



# Métastases hépatiques des tumeurs neuroendocrines :

*SIRT et ses perspectives*

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Université  
de Paris

Université  
Paris Cité

# Conflits d'intérêt

Consultant :

Boston scientific

Terumo

Merit Medical

Centre utilisateur exclusif de Therasphere®

# Remerciements

Oncologie digestive Cochin : Pr Coriat, Dr Pellat

Médecine nucléaire : Pr Cottreau – Dr Dechmi – Dr Monssarat

# Contexte et rationnel

# Les néoplasies neuroendocrines : généralités

Origine : Digestive (2/3)

Intestin grêle  
Pancréas – Appendice  
Estomac – rectum  
Œsophage - foie – Voies biliaires

1,45/100 000/an  
0,95/100 000/an  
0,45/100 000/an  
Exceptionnelles

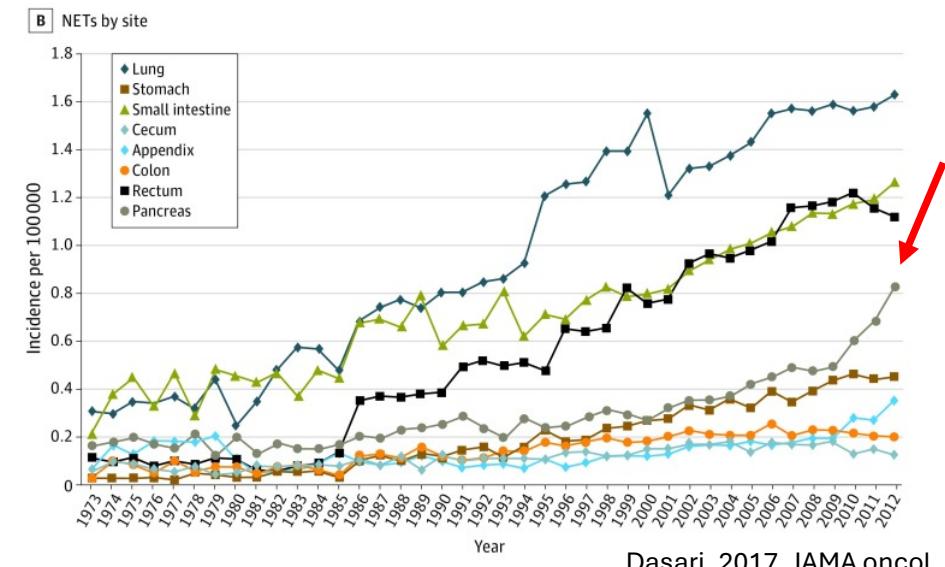
Autre  
Paragangliome, pulmonaire, sein, prostate, primitif inconnu

Présentation hétérogène :

Symptômes et stade au diagnostic  
Prédisposition héréditaire  
Sécrétion fonctionnelle (20% des pTNE – plus pour le grêle) et récepteurs à la somatostatine

Pronostic variable :

Différenciation histologique (TNE / CNE)  
Ki67 et grade (>3% ; 3-20%, ≥20%)  
Stade tumoral  
Organe d'origine



Dasari, 2017, JAMA oncol

Survie globale à 5 ans :

Pancréatiques : 25 à 75%  
Iléales : 55% à 70%  
CNE : 5 à 40%

# Les néoplasies neuroendocrines : particularités

Rare et hétérogène : peu d'essais randomisés – beaucoup d'accords d'experts

- ➔ RCP RENATEN-ENDOCAN – Réseau TENpath
- ➔ Inclusion dans des essais +++



Survie longue : objectifs thérapeutiques personnalisés :

Guérison

Augmentation de la survie

Contrôle local

Contrôle des symptômes

Amélioration / maintien de la qualité de vie...



# Bilan préthérapeutique

## Radiologie :

Scanner : Abdomino-pelvien injecté 30 et 70s

Thoracique

IRM :  
Abdominale – Foie ( $\pm$ Hépatospécifique)  
Pelvienne : TNE rectales  
*Cérébrale et rachidienne : si symptômes ou surveillance si connue*

**Primitif**

## Endoscopie digestive :

EOGD – iléocoloscopie : diagnostic, biopsies, complications

Echoendoscopie : biopsies – résecabilité – Si imagerie normale

**Résécabilité**

## Imagerie nucléaire :

PET au  $^{68}\text{Ga}$ -DOTATOC : bilan de toute NNE métastatique ( $^{177}\text{Lu}$ -DOTATATE)

PET au  $^{18}\text{FDG}$  : Pronostic indépendant des TNE, préthérapeutique des CNE

PET au  $^{18}\text{F}$ -DOPA : meilleur sensibilité que DOTATOC pour grêle : lésion introuvable, bilan exhaustif

*Essai clinique en cours BRD 11/5-K : TEP 68Ga-DOTANOC*

**Extension**

ETT : Bilan des complications

**Complication**

# Synthèse et traitement

Décision thérapeutique complexe basée sur tous les éléments cités



## Du syndrome sécrétant

Antisécrétant

Symptomatique

## Du primitif

Non métastatique

Résection endoscopique

Chirurgie ± Curage

Métastatique

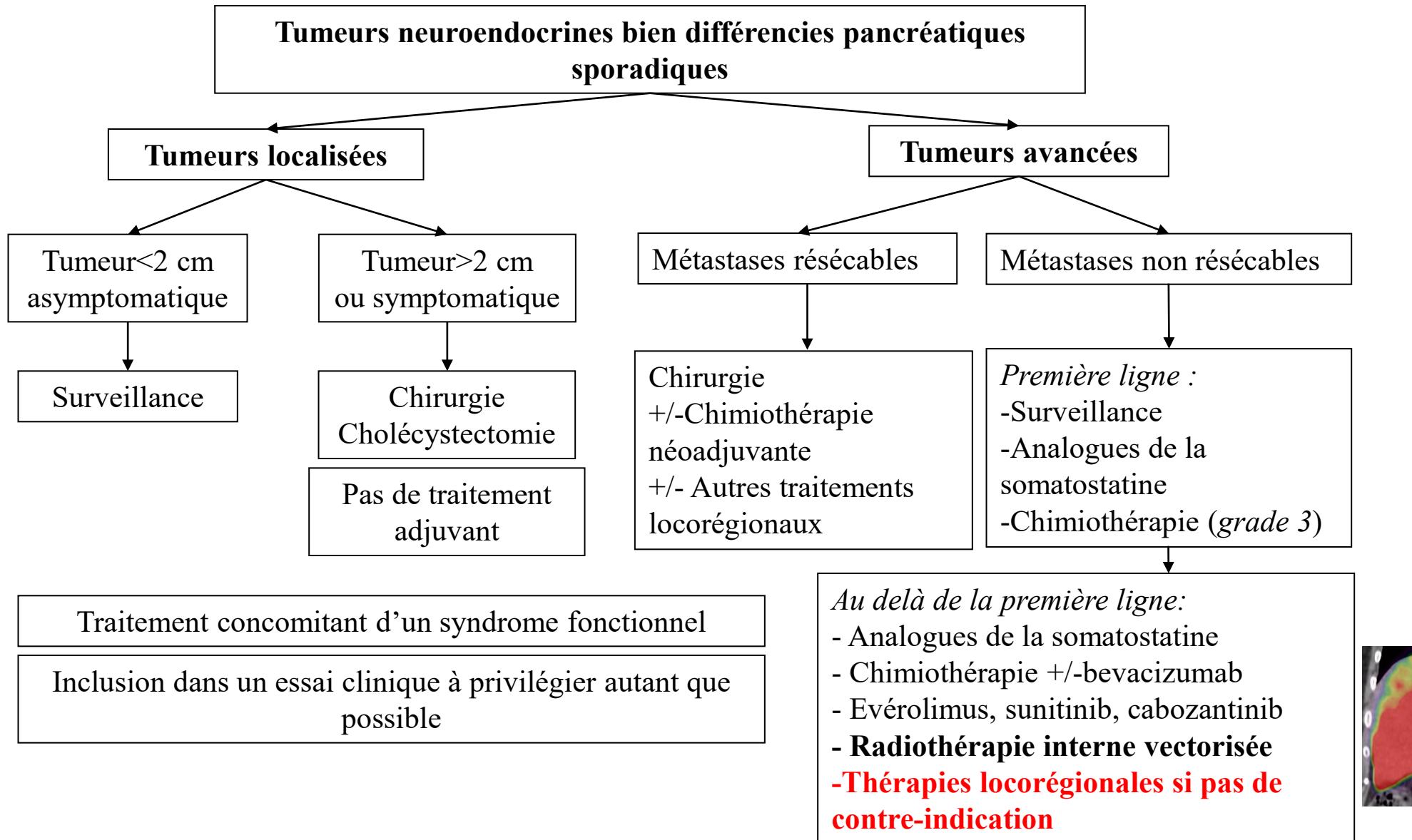
Exérèse du primitif

## Des métastases

*Place de la SIRT*  
Indication  
Progression

Symptômes/ sécrétion non contrôlée  
**Envahissement hépatique >50%**

# Synthèse et traitement



# Pourquoi un traitement « ciblé » des métastases hépatiques?

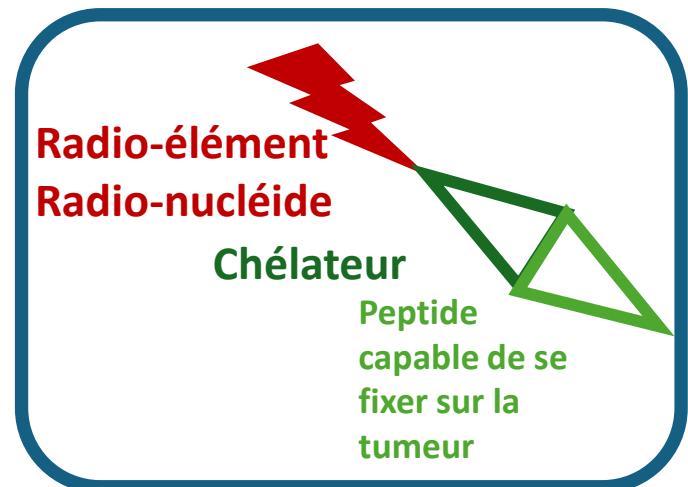
Pronostic  
*si invasion hépatique majeure*

Dénutrition  
Complication cardiaques  
**Insuffisance hépatocellulaire**  
Compression osseuse

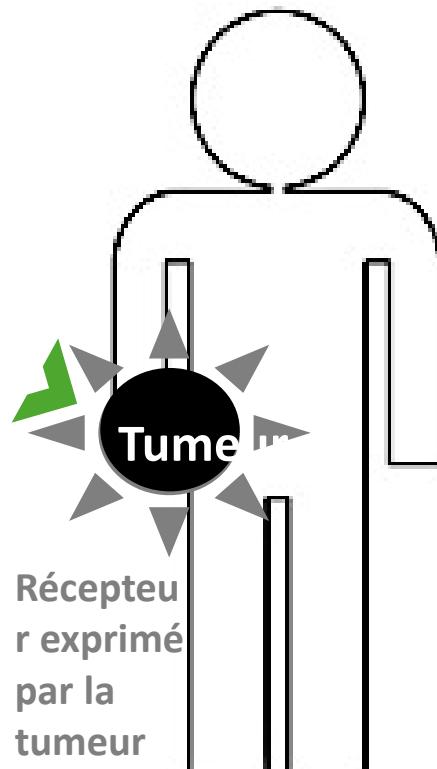
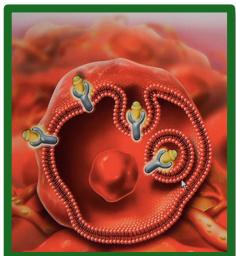


**Jusqu'à 25% COD tout patients confondus\***

# Radiothérapie interne vectorisée (RIV)



TRAITEMENT anti-cancéreux



## Radiothérapie métabolique par Lu<sup>177</sup> et TNE bien différencierées

combinaison d'une molécule vectrice dirigée spécifiquement sur une cible, avec un isotope radioactif

- Ciblage des récepteurs SST
- Vecteur : agoniste en routine clinique du sous-type 2 des récepteurs à la somatostatine
- Isotope : Lutetium 177

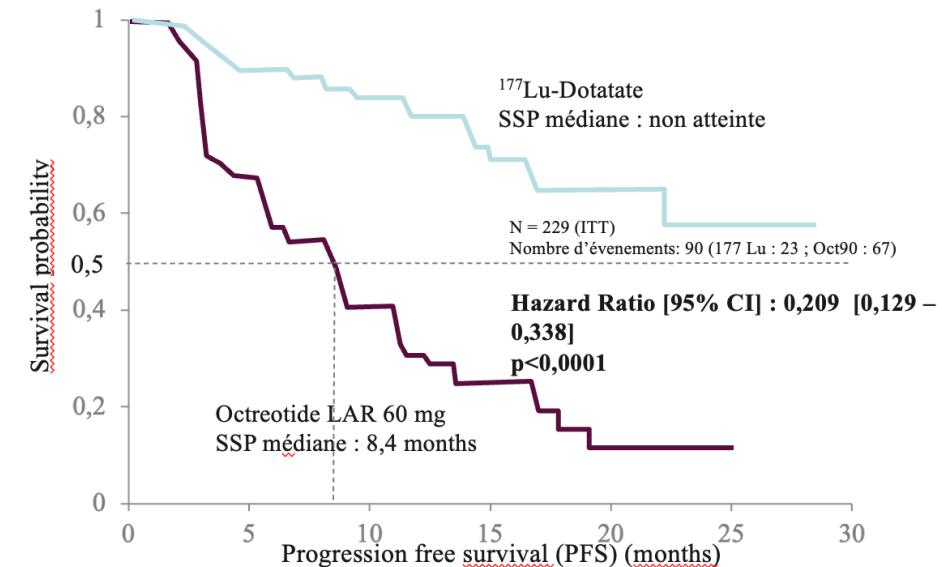
# Radiothérapie interne vectorisée (RIV)

**NETTER 1 :** 1<sup>ère</sup> étude randomisée sur la RIV phase 3 / preuve d'efficacité - TNE du grêle G1 G2

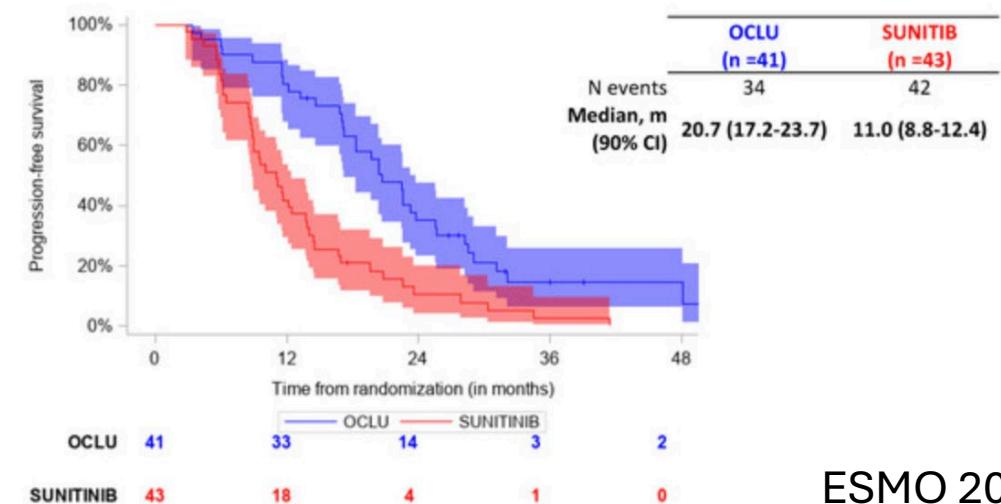
Conforté par **Netter 2** pour les G2 et G3

Strosberg J. et al. –NEJM 2017

**Occlurandom :** Phase II : étude de phase 2, randomisée comparant l'efficacité et la tolérance de la RIV versus sunitinib - TNE pancréatiques



Progression-free survival : real time blinded central review RECIST 1.1



ESMO 2022

# Radiothérapie interne vectorisée (RIV) : remboursement

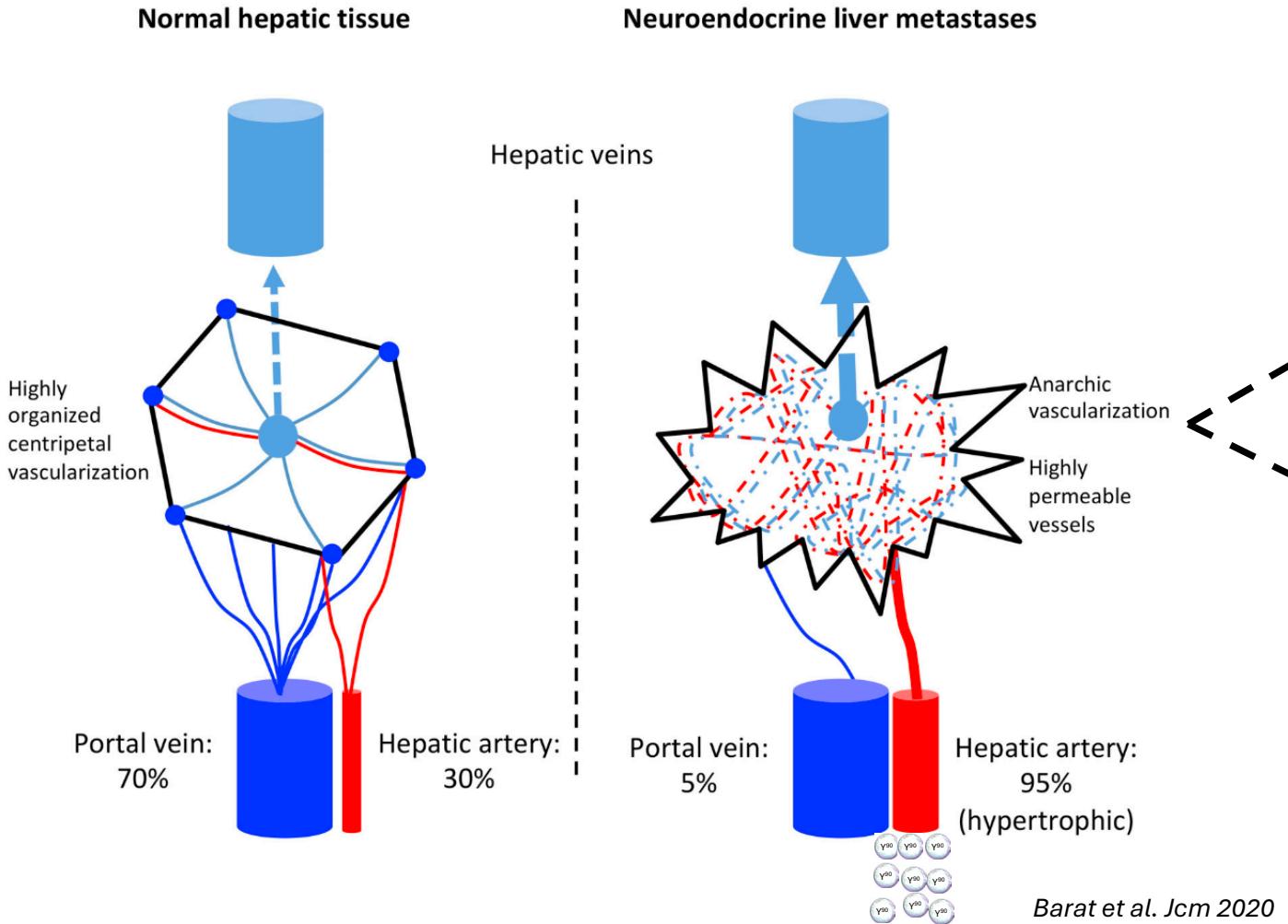
Remboursement : TNE du grêle uniquement

- 2ème ligne, après progression de la maladie avec octréotide.
- 1ère ligne si tumeurs d'emblée progressives ou avec une masse tumorale hépatique importante (> 50 %).

Doses compassionnelles :

- Phéochromocytome/paragangliome (PPGL) métastatique ou localement avancé inopérable
- TNE **bronchique**, métastatique ou localement avancée inopérable, progressive ou de sécrétante non contrôlée
- TNE **thymique**, métastatique ou localement avancée inopérable, progressive ou de forme sécrétante non contrôlée et exprimant les récepteurs de la somatostatine sur l'imagerie TEP des récepteurs de la somatostatine, en relation avec les résultats de la TEP au FDG et sur proposition de la RCP nationale Renaten.
- TNE y compris une TNE de **primitif inconnu**, NE correspondant **PAS à l'indication** de l'AMM
- Méningiome de tous grades, exprimant les récepteurs de la somatostatine de type 2

# La radio-embolisation : rationnel



Risque de shunt veineux :

- Poumon +++
- Digestif
- Autre

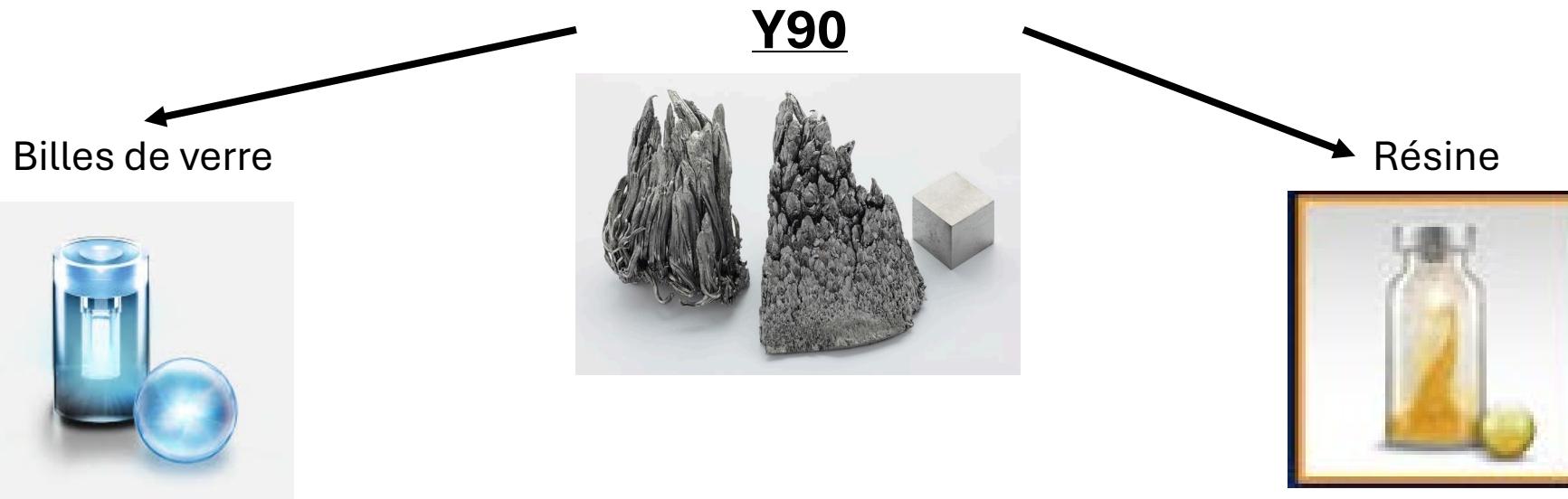
→ Irradiation non cible

Recrutement artériel anarchique

- Hépatique multiple
- Extra-hépatique
  - Digestif
  - Diaphragmatique
  - ...

→ Irradiation foie « sain »

# SIRT et TNE : Devices disponibles



Ho 166



Terumo Announces Perma  
Product Range Due to Oper

tion of Holmium Platform SIRT  
city

# Cas clinique

# Cas clinique

Homme 53 ans

TNE du grêle (duodénale) de grade II (ki-67 : 14%)

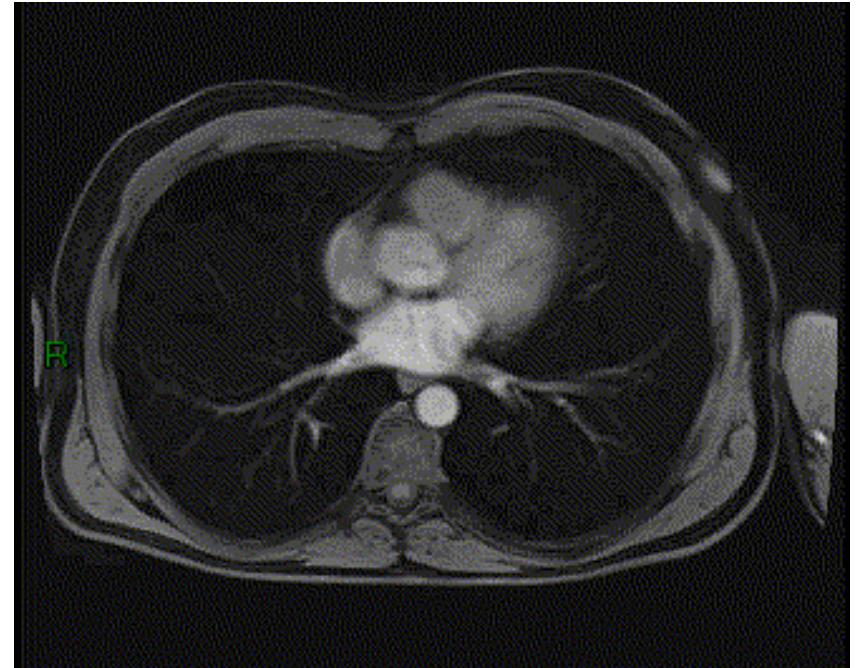
DPC en 2015.

Métastases hépatiques en 2019 : Xeloda – Temodal

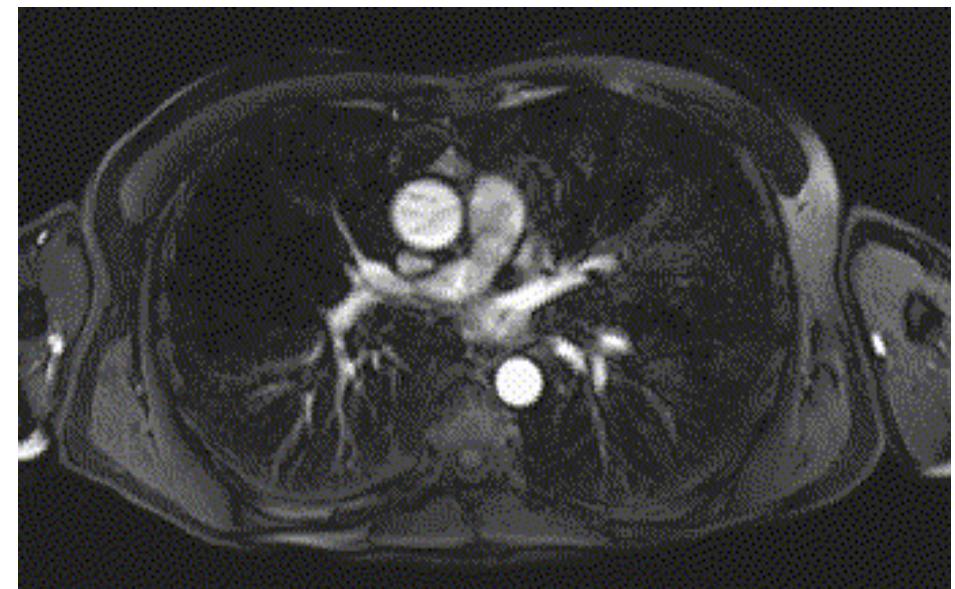
Progression en 2021 : Everolimus puis Sutent

2023 : Progression isolée et douloureuse d'une lésion du VI

2022



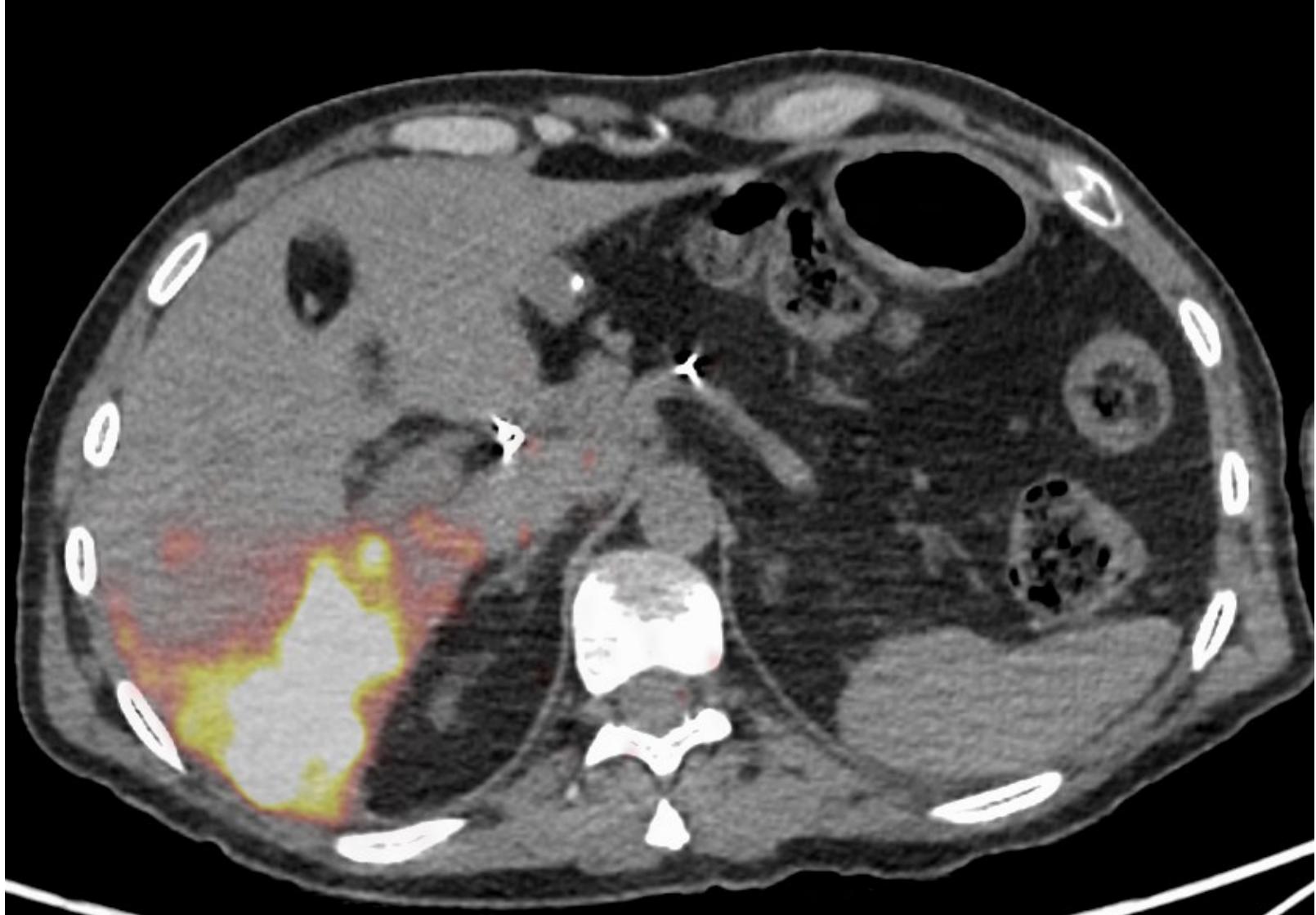
2023



# Cas clinique

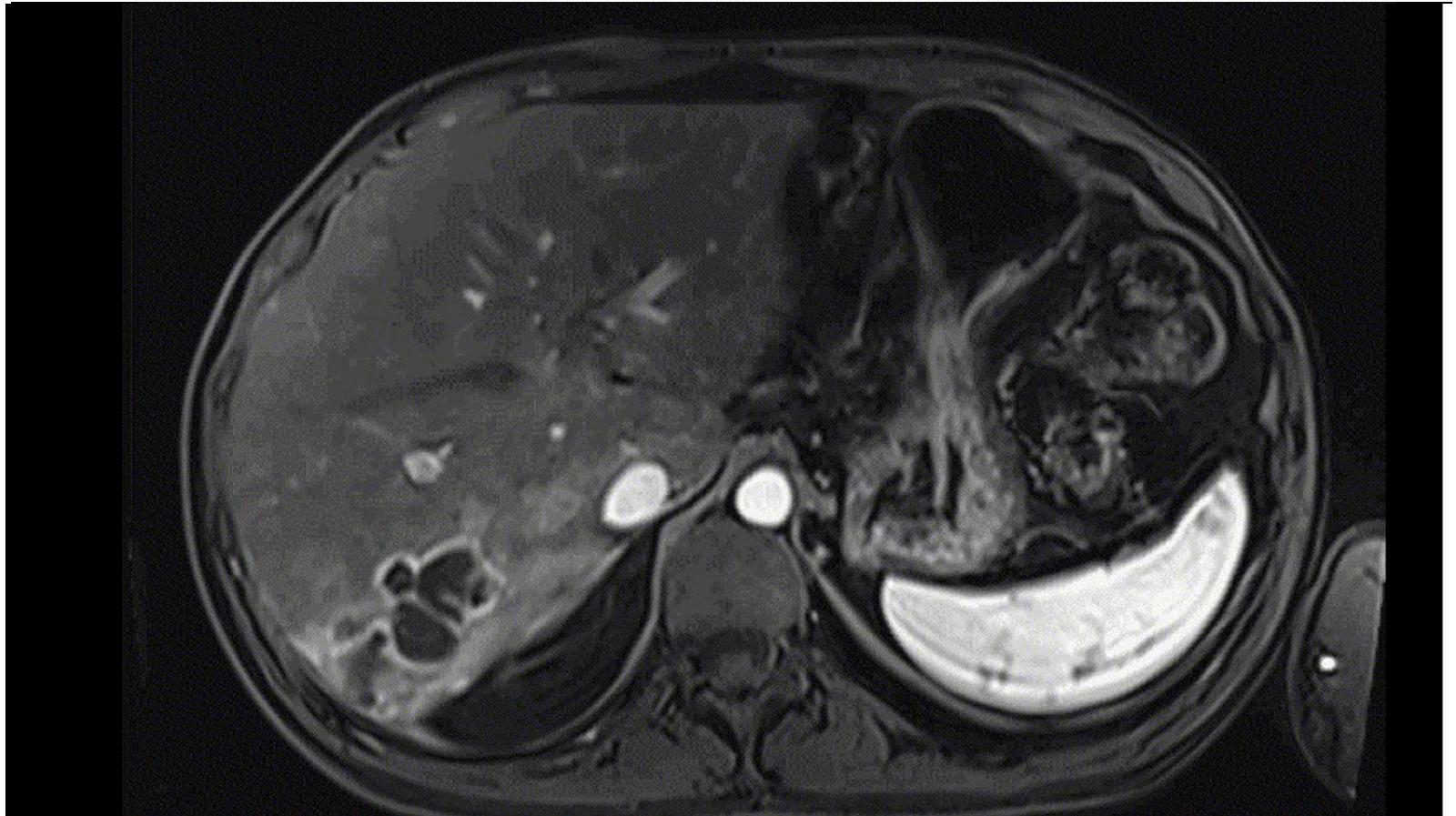
Traitement : segment VI

335 Gy à la tumeur



# Cas clinique

IRM 3 mois

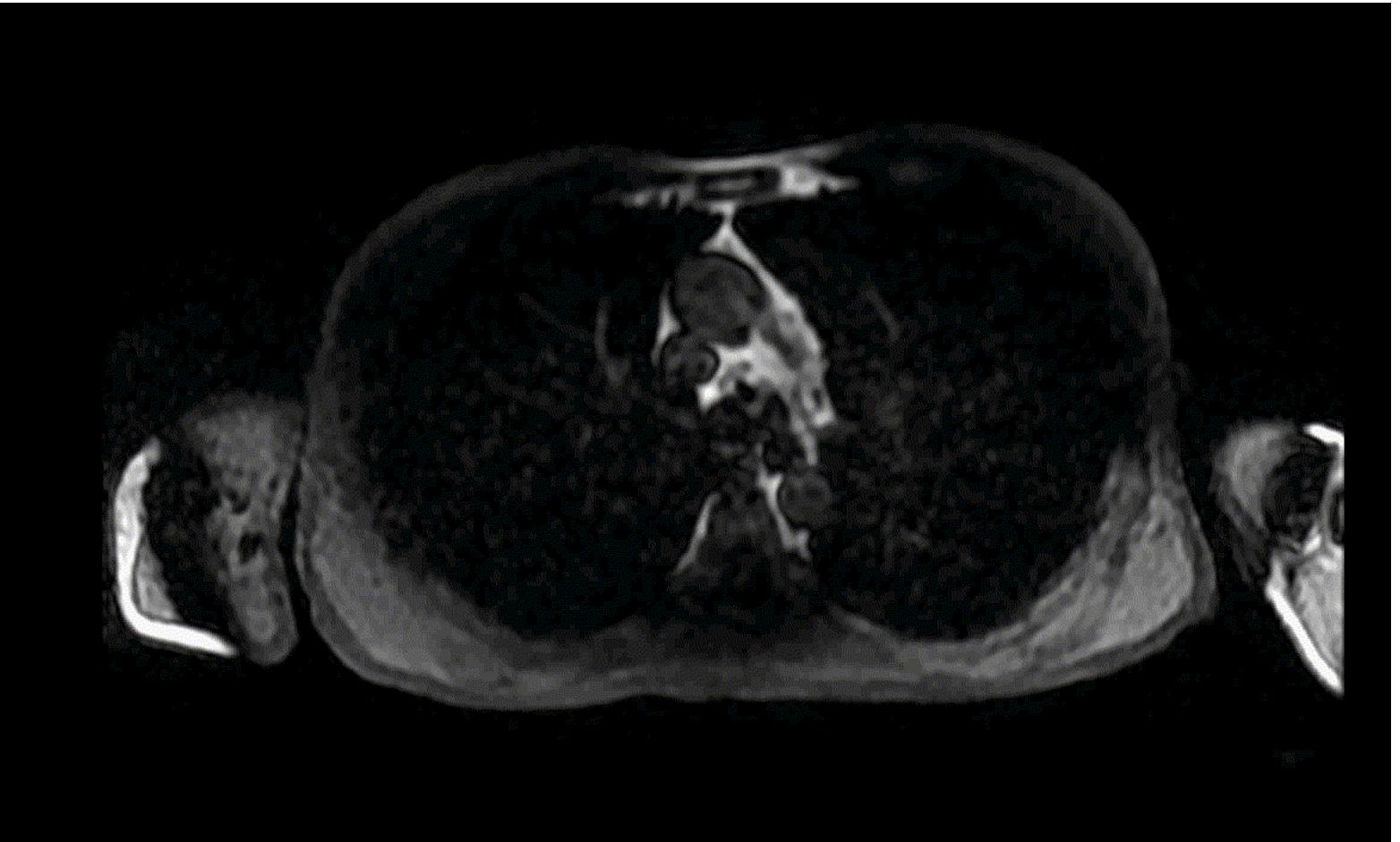


# Cas clinique

Progression des lésions non traitées :

Carbozantinib 60mg/j

IRM à 1 an : nécrose segmentaire

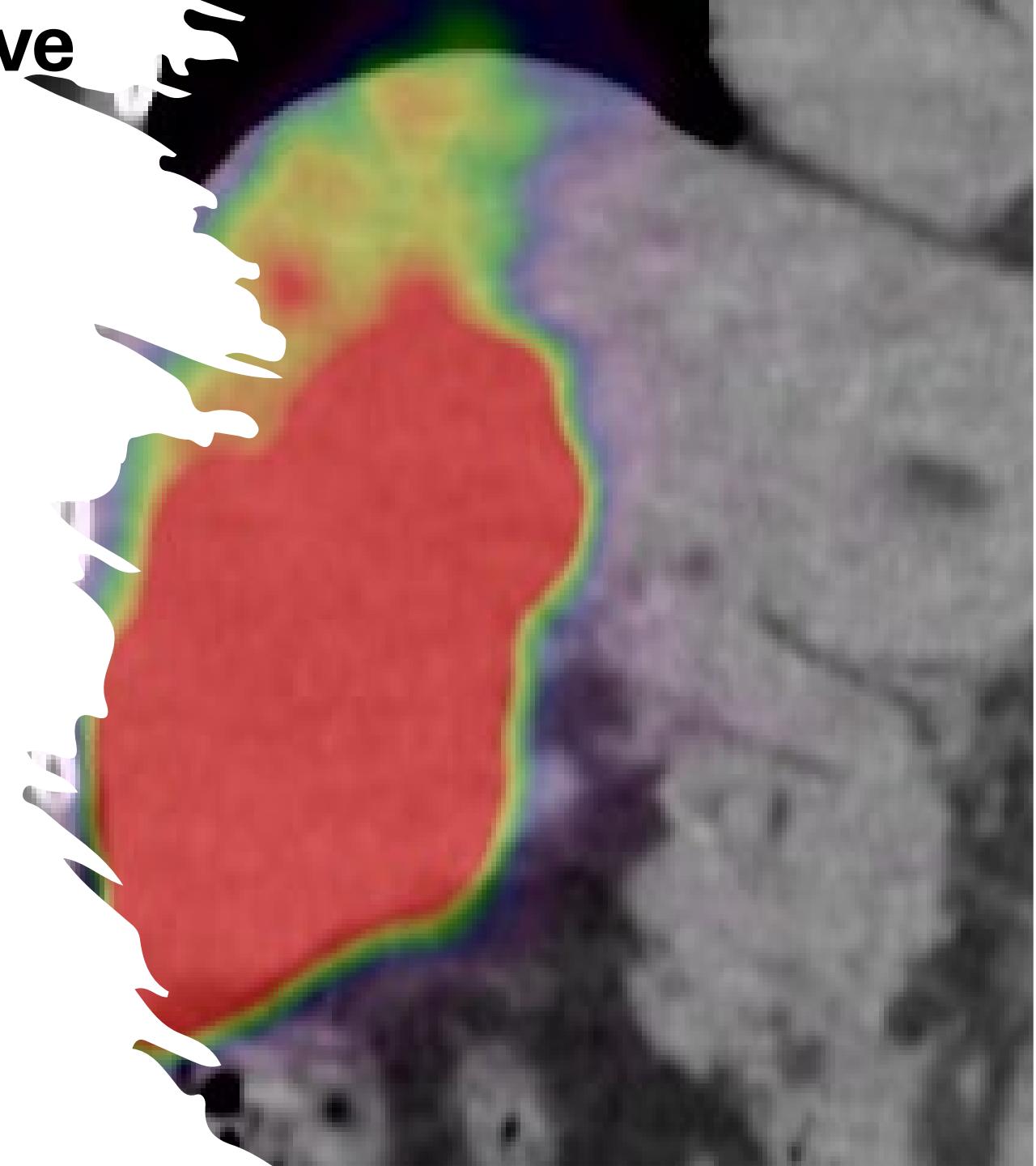


# Connaissances actuelles

# SIRT et TNE : niveau de preuve

**Difficultés :** Pathologie rare  
Peu d'évènements « décès »

**Conséquences :**  
Peu de patients inclus  
Rétrospectif  
Critère de substitution « DFS »



# SIRT et TNE : niveau de preuve – « Glass »

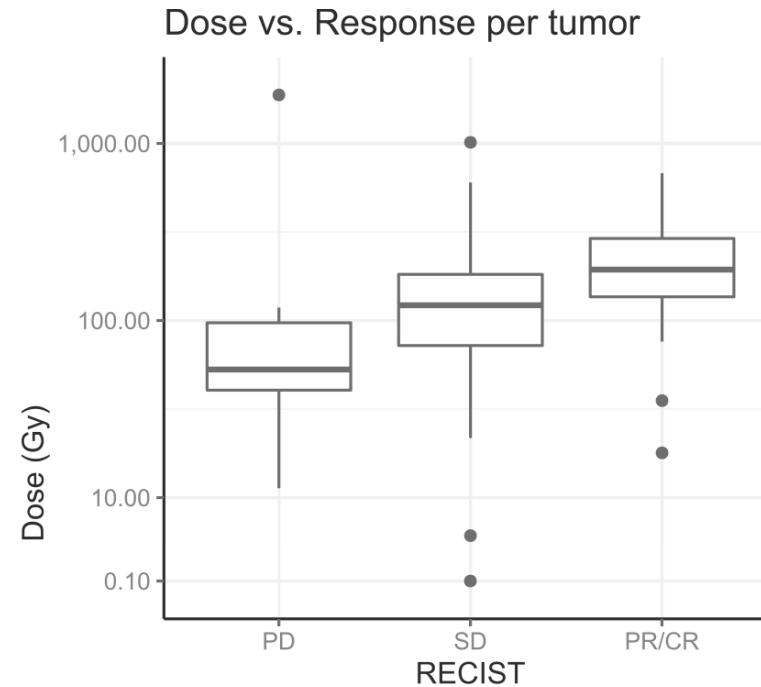
Ebbers et al. EJNMMI 2021

Rétrospectif, monocentrique

128 TNE (toute origine) – 26 patients – 31 SIRT

CJP : réponse RECIST 1.1

>150 (voir 170) Gy pour Réponse



Watanabe et al. J Nucl Med 2024

Rétrospectif 99 patients

→ Même trend

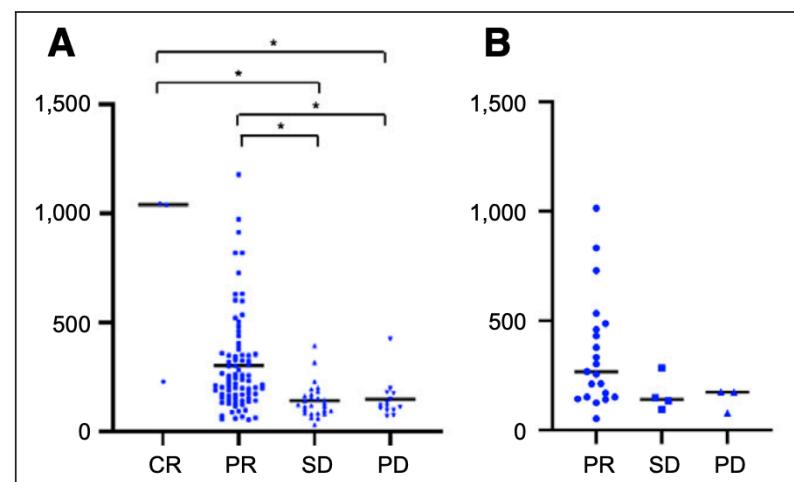


FIGURE 2. Lesion-based (A) and patient-based (B) comparisons of MAD

# SIRT et TNE : niveau de preuve – « Glass »

Egger et al. 2020

Rétrospectif, 2 centres 2000 - 2018

248 patients (TNE toute origine)

197 TACE

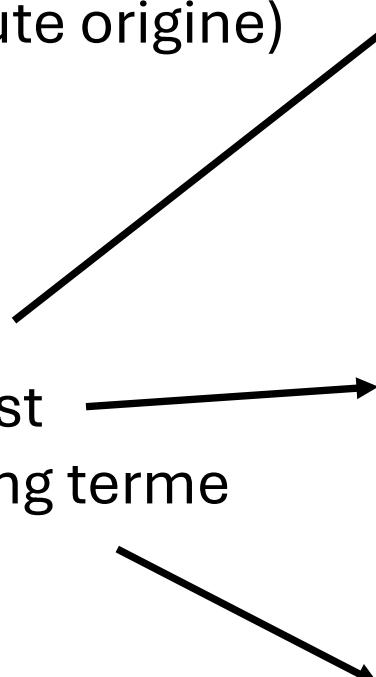
vs 51 SIRT

Évaluation :

Morbidité

Réponse Recist

Résultats à long terme



**Table 2.** Periprocedural Outcomes after Transarterial Chemoembolization vs Transarterial Radioembolization among Patients with Neuroendocrine Liver Metastases

Outcome	TARE (n = 51)	TACE (n = 197)	p Value
Length of stay, d, median (IQR)	0 (0, 0)	1 (1, 1)	<0.0001
Any complication, n (%)	7 (13.7)	44 (22.6)	0.17
Major complication, n (%)	3 (5.9)	18 (9.2)	0.58
30-d mortality, n (%)	1 (2.0)	6 (3.1)	1.0
90-d mortality, n (%)	5 (9.8)	10 (5.2)	0.21
Laboratory, median (IQR)			
Bilirubin change, mg/dL	0 (-0.3, +0.1)	+0.4 (+0.1, +0.8)	<0.0001
Platelet change, 10 <sup>3</sup> /µL	-29 (-78, +19)	-42 (-82, -4)	0.31
INR change	+0.1 (-0.1, +0.3)	+0.2 (+0.1, +0.3)	0.07
Creatinine change, mg/dL	-0.1 (-0.2, 0)	+0.1 (0, +0.2)	<0.0001
% chromogranin change	-16 (-64, +49)	-43 (-77, -4)	0.07
Radiographic			
% change in size, median (IQR)	-9 (0, -27)	-19 (-6, -34)	0.051
RECIST response, n (%)			0.0002
Complete response	2 (4.4)	5 (3.6)	
Partial response	9 (19.6)	37 (26.6)	
Stable disease	27 (58.7)	92 (66.2)	
Progressive disease	8 (17.4)	5 (3.6)	

INR, international normalized ratio; IQR, interquartile range; RECIST, response evaluation criteria in solid tumors; TACE, transarterial chemoembolization; TARE, transarterial radioembolization.

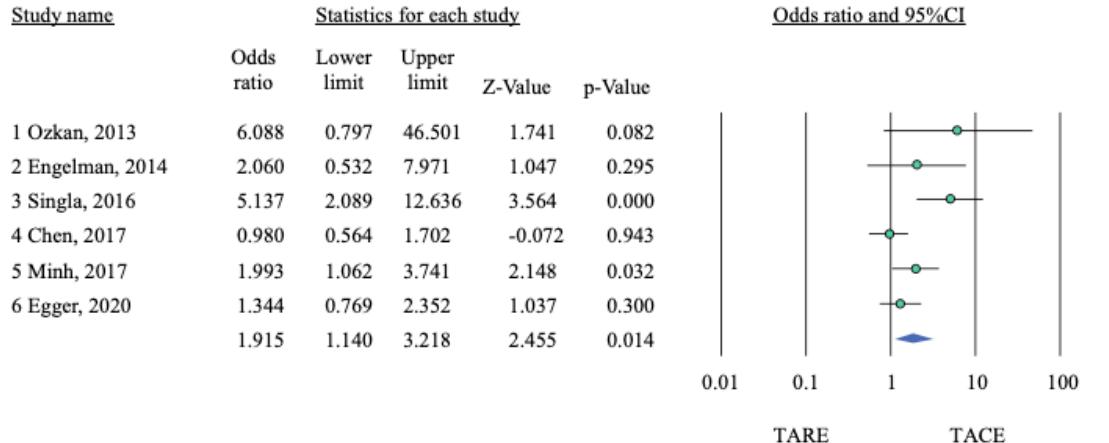
Pas de différence

# TNE : SIRT versus TACE

Ngo et al. An Surg Oncol 2021

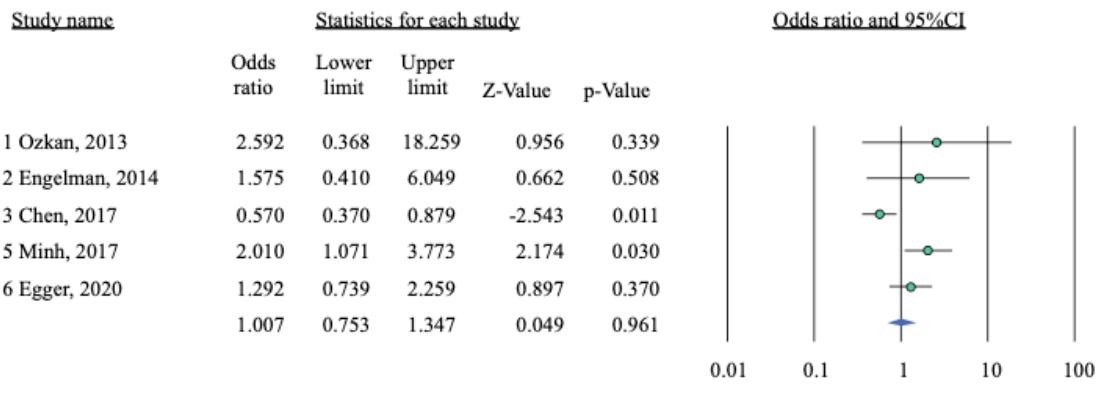
(a)

## Median overall survival



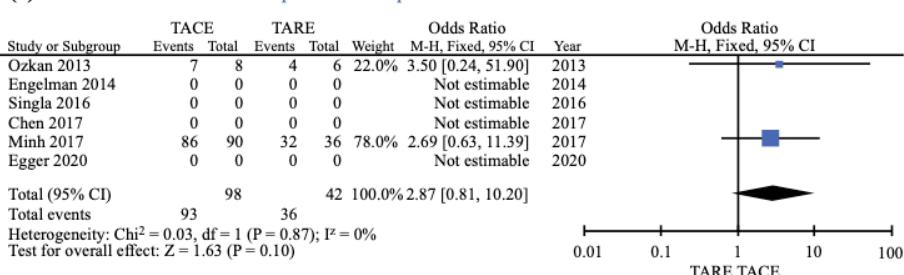
(b)

## Median hepatic progression-free survival



(a)

## Hepatic tumor response within 3 months of treatment



(b)

## Hepatic tumor response more than 3 months after treatment

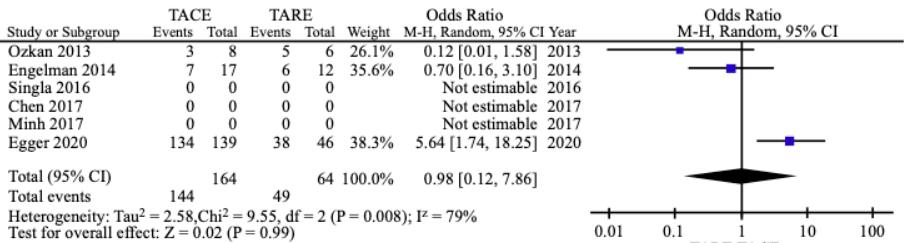
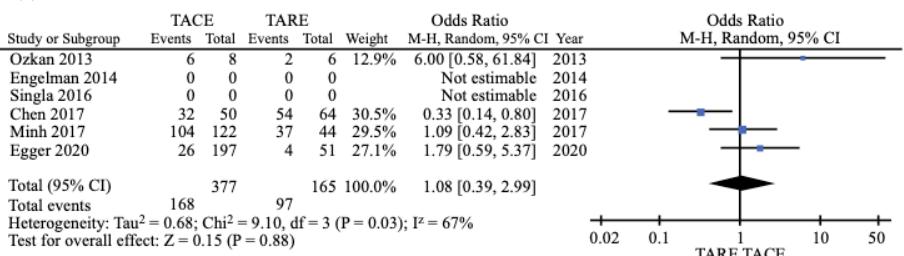


FIG. 4 Meta-analysis of hepatic tumor response. a Within 3 months of treatment. b More than 3 months after treatment

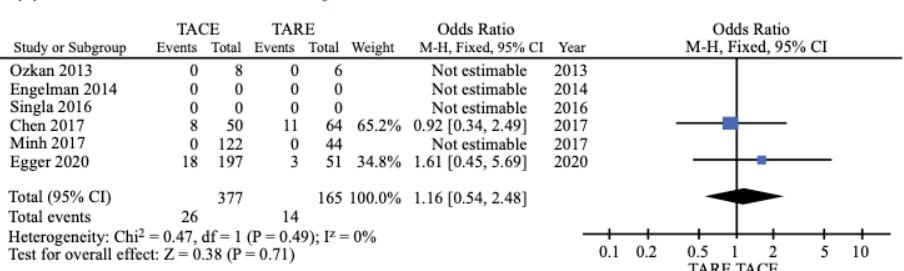
(a)

## Minor adverse events



(b)

## Major adverse events



# SIRT et TNE : niveau de preuve – « Resin »

Braat et al. CVIR 2019 :

International multicentrique

Rétrospectif

Objectif :

CJP → Efficacité (Recist 1.1)

CJS → Toxicité et OS

TNE toute origine et grade avec MH

244 patients sur 8 centres

Réponse complète (RC) : 8 %.

Réponse partielle (RP) : 35 %

Stabilisation de la maladie (SM) : 48 %

Survie globale médiane (SGm) :

G1 : 3,7 ans

G2 : 2,7 ans

G3 : 0,7

Complications spécifiques à la SIRT : < 4 %.

**Symptômes : 44 % amélioration et 34 % résolution.**

Les facteurs pronostiques significatifs de survie :

Grade de la TNE/l'indice Ki67,

Charge tumorale ( $\geq 75\%$ ),

Atteinte extrahépatique

Réponse Recist 1.1.

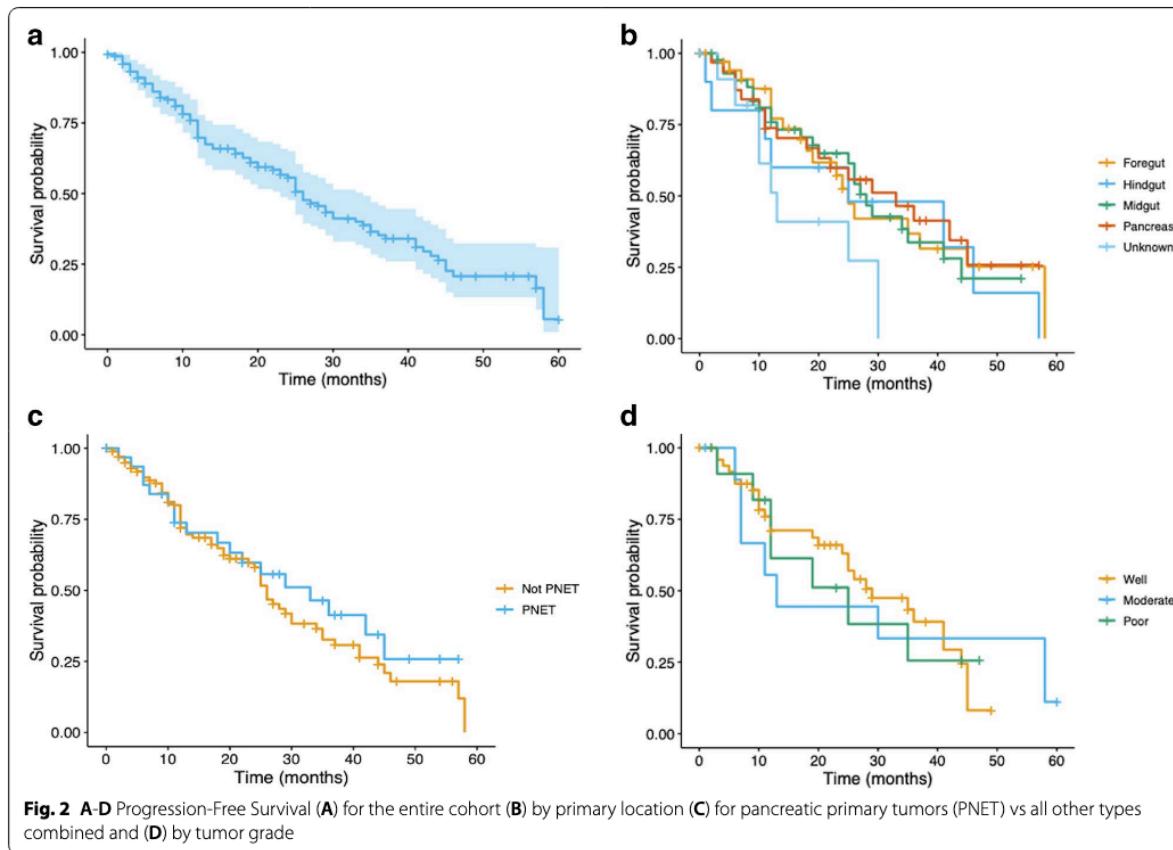
Braat et al. CVIR 2020 : SIRT après RIV 5% de complication

# SIRT et TNE : niveau de preuve – « Resin »

Wong et al. Resin registry Cancer 2022

170 patients suivi >2 ans

Survie globale 33 mois  
Survie sans progression : 25 mois  
Toxicité > grade 3 : 7,6%



Pas de différence significative  
OS ni PFS selon l'origine de la TNE

# SIRT et TNE : niveau de preuve – « Resin »

Schaarschmidt et al. J Nucl Med 2022

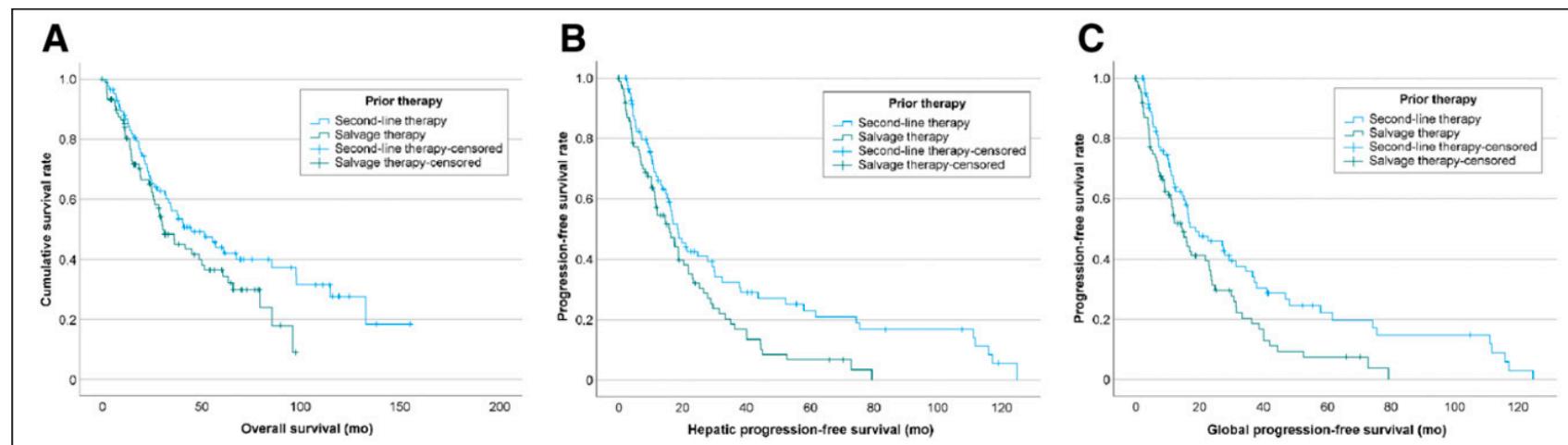
Multicentrique, rétrospective, descriptive

Comparaison originale :

128 patients SIRT de **sauvetage**

102 SIRT en **seconde ligne**

TNE toute origine et tout grade



→ Meilleur en 2<sup>e</sup> ligne

**FIGURE 4.** Kaplan-Meier survival curves investigating influence of extent of prior therapy (second-line therapy: prior surgery for primary tumor or metastases and somatostatin analog treatment before  $^{90}\text{Y}$  RE vs. salvage therapy) on OS (A), hepatic PFS (B), and global PFS (C).

# SIRT et TNE : Dosimétrie

## « Recommandations » dosimétriques

4. Long-term hepatic fibrosis may have a more negative impact than with other tumour types [50]
5. The preference for bland embolization, chemoembolization, or radioembolization remains a subject of scientific debate [10, 50, 58, 59]. Radioembolization typically has better short-term tolerability and durations of response [56]

Euro

### Clinical and dosimetric considerations for yttrium-90 glass

### microspheres radioembolization of intrahepatic cholangiocarcinoma, metastatic colorectal carcinoma, and metastatic neuroendocrine carcinoma: recommendations from an international multidisciplinary working group

Marnix Lam<sup>1,2</sup>  · Riad Salem<sup>3</sup> · Beau Toskich<sup>4</sup> · S. Cheenu Kappadath<sup>5</sup> · Carlo Chiesa<sup>6</sup> · Kirk Fowers<sup>7</sup> · Paul Haste<sup>8</sup> ·

Joseph M. Herman<sup>9</sup> · Edward Kim<sup>10</sup> · Thomas Leung<sup>11</sup> · Siddharth A. Padia<sup>12</sup> · Bruno Sangro<sup>13</sup> · Daniel Y. Sze<sup>14</sup> ·

Etienne Garin<sup>15</sup>

Table 2 mNET radioembolization recommendations

Treatment Intent	<ol style="list-style-type: none"><li>1. In eligible patients, radiation segmentectomy or radiation lobectomy are recommended [10, 57]</li><li>2. Palliation (i.e., symptom and tumour control) and/or consolidation complementary to systemic treatment is the intent for advanced disease</li><li>3. Most patients are treated with palliative intent due to late stage, multifocal disease with large tumour load [50]. Liver function should be preserved to prevent hepatotoxicity that precludes subsequent treatment (locoregional or systemic therapies)</li><li>4. Long-term hepatic fibrosis may have a more negative impact than with other tumour types [50]</li><li>5. The preference for bland embolization, chemoembolization, or radioembolization remains a subject of scientific debate [10, 50, 58, 59]. Radioembolization typically has better short-term tolerability and</li></ol>
Dose Calculation and Dosimetry Considerations	<ol style="list-style-type: none"><li>1. For radiation segmentectomy/lobectomy, it is recommended to use HCC, mCRC, iCCA guidance as a reference due to limited mNET data</li><li>2. MCD is preferred over SCD to evaluate TAD and NTAD [11, 62]. Data on specific dose thresholds is limited to a single institution case series and should be further investigated. Routine clinical use can not be recommended at this time</li><li>3. SCD reference average absorbed dose to the perfused volume is 120 Gy. The perfused volume may be the whole liver or a fraction thereof, which has proven to be safe and effective [50, 52]. TAD and NTAD may vary considerably between patients. Caution in case of poor targeting (<sup>99m</sup>Tc]TcMAA SPECT/CT) and/or low tumour burden (&lt; 10%), which could lead to low efficacy and/or high toxicity, respectively</li><li>4. MCD NTAD prediction is typically more accurate than TAD, especially for (multiple) small and/or infiltrative tumours. In case of multiple smaller tumours, segmentation may be challenging, and instead can use a count-based isocontour thresholding technique on <sup>99m</sup>Tc]TcMAA SPECT/CT</li><li>5. Optimal tumour response and OS are attained when the TAD is ≥ 200 Gy with a minimum TAD ≥ 150 Gy (i.e., survival benefit threshold). Early studies showed a median whole liver NTAD of 54 Gy (range 11–128 Gy) with safe maximum administration ≤ 106 Gy [11]. A maximum NTAD of 75 Gy for single-session whole liver treatment is recommended to ensure safety. Proportionally higher NTAD is acceptable for smaller treatment fractions</li><li>6. Timing of radioembolization and extent of prior liver treatments should be considered. Evidence of long-term radioembolization effects is sparse and current studies lack clinical and dosimetric parameters [50]</li><li>7. In lower grade disease, emphasis is on safety (NTAD); in higher grade disease, emphasis is on efficacy (TAD)</li></ol>
Treatment Delivery	<ol style="list-style-type: none"><li>1. The interval between sequential lobar treatments ranges from 3–6 months. Interval progression in the untreated lobe is uncommon (except for grade 3/NEC); longer intervals may decrease the risk of liver decompensation. In cases of palliation, a shorter interval may be preferred</li><li>2. Bilobar disease can be treated using single-session bilobar or staged sequential lobar treatment. In general, staged sequential lobar treatment is preferred in low-grade NET to avoid potential long-term toxicity. In case of staged sequential lobar treatment, the lobe with more extensive disease should be treated first. For highly aggressive (i.e., grade 3/NEC) bilobar disease in a patient with preserved liver function and with <sup>99m</sup>Tc]TcMAA tumour targeting (i.e., high TAD; low NTAD), single-session bilobar treatment (i.e., two unilobular injections) should be based on MCD</li></ol>
Outcome Assessment and Follow-up	<ol style="list-style-type: none"><li>1. Contrast-enhanced MRI or CT should be performed every 3–6 months following treatment</li><li>2. Late responses are common and may take up to 4–9 months [63]</li><li>3. In the palliative intent setting, caution is warranted with an overly aggressive retreatment approach in patients with SD or PR. Retreatment (radioembolization, embolization, or systemic therapy) should be considered only in the setting of PD</li></ol>
Strength of Recommendation Degree of Consensus	B Strong

# SIRT et TNE : Dosimétrie

## « Recommandations » dosimétriques

1. All grade disease is acceptable, including neuroendocrine carcinoma (NEC)
2. Acceptable intrahepatic (maximum 50–70%; no maximum in case of palliation and preserved liver function) and extrahepatic disease burden (no defined maximum, as long as prognosis is defined by intrahepatic disease). In general, more extrahepatic disease is acceptable (i.e., grade 1–2 NET) com-

Joseph M. Herman<sup>9</sup> · Edward Kim<sup>10</sup> · Thomas Leung<sup>11</sup> · Siddharth A. Padia<sup>12</sup> · Bruno Sangro<sup>13</sup> · Daniel Y. Sze<sup>14</sup> ·

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Table 2 mNET radioembolization recommendations

Treatment Intent	1. In eligible patients, radiation segmentectomy or radiation lobectomy are recommended [10, 57] 2. Palliation (i.e., symptom and tumour control) and/or consolidation complementary to systemic treatment is the intent for advanced disease 3. Most patients are treated with palliative intent due to late stage, multifocal disease with large tumour load [50]. Liver function should be preserved to prevent hepatotoxicity that precludes subsequent treatment (locoregional or systemic therapies) 4. Long-term hepatic fibrosis may have a more negative impact than with other tumour types [50] 5. The preference for bland embolization, chemoembolization, or radioembolization remains a subject of scientific debate [10, 50, 58, 59]. Radioembolization typically has better short-term tolerability and durations of response [56]
Patient Selection	1. All grade disease is acceptable, including neuroendocrine carcinoma (NEC) 2. Acceptable intrahepatic (maximum 50–70%; no maximum in case of palliation and preserved liver function) and extrahepatic disease burden (no defined maximum, as long as prognosis is defined by intrahepatic disease). In general, more extrahepatic disease is acceptable (i.e., grade 1–2 NET) com-
Treatment Delivery	SPECT/CT) and/or low tumour burden (< 10%), which could lead to low efficacy and/or high toxicity, respectively 4. MCD NTAD prediction is typically more accurate than TAD, especially for (multiple) small and/or infiltrative tumours. In case of multiple smaller tumours, segmentation may be challenging, and instead can use a count-based isocountour thresholding technique on <sup>99m</sup> TcTcMAA SPECT/CT 5. Optimal tumour response and OS are attained when the TAD is $\geq$ 200 Gy with a minimum TAD $\geq$ 150 Gy (i.e., survival benefit threshold). Early studies showed a median whole liver NTAD of 54 Gy (range 11–128 Gy) with safe maximum administration $\leq$ 106 Gy [11]. A maximum NTAD of 75 Gy for single-session whole liver treatment is recommended to ensure safety. Proportionally higher NTAD is acceptable for smaller treatment fractions 6. Timing of radioembolization and extent of prior liver treatments should be considered. Evidence of long-term radioembolization effects is sparse and current studies lack clinical and dosimetric parameters [50] 7. In lower grade disease, emphasis is on safety (NTAD); in higher grade disease, emphasis is on efficacy (TAD)
Outcome Assessment and Follow-up	1. The interval between sequential lobar treatments ranges from 3–6 months. Interval progression in the untreated lobe is uncommon (except for grade 3/NEC); longer intervals may decrease the risk of liver decompensation. In cases of palliation, a shorter interval may be preferred 2. Bilobar disease can be treated using single-session bilobar or staged sequential lobar treatment. In general, staged sequential lobar treatment is preferred in low-grade NET to avoid potential long-term toxicity. In case of staged sequential lobar treatment, the lobe with more extensive disease should be treated first. For highly aggressive (i.e., grade 3/NEC) bilobar disease in a patient with preserved liver function and with <sup>99m</sup> TcTcMAA tumour targeting (i.e., high TAD; low NTAD), single-session bilobar treatment (i.e., two unilobular injections) should be based on MCD 1. Contrast-enhanced MRI or CT should be performed every 3–6 months following treatment 2. Late responses are common and may take up to 4–9 months [63] 3. In the palliative intent setting, caution is warranted with an overly aggressive retreatment approach in patients with SD or PR. Retreatment (radioembolization, embolization, or systemic therapy) should be considered only in the setting of PD
Strength of Recommendation	B
Degree of Consensus	Strong

# SIRT et TNE : Dosimétrie

## « Recommandations » dosimétriques

European Journal of Nuclear Medicine and Molecular Imaging

### Clinical and dosimetric considerations for yttrium-90 glass microspheres radioembolization of intrahepatic cholangiocarcinoma, metastatic colorectal carcinoma, and metastatic neuroendocrine carcinoma: recommendations from an international multidisciplinary working group

Marnix Lam<sup>1,2</sup>  · Riad Salem<sup>3</sup> · Beau Toskich<sup>4</sup> · S. Cheenu Kannadath<sup>5</sup> · Carlo Chiesa<sup>6</sup> · Kirk Fowers<sup>7</sup> · Paul Haste<sup>8</sup>.

5. Optimal tumour response and OS are attained when the TAD is  $\geq 200$  Gy with a minimum TAD  $\geq 150$  Gy (i.e., survival benefit threshold). Early studies showed a median whole liver NTAD of 54 Gy (range 11–128 Gy) with safe maximum administration  $\leq 106$  Gy [11]. A maximum NTAD of 75 Gy for single-session whole liver treatment is recommended to ensure safety. Proportionally higher NTAD is acceptable for smaller treatment fractions

Table 2 mNET radioembolization recommendations

Treatment Intent	1. In eligible patients, radiation segmentectomy or radiation lobectomy are recommended [10, 57] 2. Palliation (i.e., symptom and tumour control) and/or consolidation complementary to systemic treatment is the intent for advanced disease 3. Most patients are treated with palliative intent due to late stage, multifocal disease with large tumour load [50]. Liver function should be preserved to prevent hepatotoxicity that precludes subsequent treatment (locoregional or systemic therapies) 4. Long-term hepatic fibrosis may have a more negative impact than with other tumour types [50] 5. The preference for bland embolization, chemoembolization, or radioembolization remains a subject of scientific debate [10, 50, 58, 59]. Radioembolization typically has better short-term tolerability and durations of response [56]
Patient Selection	1. All grade disease is acceptable, including neuroendocrine carcinoma (NEC) 2. Acceptable intrahepatic (maximum 50–70%; no maximum in case of palliation and preserved liver function) and extrahepatic disease burden (no defined maximum, as long as prognosis is defined by intrahepatic disease). In general, more extrahepatic disease is acceptable (i.e., grade 1–2 NET) compared with other grade NET and/or other tumour types 3. Patients may receive treatment before or after PRRT. Radioembolization demonstrates acceptable tolerability post-PRRT and does not limit subsequent treatment [11, 60, 61]
Pretreatment Imaging	1. Multiple phase contrast-enhanced MRI or CT is recommended within 4–8 weeks of treatment. Depending on tumour grade, a longer interval is acceptable 2. Somatostatin receptor imaging may be performed for staging in somatostatin receptor positive grade 1–2 NET ( $^{18}\text{F}$ FDG-PET/CT in grade 3/NEC)
Dose Calculation and Dosimetry Considerations	1. For radiation segmentectomy/lobectomy, it is recommended to use HCC, mCRC, iCCA guidance as a reference due to limited mNET data 2. MCD is preferred over SCD to evaluate TAD and NTAD [11, 62]. Data on specific dose thresholds is limited to a single institution case series and should be further investigated. Routine clinical use can not be recommended at this time 3. SCD reference average absorbed dose to the perfused volume is 120 Gy. The perfused volume may be the whole liver or a fraction thereof, which has proven to be safe and effective [50, 52]. TAD and NTAD may vary considerably between patients. Caution in case of poor targeting ( $^{99m}\text{Tc}$ TcTeMAA)
Outcome Assessment and Follow-up	1. Contrast-enhanced MRI or CT should be performed every 3–6 months following treatment 2. Late responses are common and may take up to 4–9 months [63] 3. In the palliative intent setting, caution is warranted with an overly aggressive retreatment approach in patients with SD or PR. Retreatment (radioembolization, embolization, or systemic therapy) should be considered only in the setting of PD
Strength of Recommendation	B
Degree of Consensus	Strong

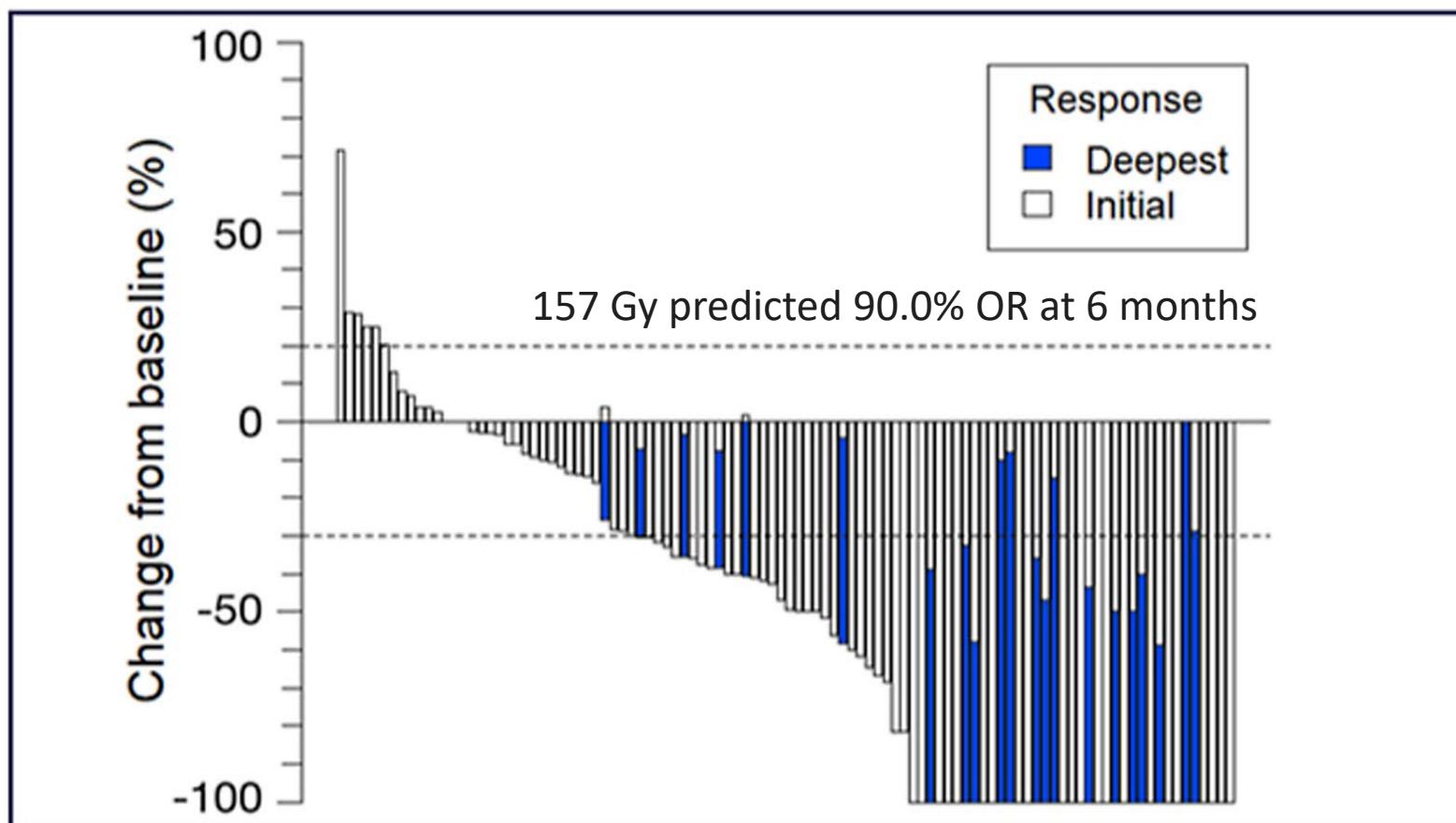
# SIRT et TNE : Dosimétrie

Dosimetrie :

Doyle 2024 ( JVIR)

Tumeurs >120Gy → Réponse objective

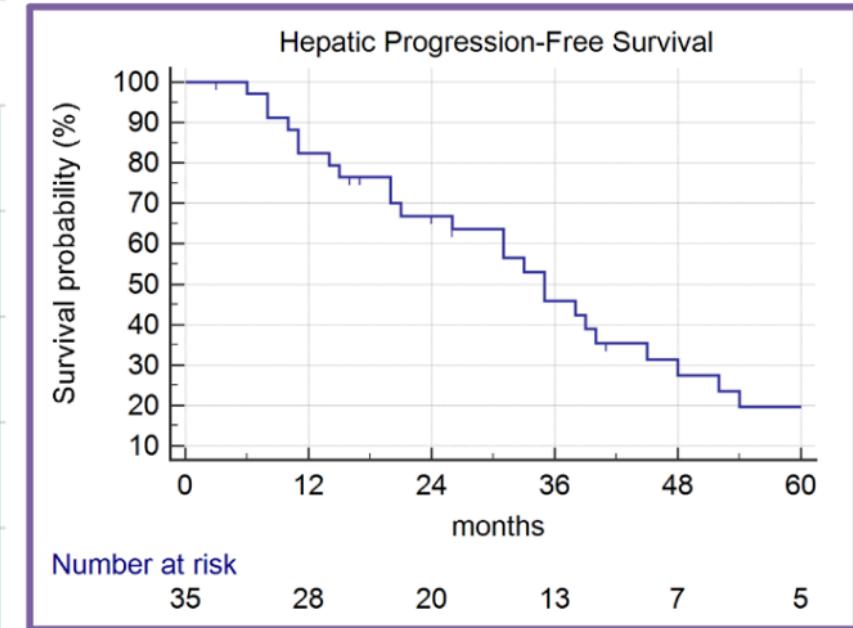
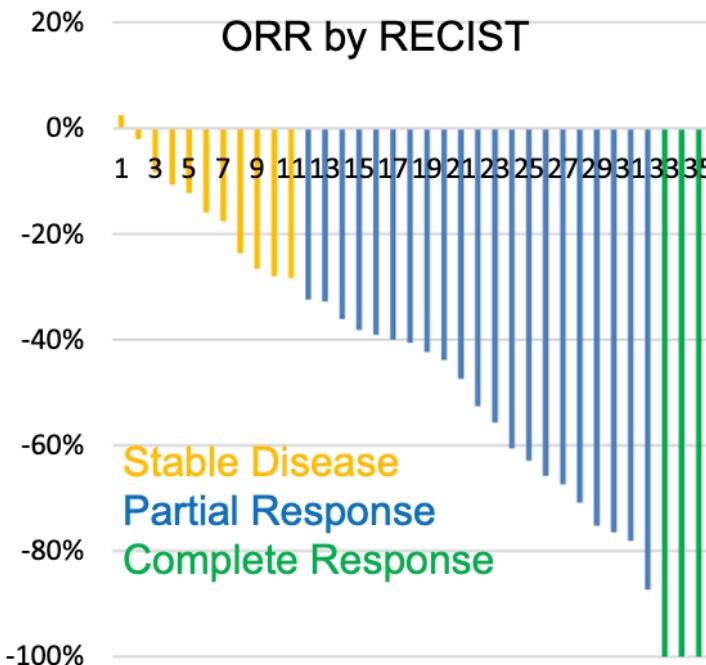
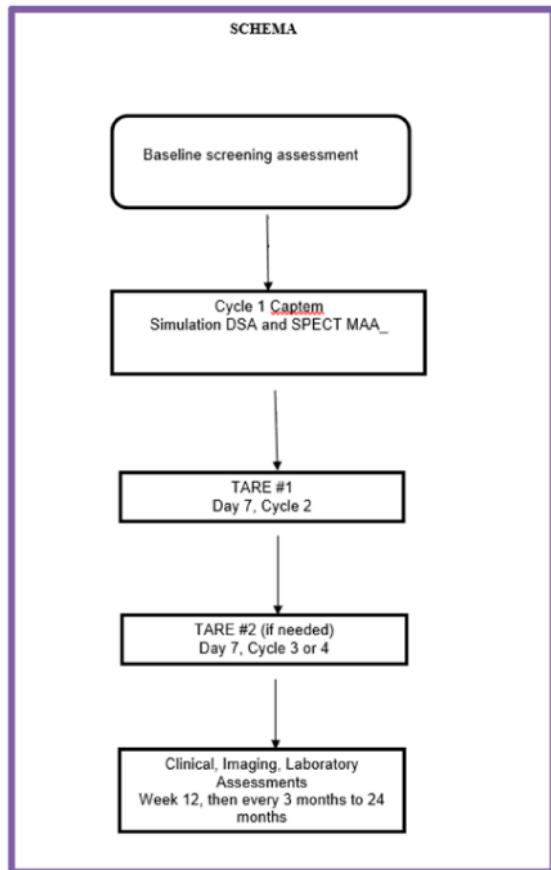
Rétrospective monocentrique – 35 patients 56 tumeurs



Avenir et perspectives?

# SIRT et TNE : combinaison thérapeutique

Soulen et al. CVIR 2023 : SIRT + Capecitabine-Temozolomide pour TNE Grade 2  
Monocentrique – rétrospective, 37 patients

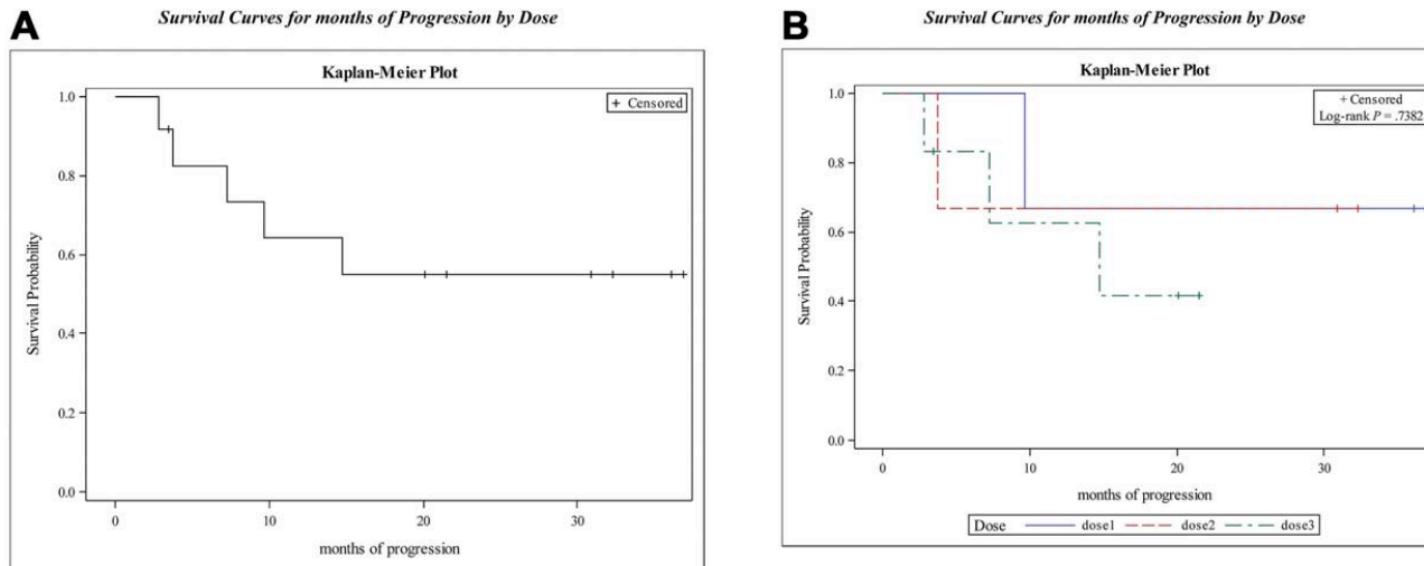


The combination of radiosensitizing chemotherapy with CapTem and  $^{90}\text{Y}$ -TARE provided durable control of G2 NET liver metastases for substantially longer than expectations for embolotherapy or chemotherapy alone.

# SIRT et TNE : combinaison thérapeutique

Kim et al. Cancer 2018

Phase 1b Pasireotide, Everolimus et SIRT – 13 patients



No. of Subjects	Event	Censored	Median Survival (95% CI)		
			12-mo Survival	24-mo Survival	36-mo Survival
12	5 (42%)	7 (58%)	NA (3.8, NA)	64.2% (30.2%, 84.8%)	55.0% (23.2%, 78.3%)

Dose	No. of Subjects	Event	Censored	Median Survival (95% CI)		
				12-mo Survival	24-mo Survival	36-mo Survival
dose1	3	1 (33%)	2 (67%)	NA (9.6, NA)	66.7% (5.4%, 94.5%)	66.7% (5.4%, 94.5%)
dose2	3	1 (33%)	2 (67%)	NA (3.8, NA)	66.7% (5.4%, 94.5%)	NA (NA, NA)
dose3	6	3 (50%)	3 (50%)	14.7 (2.8, NA)	62.5% (14.2%, 89.3%)	NA (NA, NA)

**Figure 2.** (A) Kaplan-Meier curve for progression-free survival showing a median of 18.6 months (95% CI, 7.3 months to not reached). The progression-free survival rates were 61.5% at 1 year, 46.2% at 2 years, 38.5% at 3 years, and 28.8% at 4 years of follow-up. (B) Kaplan-Meier curve for overall survival showing a median of 46.3 months (95% CI, 18 months to not reached). The overall survival rates were 92.3% at 1 year, 69.2% at 2 years, 61.5% at 3 years, and 38.5% at 4 years of follow-up. CI indicates confidence interval; NA, not available.

Dose recommandée 10mg/j

Pas de suite...

# SIRT et TNE : niveau de preuve – A suivre

ArTisAn trial – Sharma et al. BMC 2022  
efficacité de la SIRT dans le traitement  
des (NEL) métastatiques hépatiques  
inopérables.

Phase 2 – Ouvert - Prospectif  
24 patients  
CJP : réponse Recist 1.1

En cours

CapTemY90 for Grade 2/3 NET Liver Metastases  
(CapTemY90)

Phase 2 - Prospectif  
70 patients attendus  
4 centres

Critères de jugement : PFS/OS

# SIRT et TNE : Actualité

été 2025

Sirtex reçoit une approbation de marquage CE élargi pour ses microsphères en résine SIR-Spheres® Y-90

[USA](#) - [Français](#) ▾

À propos de SIR-Spheres® en Europe

Les microsphères de résine SIR-Spheres® Y-90 sont indiquées pour le traitement :

- du carcinome hépatocellulaire (CHC) non résécable, ou
- des tumeurs hépatiques métastatiques non résécables d'un cancer colorectal primaire...
- du cholangiocarcinome intrahépatique non résécable, ou
- **des métastases hépatiques de tumeurs neuroendocrines (mNET), ou**
- d'autres métastases hépatiques

# SIRT et TNE : Conclusion

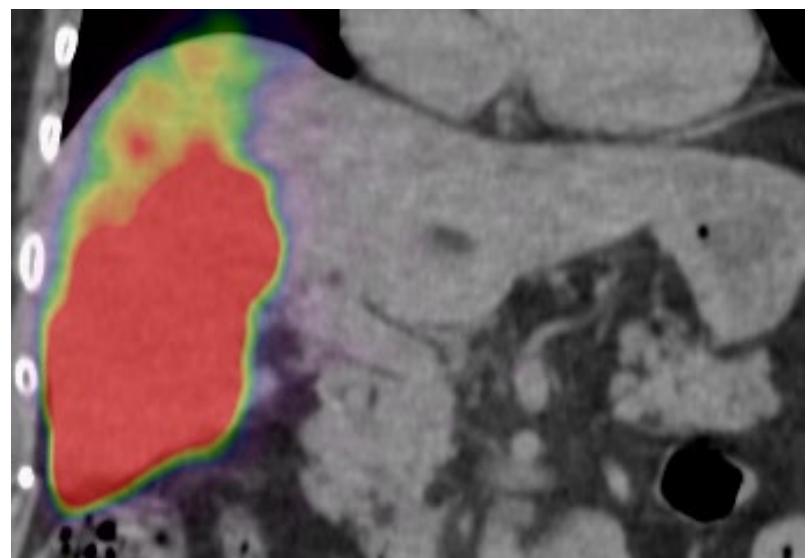
Outil de l'arsenal thérapeutique

Spécificité du type tumoral – faible niveau de preuve

→ Dosimétrie

→ Association thérapeutique

→ Tôt dans l'évolution



## **PATIENT SELECTION: NORTHWESTERN**

- DISEASE BURDEN
  - Large, bulky tumors
  - Bilobar multi-focal disease
  - Infiltrative disease
  - Hypovascular Tumors
- CLINICAL INDICATORS
  - Significant carcinoid symptoms
  - Compromised performance status
- SPECIAL CONSIDERATIONS
  - Failed other embolic therapy
  - Biliary tree compromised

*NORTHWESTERN  
UNIVERSITY  
RADIOLoGY*



# Métastases hépatiques des tumeurs neuroendocrines :

*SIRT et ses perspectives*

Maxime Barat

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Université  
de Paris



Université  
Paris Cité