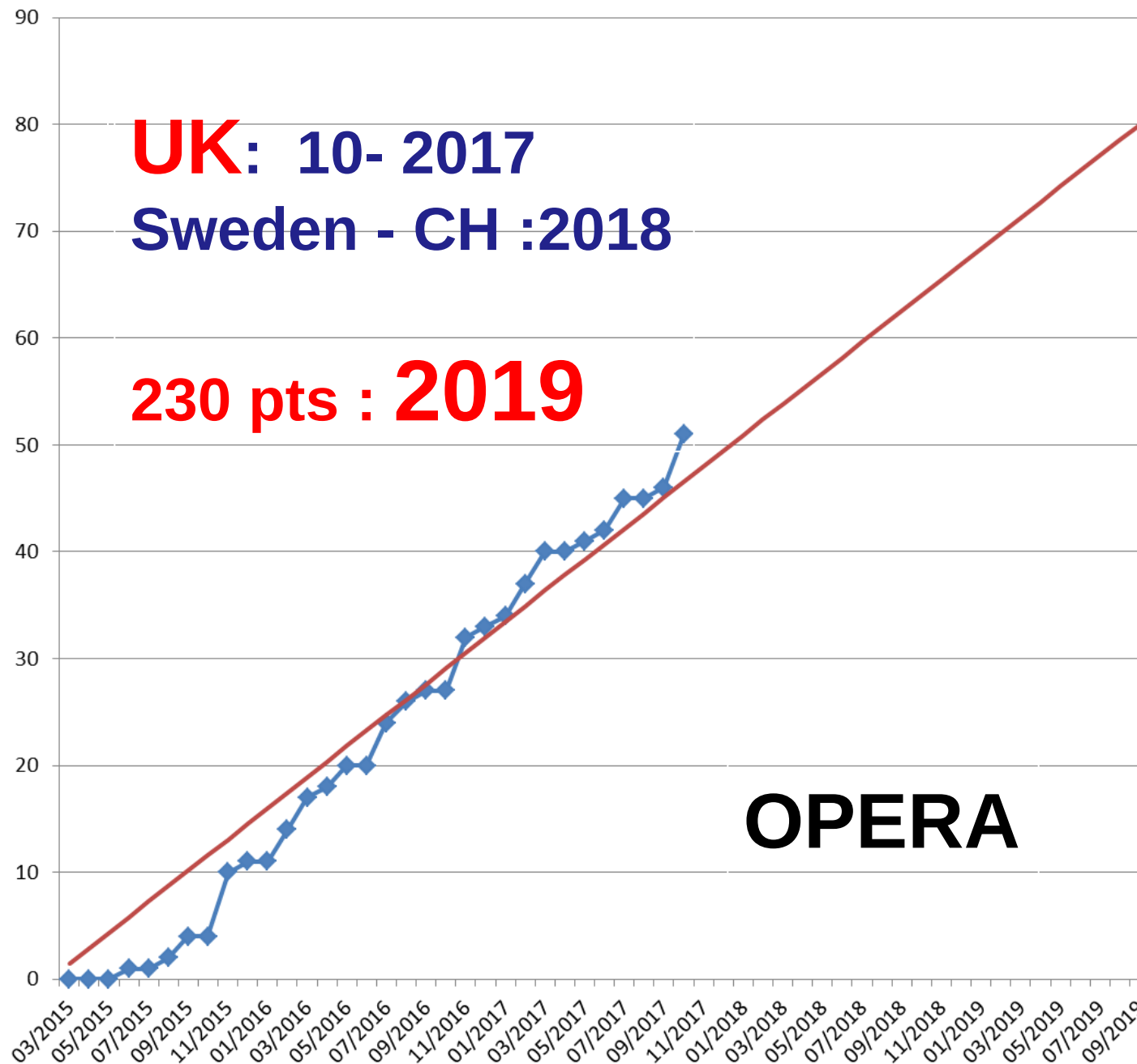
Organ Preservation Early Rectal Adenocarcinoma

T 2-3a-b < 5 cm $\emptyset$ N0 M0 ; Distal- Middle Rectum



End point : organ (rectal) preservation 3 years (stoma- Loc R)

Hypothesis : 20 % to 40 % 236 pts (HR:0.56)



France
90 pts

start
June 2015

November
2017
55 patients

50 pts randomized

- Med age : 68 y
- T2 : 32 / T3 : 18 ≤ 3 cm $\varnothing$: 21 / > 3 cm : 29
- Distal : 36 / Middle : 14
- **CRF : 27 pts** **cCR** ₍₁₂₎ + near cCR ₍₉₎ : **21/27 (78%)**
-

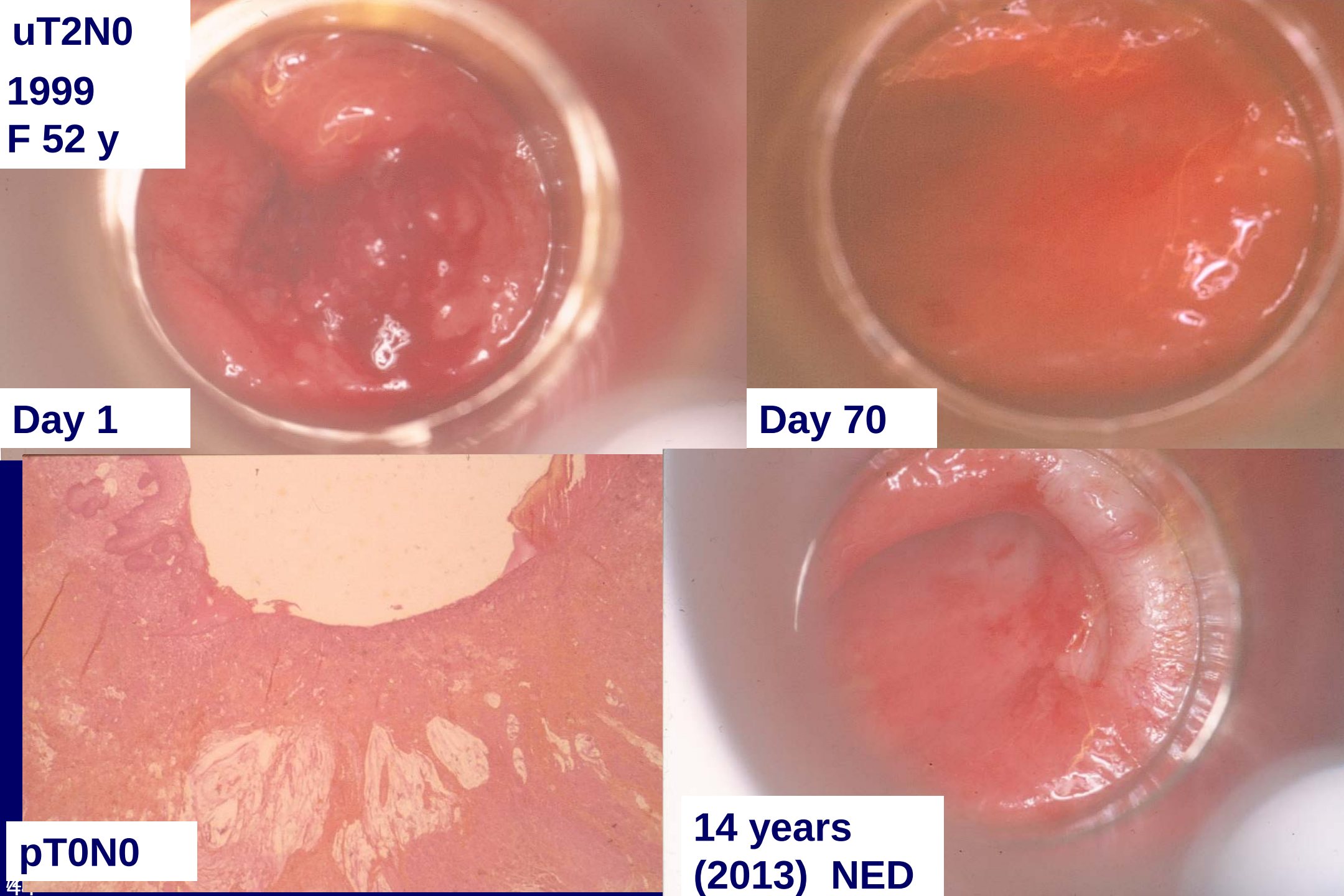
No SAE

Gr 3 Toxicity : 3

Death : 0

Randomisez trial : Lyon R 96-02

- **T2-3** Distal rectum Opérable : 90 pts 1996-2001
Preop alone EBRT vs EBRT + **Boost CXB** : TME to all pts
- Main End Point : sphincter preservation
- **Clinical CR: 2% vs 30% Sph pr : 44% vs 76% (p< 0.05)**
- At 10 years , 10 patients cCR : **rectum Conservation**
- TME after boost contact : Feasible – **No extra toxicity**
- **No isolated perirectal recurrence**



uT2N0
1999
F 52 y

Day 1

Day 70

pT0N0

14 years
(2013) NED



« Magic stick **2017** »

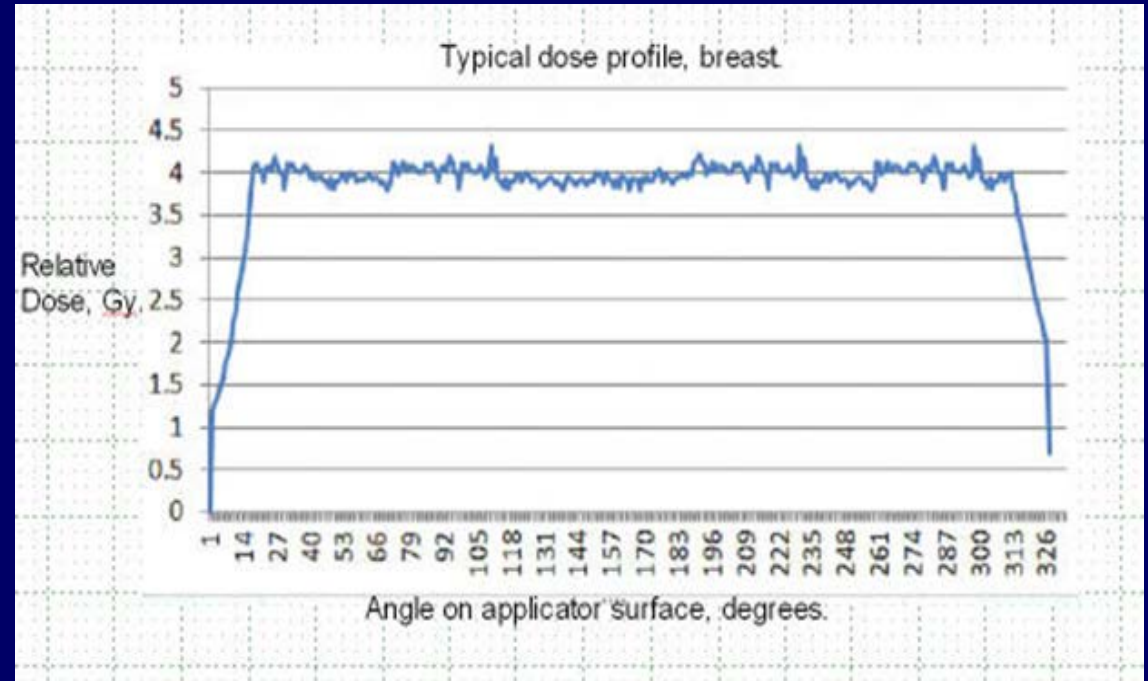
Papillon + X Ray Tube : **Micronode 310 °**

N B A



NICE BREAST APPLICATOR™

Gaf Chromic Films 3/2013 Derby



NBA Dose profile Flatness

JP GERARD 2013 Rad Oncol ; 106 supp 2 OC- 0479

DEPOT D'UNE DEMANDE DE BREVET

PAYS : FRANCE

DONNEES DE DEPOT :

Numéro : FR.11/53529

Date de dépôt : 22.04.2011

DEPOSANT(S) : Centre Antoine Lacassagne
Gérard Jean-Pierre

INVENTEUR(S) :

TITRE DE L'INVENTION :

Dispositif muni d'un jeu d'applicateurs pour la radiothérapie de contact et système comprenant ledit dispositif

REFERENCE : NICE BREAST APPLICATOR - DOSSIER : B2262

N/Réf : B2262/PCT - RRI (JPD/VS)

OBJET : DEMANDE DE BREVET INTERNATIONALE PCT/EP2012/057317 DEPOSEE LE 20.04.2012

Réf : NICE BREAST APPLICATOR – Dossier B2262

RAPPORT DE RECHERCHE INTERNATIONALE et OPINION ECRITE

Cher Monsieur,

Nous venons de recevoir le rapport de recherche internationale sur la demande PCT ci-dessus référencée ainsi que la première opinion de l'examineur international, dont nous vous remettons ci-joint une copie.

RAPPORT DE RECHERCHE INTERNATIONALE

Comme vous le remarquerez, ce rapport reprend les documents déjà cités dans le rapport de recherche de la demande de brevet français. Vous possédez déjà un exemplaire de chacun d'eux. Par conséquent, nous en conservons la copie dans nos dossiers.

OPINION ECRITE

Comme vous le remarquerez, l'opinion indique que les revendications répondent aux critères suivants :

REVENDEICATIONS CONCERNEES	NOUVEAUTE	ACTIVITE INVENTIVE	POSSIBILITE D'APPLICATION INDUSTRIELLE
OUI	-	7, 9, 15-17	1 – 17
NON	1 - 17	-	-

OMBI – OFFICE MEDITERRANEEN DE BREVETS D'INVENTION ET DE MARQUES

BREVETS, MARQUES, MODÈLES, DROIT D'AUTEUR, CONTRATS, LITIGES, CONTREFAÇONS, NOMS DE DOMAINE, CERTIFICATS D'OBTENTION VEGETALE

Bureau à Nice
20, RUE DE LA LIBERTÉ
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FAX. : +33 (0) 4 93 88 16 17

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S.A.R.L. AU CAPITAL DE 701 250 €
TVA N° : FR 27 519023725
SIRET : 519 023 725 00012 – APE : 7010 Z

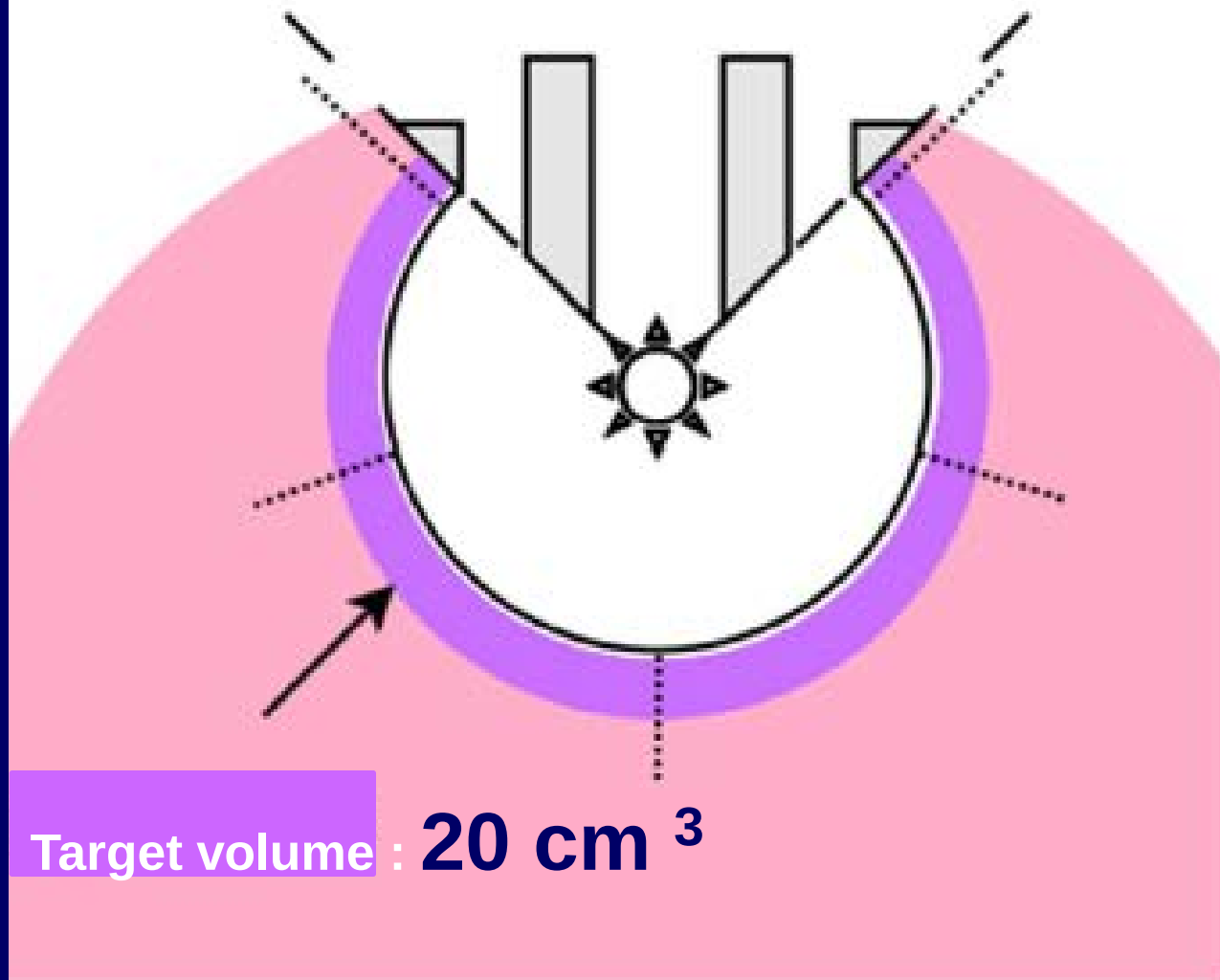
Bureau à Sophia Antipolis
LES ALGORITHMES - BAT ARISTOTE A
2000, ROUTE DES LUCIOLES
06901 SOPHIA ANTIPOLIS CEDEX – FRANCE
TEL. : +33 (0) 4 92 94 47 87

French patent
04.2011

European **PCT** patent
03.2013

NBA Patent

270 Degree Aplicator



Isotropic
beam

2017- 18



Appareil Papillon+™ : 12 - 2017
Breast IORT 2 mn → Cancer Rectum

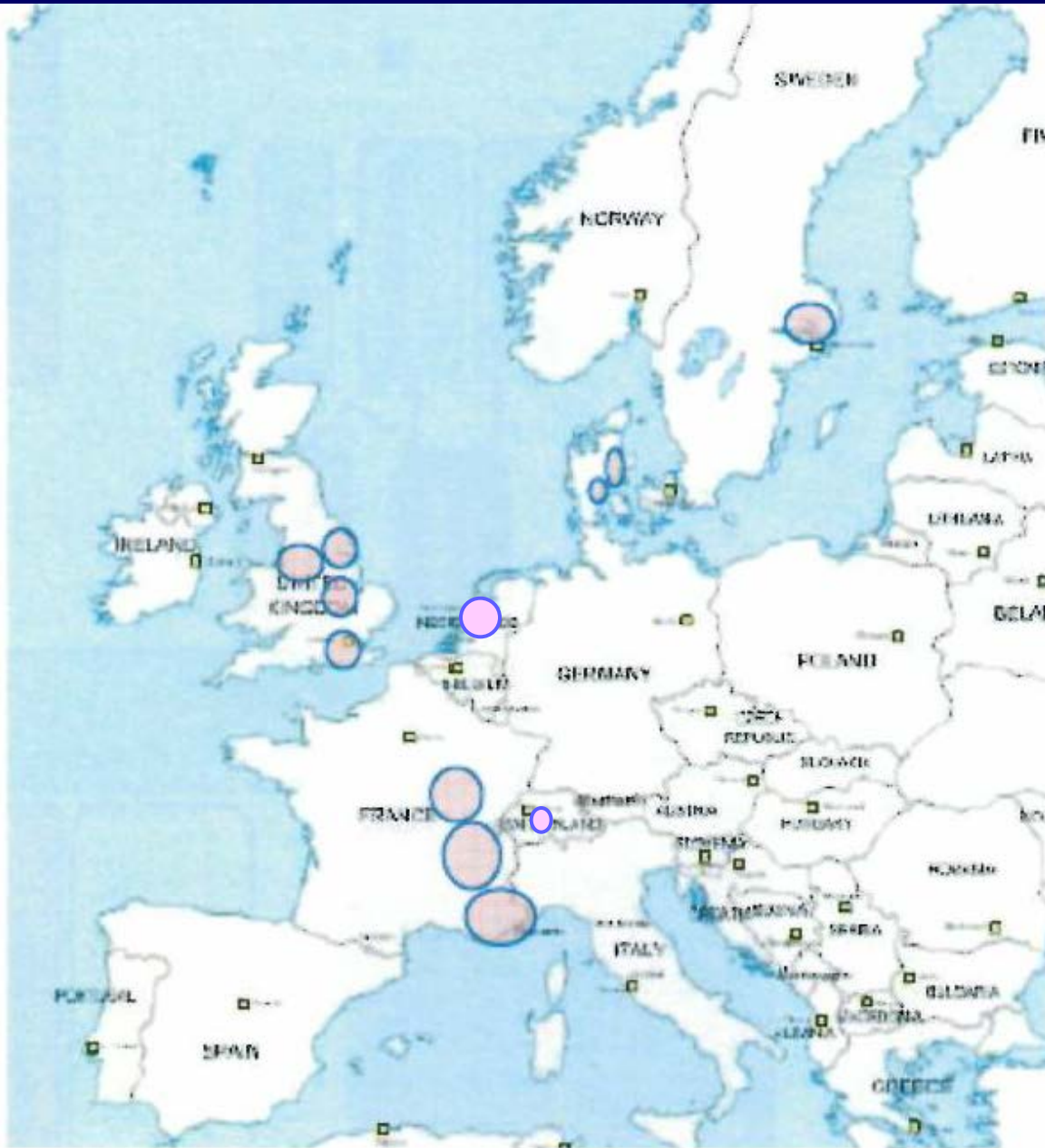
2017 – 2018

New Papillon +

**Amsterdam NKI
RMH**

...

**Amendment
OPERA**



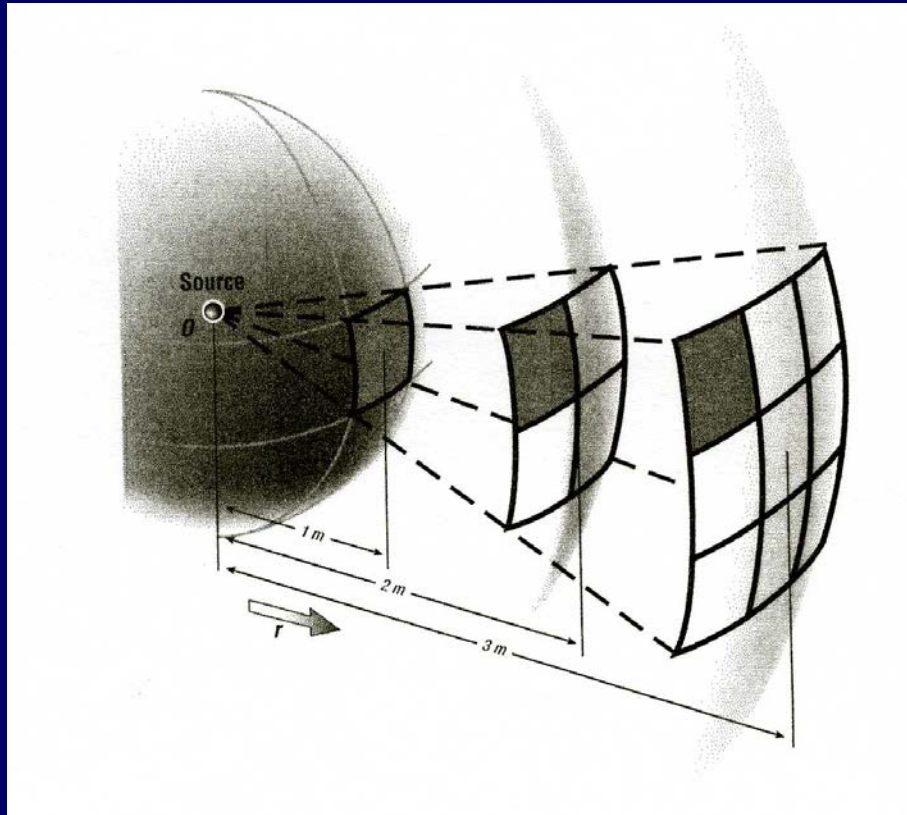
Papillon future developments

- Papillon + tm , rode (tube) $\varnothing$: 22 mm - 16 mm
- Beam distribution: isotropic (lateral beam)
- Intraluminal « stepping source »
- Larger T3 Tumor :
 - > 5 cm long > 1/2 circumference
- Shielding (efficient) normal rectum : 50 kV: lead

2018 - 2019 ??

Inverse Square Law $S : 4 \pi r^2$

ISOTROPIC RADIATION distribution : sphere



1 cm $S : 1$ Dose **10 Gy**

2 cm $S : 4$ D 25% **2.5**

3 cm $S : 9$ D 11% **1.1**

Point Source

When **FSD is short** :
0.5 to 6 cm (5 – 60 mm)

- Dose is inhomogeneous
- For penetration (% depth-Dose)
Increased FSD (Focus- surface –Distance)
is the most important

Veijle vs Novisad / Oncosmart Applicator : Iridium HDR



Veijle

HDR 300kV

FSD : 10 mm

Novisad

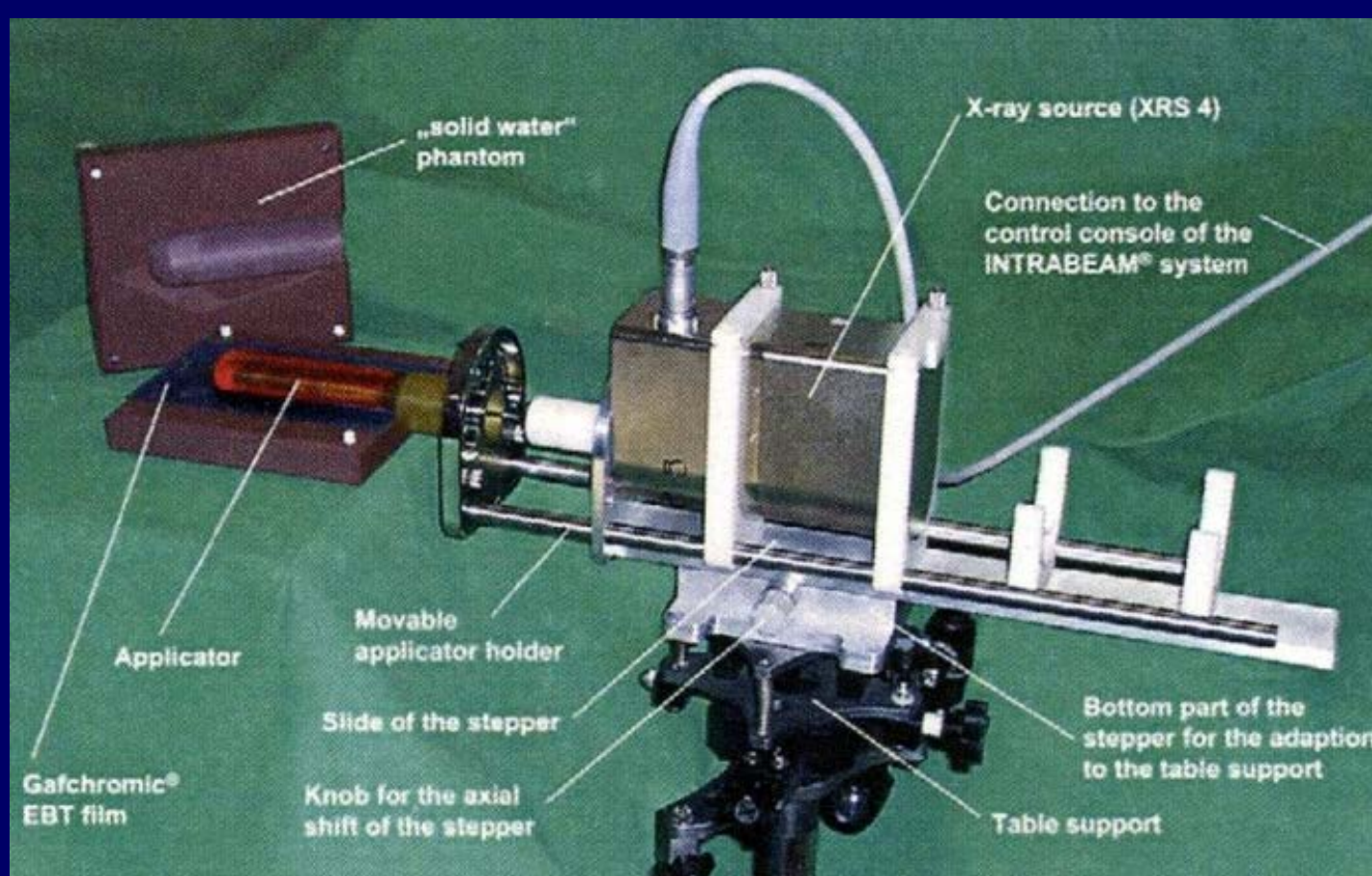
HDR 300 kV

FSD : 4 mm

Novisad / Oncosmart (4mm) vs Veijle (10 mm)

FSD 4mm	r	r ²	dose	↗ surface	dose ↘	dose
	4	16	100			100
5mm	9	81		81/16 : 5	5	20
10mm	14	196		196/16 : 12	12	8
FSD 10mm	r	r ²	dose	↗ surface	dose ↘	dose
	10	100	100			100
5 (15)	15	225		225/100 : 2.25	2.25	40
10 (20)	20	400		400/100 : 4	4	25

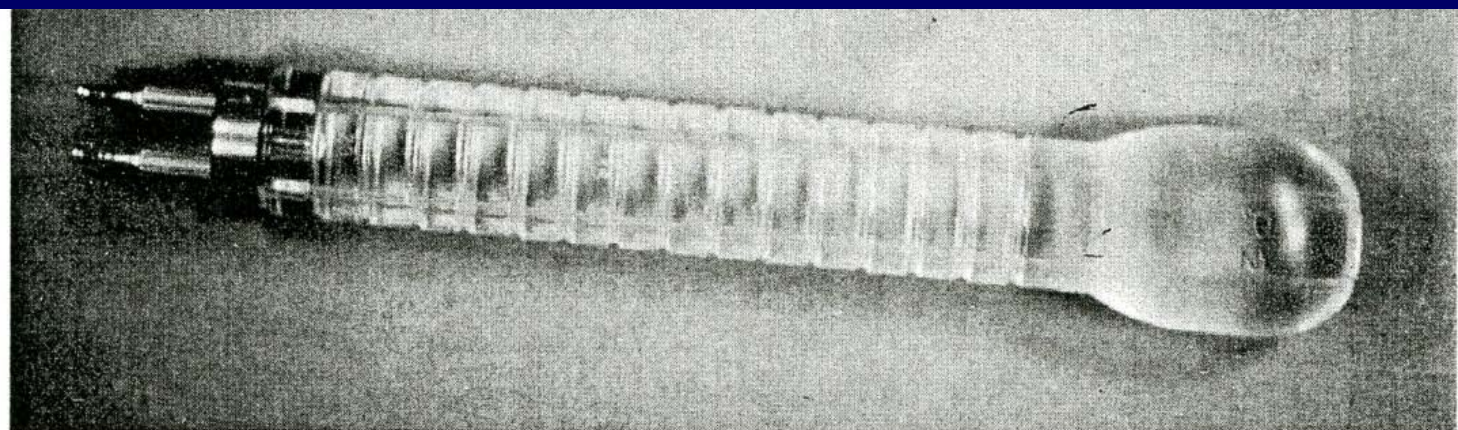
Brachytherapy : FSD Key parameter for Depth Dose



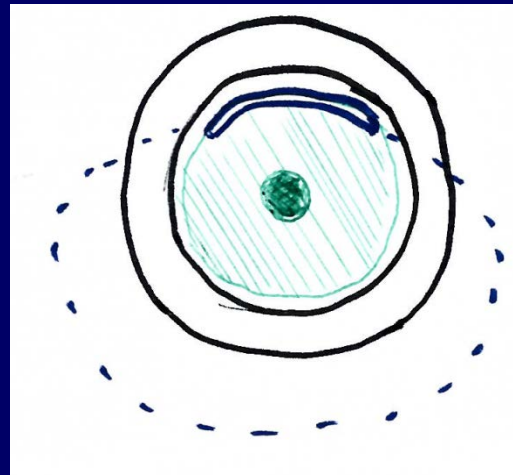
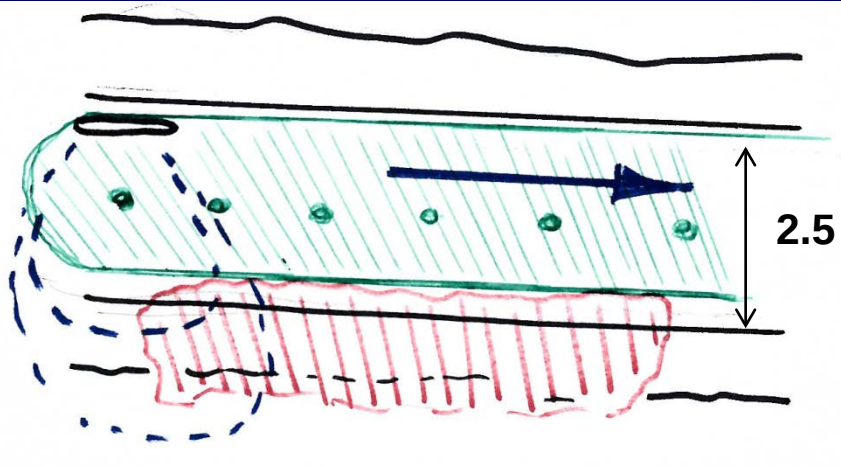
HDR
vaginal vault
Stepping source
X 50 kV

Schneider IJROBP 2009

Papillon +



Rectal Applicator
2.5 cm $\varnothing$
FSD : ≥ 1 cm
T3 b-c > 4 cm

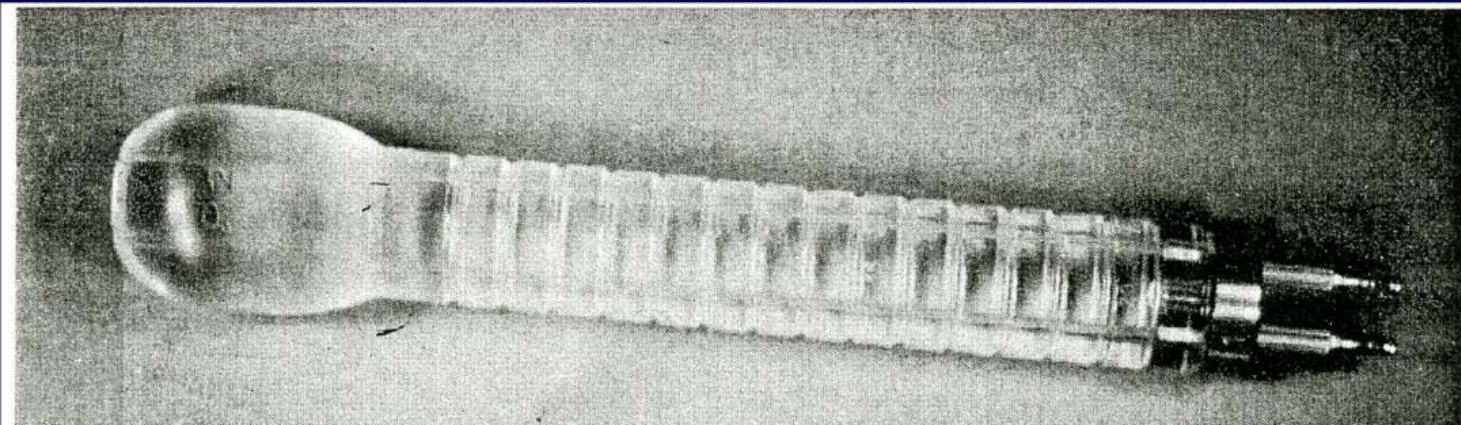


Papillon + 50 kV Rectal Applicator

2.5 cm ϕ

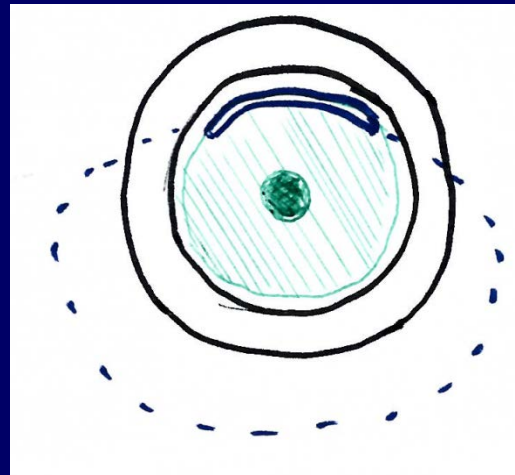
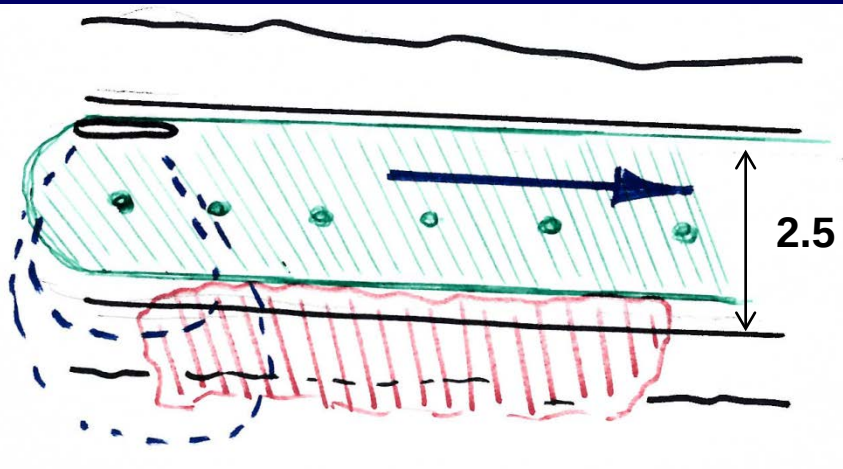
FSD : ≥ 1 cm

T3 b-c > 4 cm



*Lateral beam
Lead shield
Stepping source*

Montreal – Veijle - Leyden



Papillon + 50 kV Rectal Applicator

2.5 cm ϕ

FSD : ≥ 1 cm

T3 b-c > 4 cm



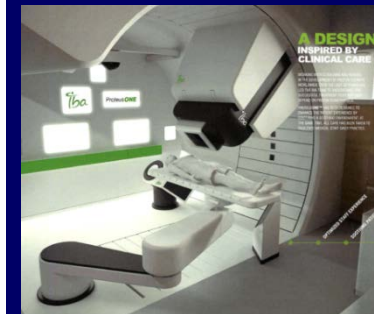
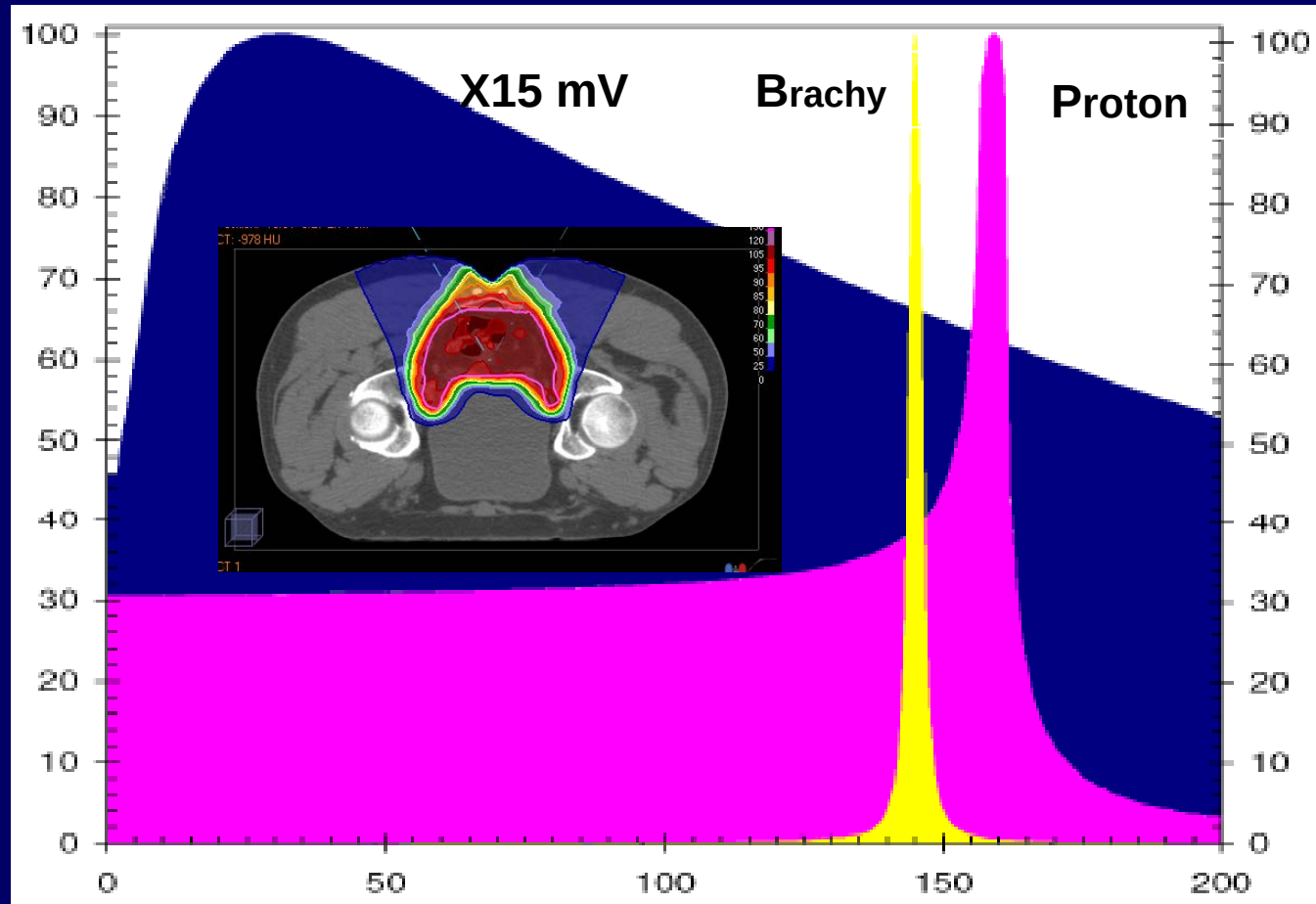
*Lateral beam
Lead shield
Stepping source*

Take home message

- OPERA - Morpheus
- Radioprotection 50 kV > Iridium
- Efficacy : T targeting - High Dose
OAR protection
Therapeutic ratio
- Cooperation (cost/benefit ratio)

Rectal Cancer Radiotherapy 2024 !

Proton
+
Brachy

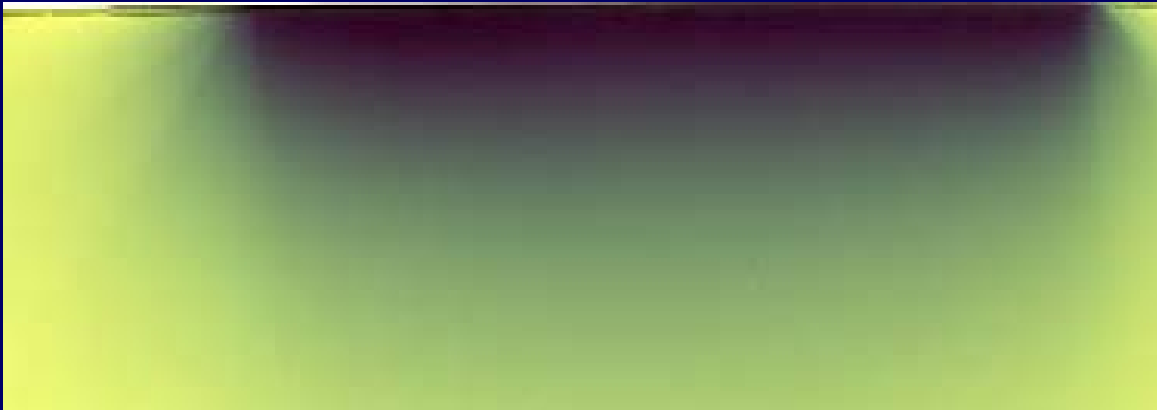


Paris - BATACLAN - Bruxelles -
NICE -Londres- Barcelone – New York
Not afraid !

MERCI

Step and shoot linear rectal treatment with Papillon X ray brachytherapy system.

Example



Step and shoot: 25 mm diameter applicator, 60mm beam length, 7 steps

HVL: 0.65 mm Al,

SSD: 11mm

Dose at 5mm tissue depth: 36%

Total radiation beam on time: 4 minutes.

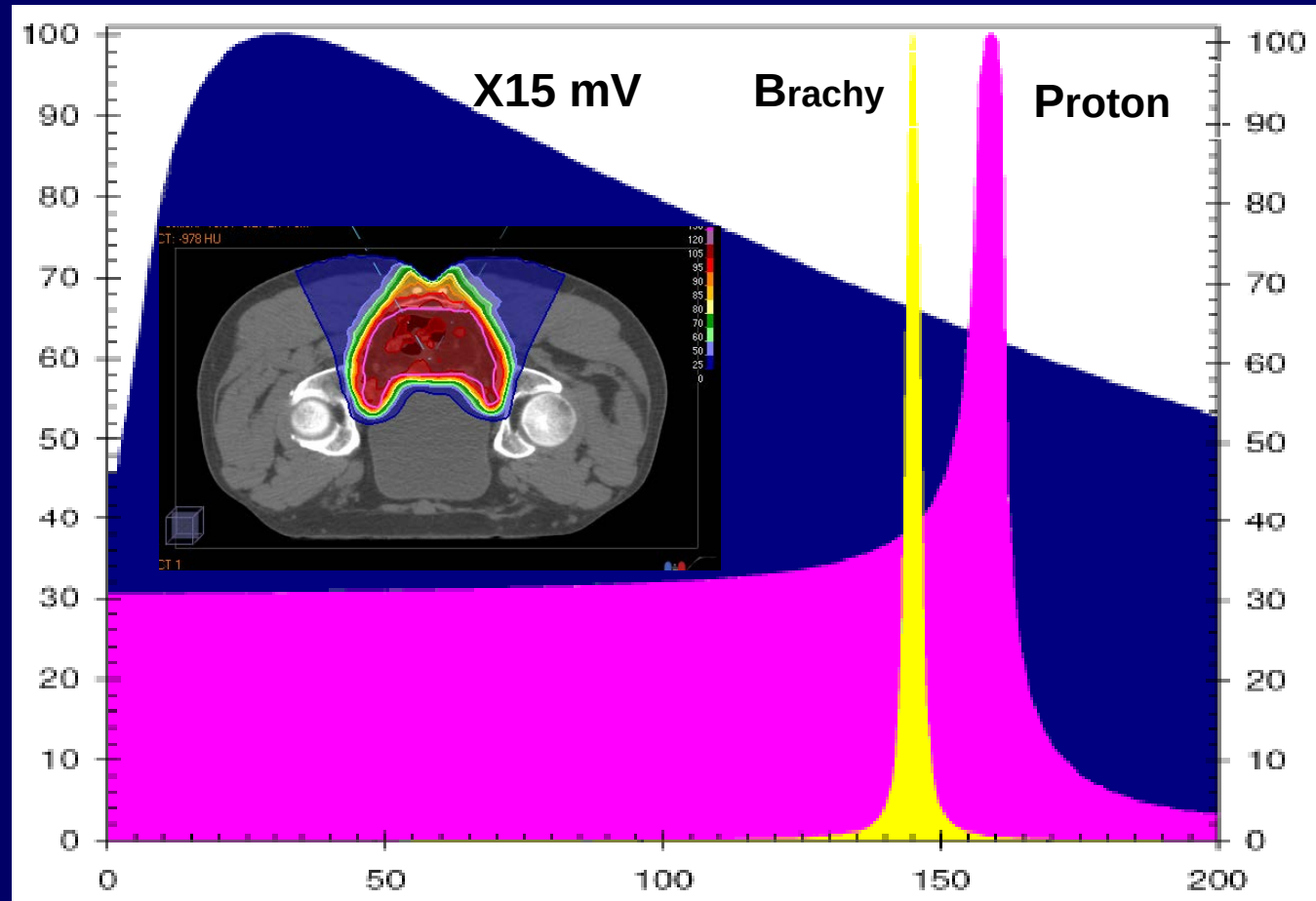
Annular shielding: 0.25 mm Pb. over 120 degree segment reduces radiation to <1%

Past-Present- Future : dose 100% PTV – 0% OAR

2024 !

Rectum:

Proton
+ Brachy



Brachytherapy – particle therapy-Hadron

But ...



EXPERT
REVIEWS

Renaissance of contact x-ray therapy for treating rectal cancer

Expert Rev. Med. Devices 8(4), 483–492 (2011)

Jean-Pierre Gérard¹¹,
Arthur Sun Myint²,
Olivier Croce³,
Jacob Lindegaard⁴,
Anie Jensen⁴,
Robert Myerson⁵,
Jean-Michel
Hannoun-Lévi¹
and Serge Marcie¹

Contact x-ray therapy (CXRT) with 50 kV has proven to be an efficient radiation therapy technique to achieve local control and rectal preservation for early rectal adenocarcinoma. Despite these results, CXRT has not been used due to the shortage of the no longer manufactured Philips RT 50™ unit. Recently, a new CXRT machine (Papillon 50™) became available on the market. This machine delivers a beam of 50 kV with a dose rate close to 15 Gy/min and has a percentage depth dose of 50% at 6–7 mm. The applicator size varies from 2–3 cm in diameter. Due to the original design of the main tube, treatment delivery is quick and more comfortable for the patients. An online viewing system incorporated in the tube allows a good visualization of the tumor with improved accuracy of radiation delivery. An international collaborative trial (Contact Endoscopic Microsurgery [CONTEM]) was set up to accrue approximately 300 cases

➤ **2001 Philips machine Stop**

2009 : Papillon 50™
UK- French design

Exp Rev Med Dev
2010; 34: 231
JP Gerard & al



Rectum External Beam RT (EBRT) BOOST ??

EBRT : - stereotactic Body RT (SIB)

- cyberknife
- proton



ACCORD 12 phase III : 45 Gy vs 50 Gy **ypCR** : 13% - 19%

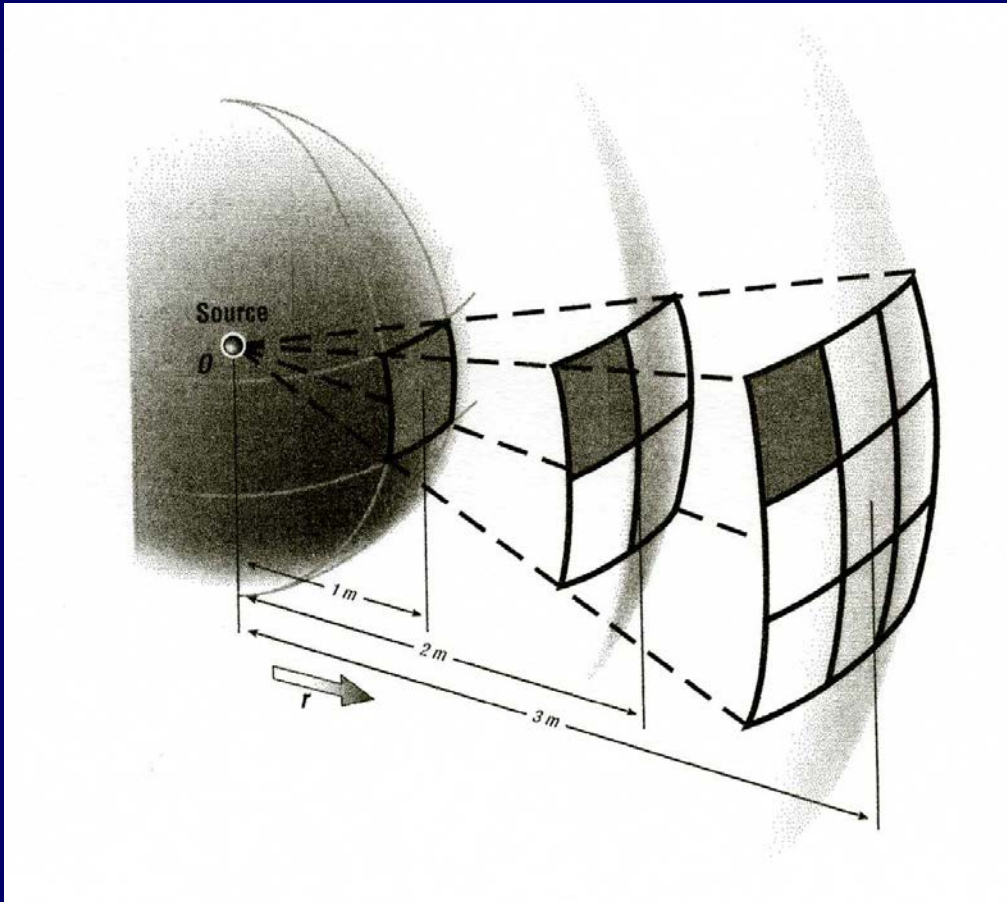
GRECCAR 12 « « : 50 Gy vs 60 Gy **R0** : 83% - 88%

But Toxicity is the ultimate Limit

With EBRT impossible to increase Dose > 80 Gy

Inverse Square Law $S : 4\pi r^2$

ISOTROPIC RADIATION source : sphere



1 cm $S : 1$ Dose **10 Gy**

2 cm $S : 4$ **25%:** **2.5**

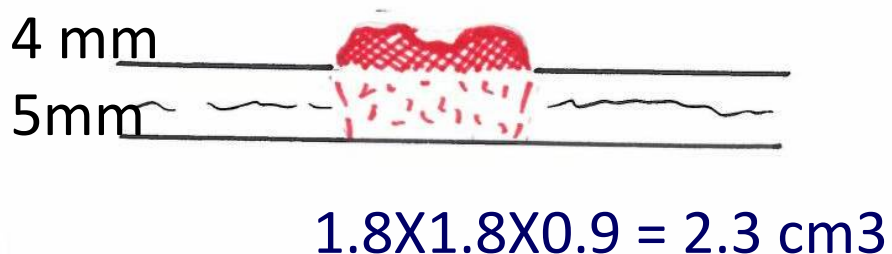
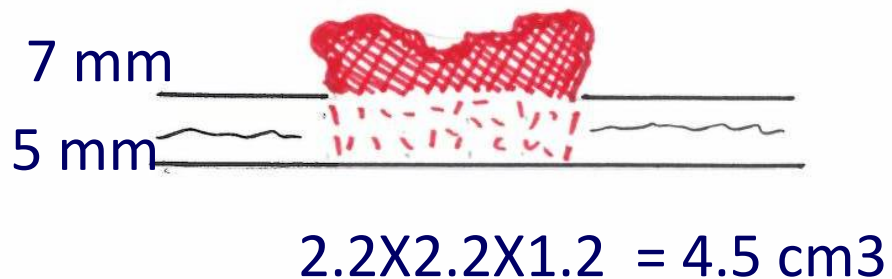
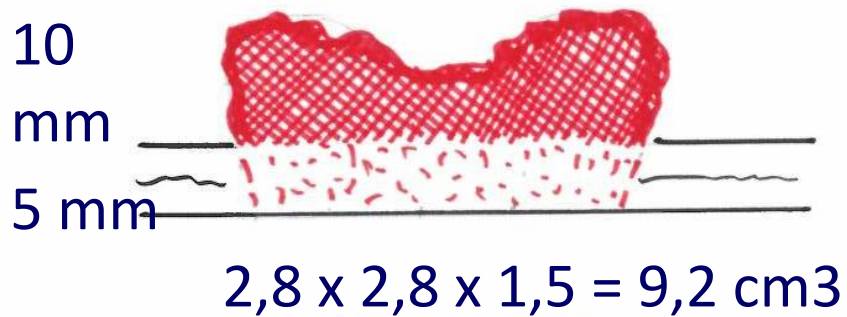
3 cm $S : 9$ **11%:** **1.1**

10 cm $S : 100$

1cm/10cm : **1% :**

1000 Gy : 10 GY

1mm : 15%



D 1
30 Gy

D 14
30 Gy

D 28
30 Gy

- « Tumor destroyed
layer by layer »
J Papillon

- Contact X ray Brachy

D1 – D 28 : 90 Gy / 3 Fr

Mean Dose: 48 Gy / 3 F

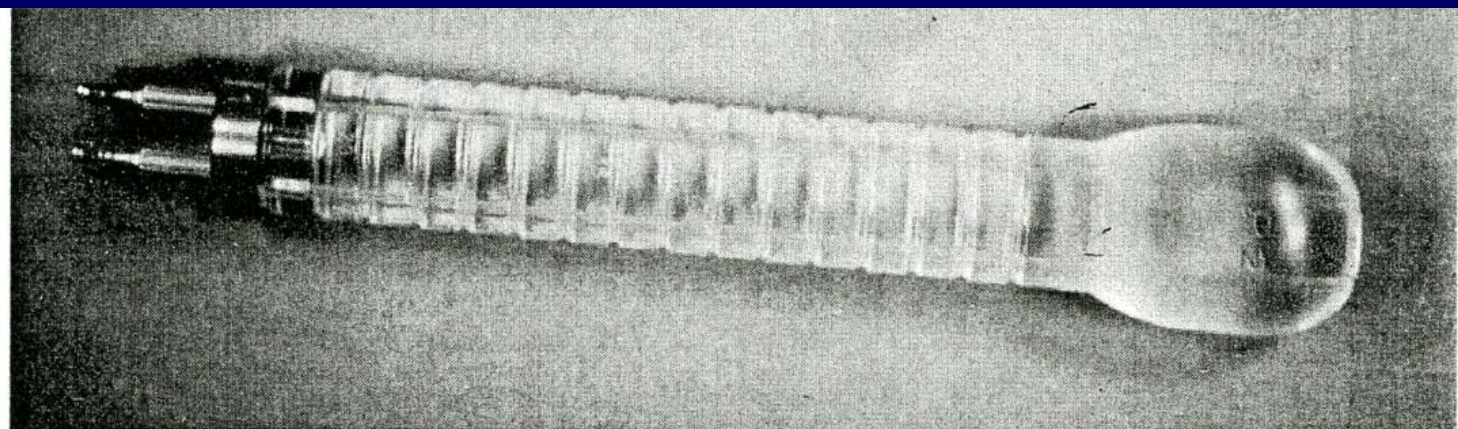
α/β : 10 (S Marcié)

EQD2 = 98 Gy

- Appelt T 3cm x 2 thick 30 Gy

α/β : 10 : EQD2 : **33.9 Gy**

**Papillon + 50 kV
Rectal Applicator
Lateral Beam
Stepping source
vs Iridium**

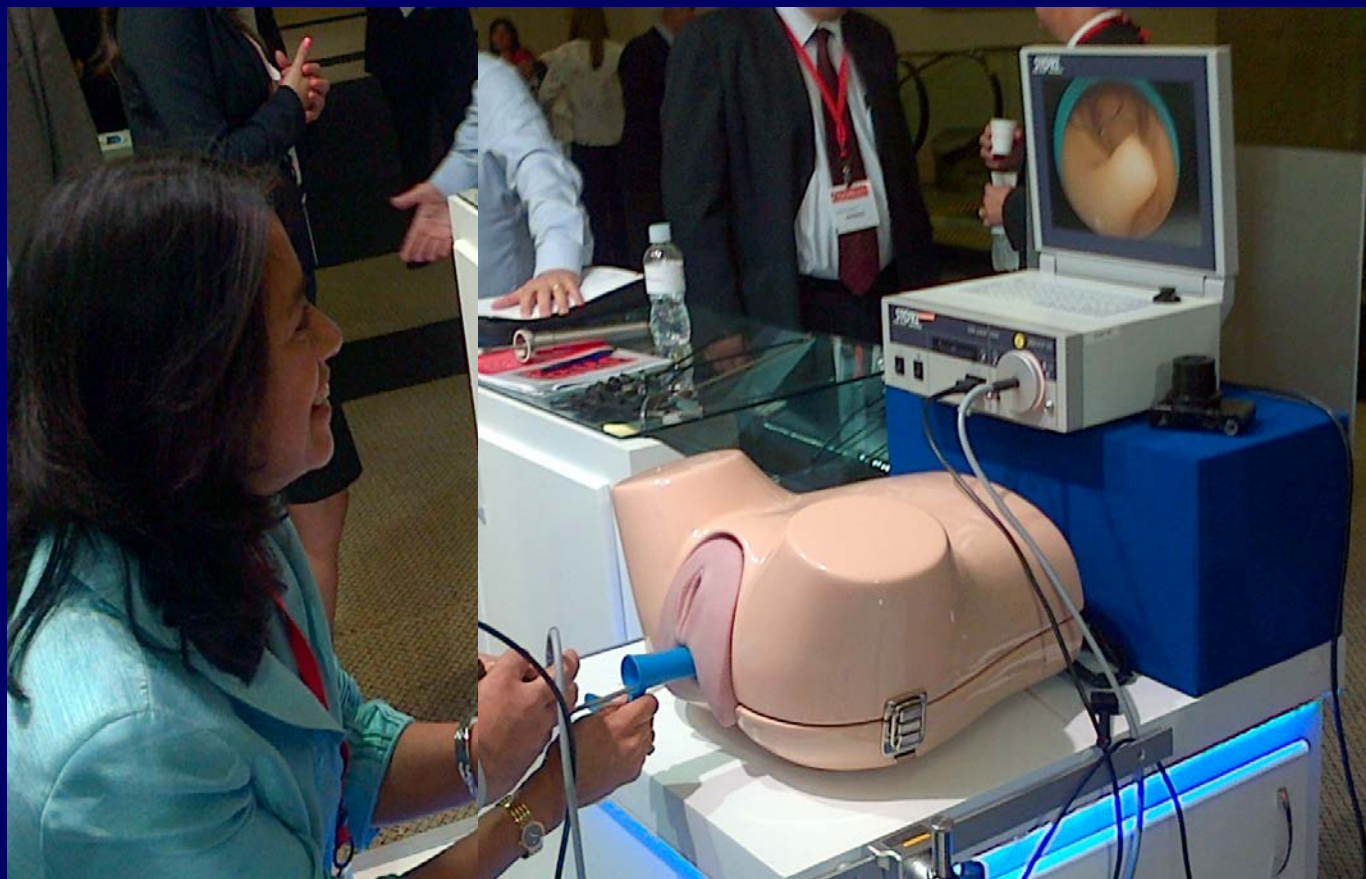


Organ preservation

- Topic of interest ++ (LARS Syndrome)
- To « hope » for organ preservation: first **Clinical CR**
- CRT alone : Sao Paulo; NY; UK; NL; France
 - IWWD : 1008 pts
 - cCR : 20-50% Loc Rec : 25% [15-40%]
- ADK rectum : **quite radioresistant tumor**
- RT dose escalation (toxicity) : BOOST

Isotropic Dose Distribution- Sphere Surface : $4 \pi r^2$

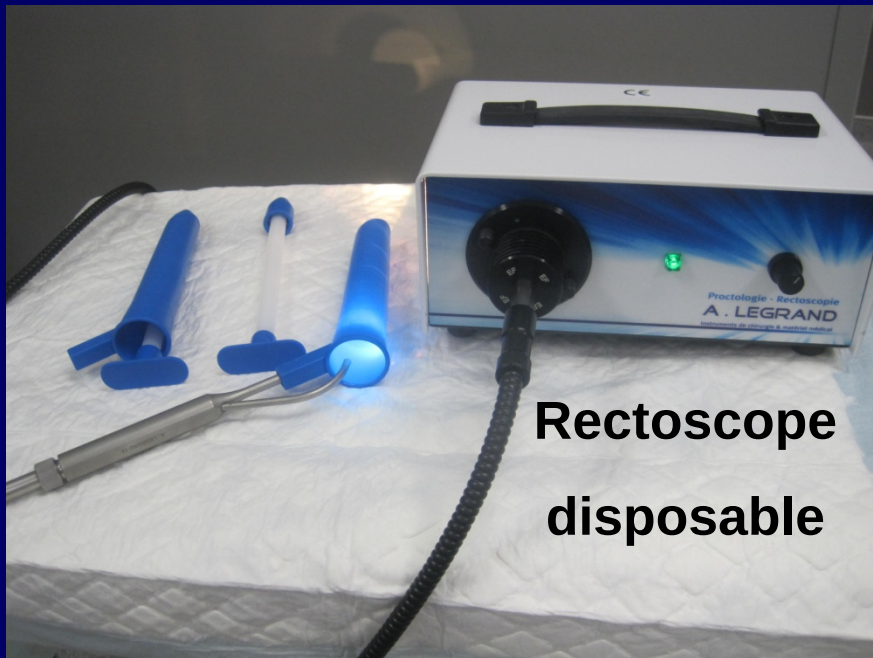
FSD 4mm	r	r ²	dose	↗surface	dose↘	dose
	4	16	100			100
5mm	9	81		81/16 : 5	5	20
10mm	14	196		196/16 : 12	12	8
FSD 40mm						
	40	1600	100			100
5 (45)	45	2025		2025/1600 : 1.2	1.2	83
10 (50)	50	2500		2500/1600 : 1.56	1.56	64



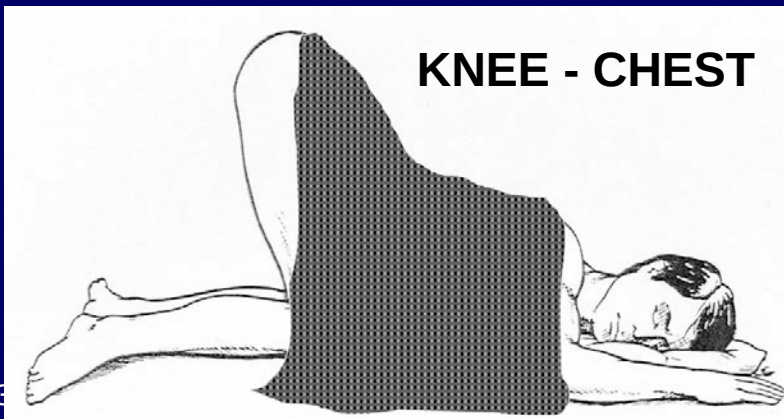
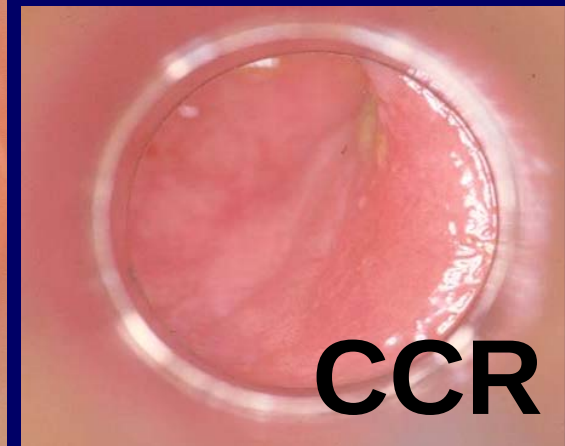
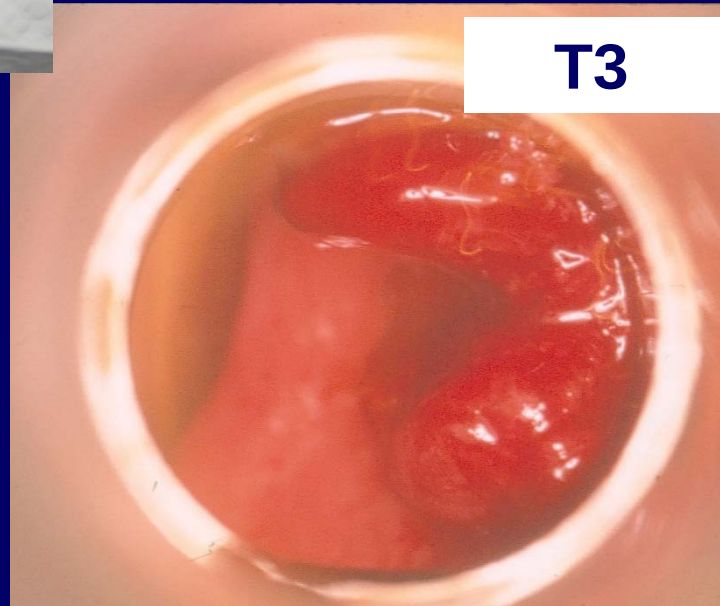
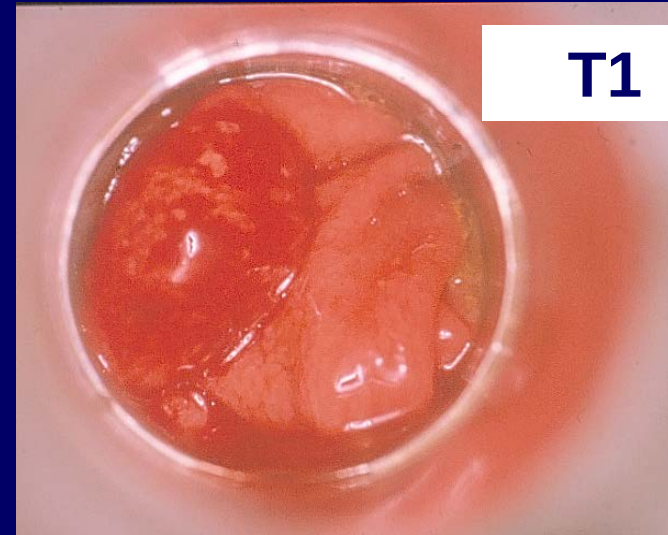
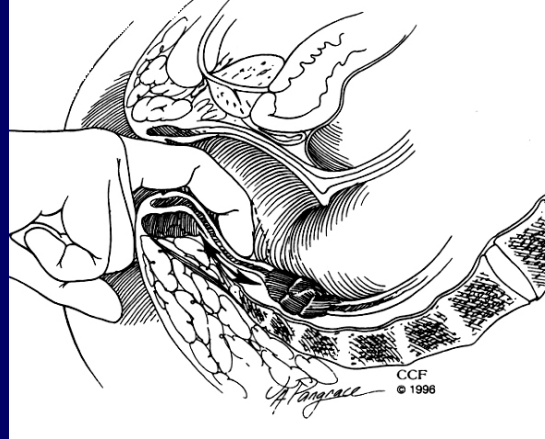
DRE + ED+NDOSCOPY :
Key examens
Rectoscope Legrand Paris



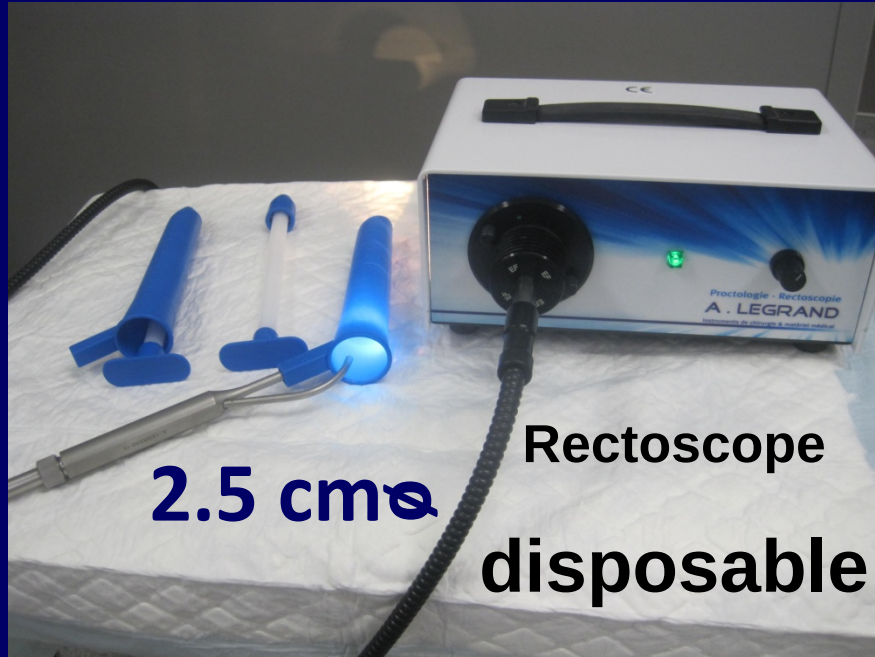
RECTOSCOPY + DRE in KNEE CHEST POSITION



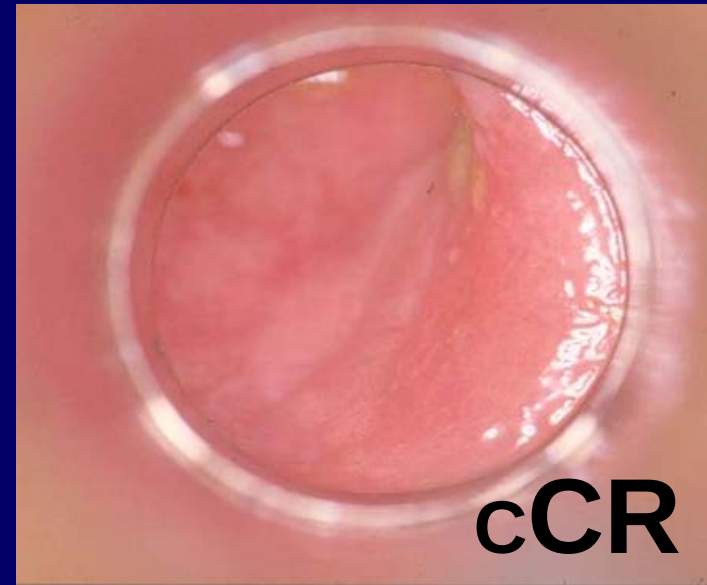
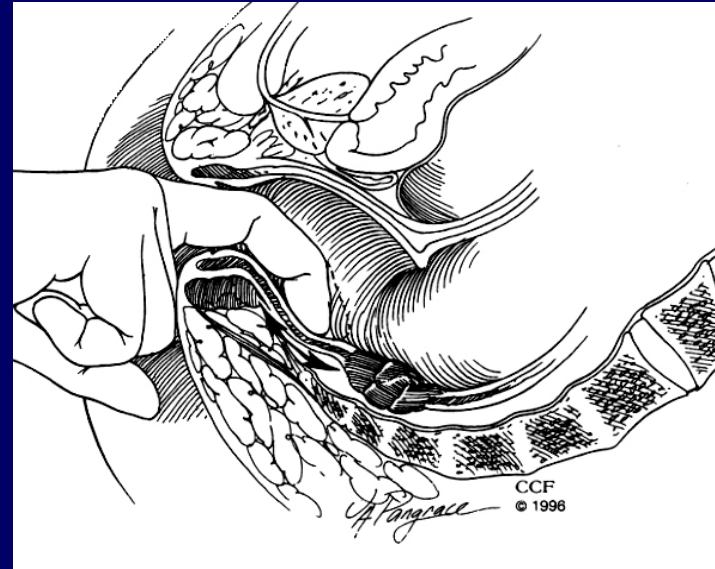
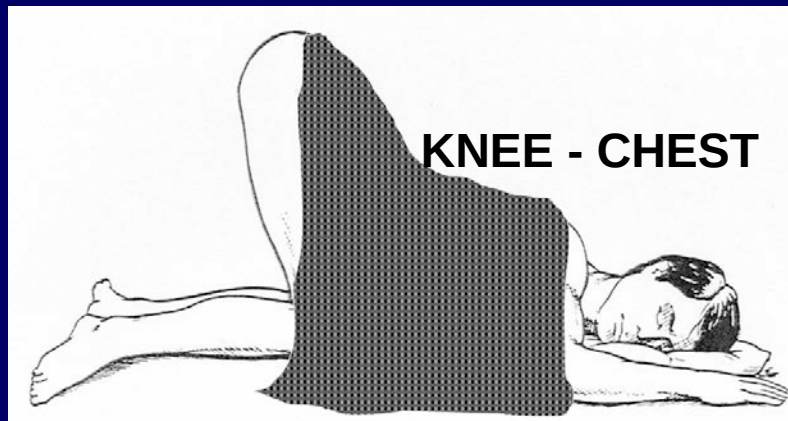
LEGRAND tm



Wek 14 : RECTOSCOPY + DRE + MRI

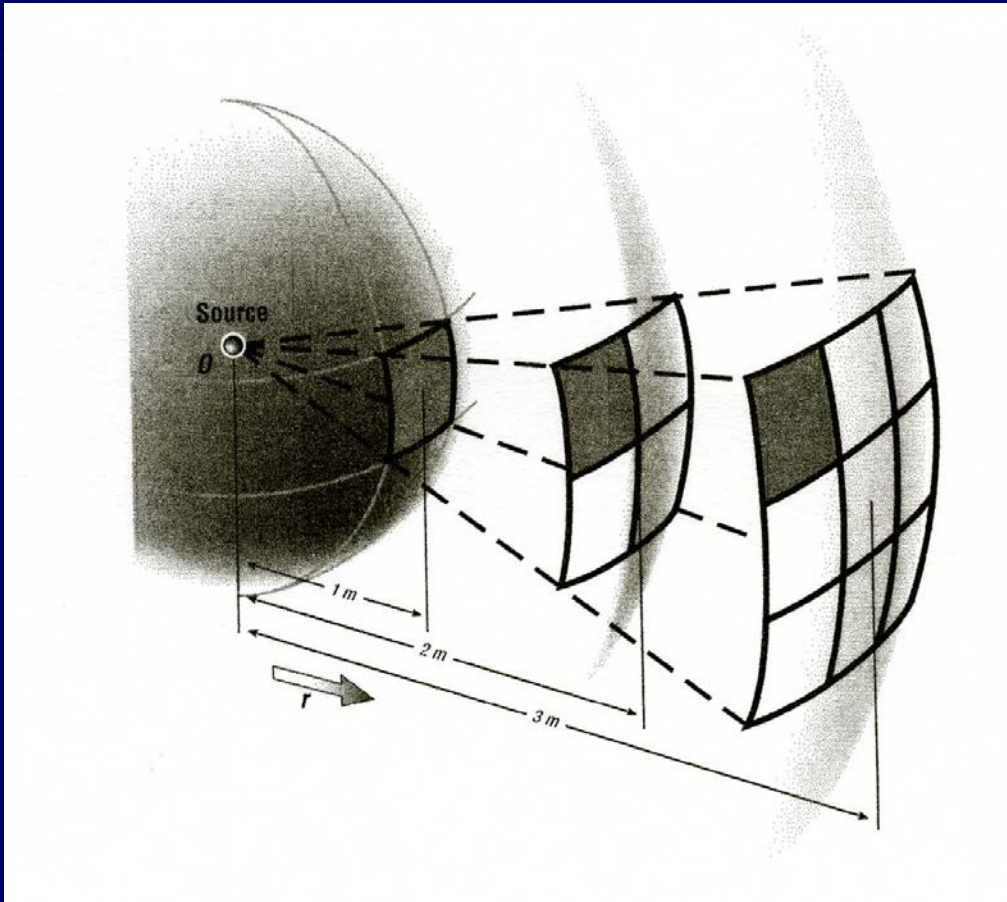


LEGRAND tm



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