



Ganglion sentinelle et mélanome

1^{ère} JFMN, La Rochelle, 2015

Dr Jean-Cyril Bourre,

**Pierre-Yves Brard, Julie Charles, Mattia Stella, Catherine Rouet,
Raphaëlle Andreani, Anne Francois-Joubert,**

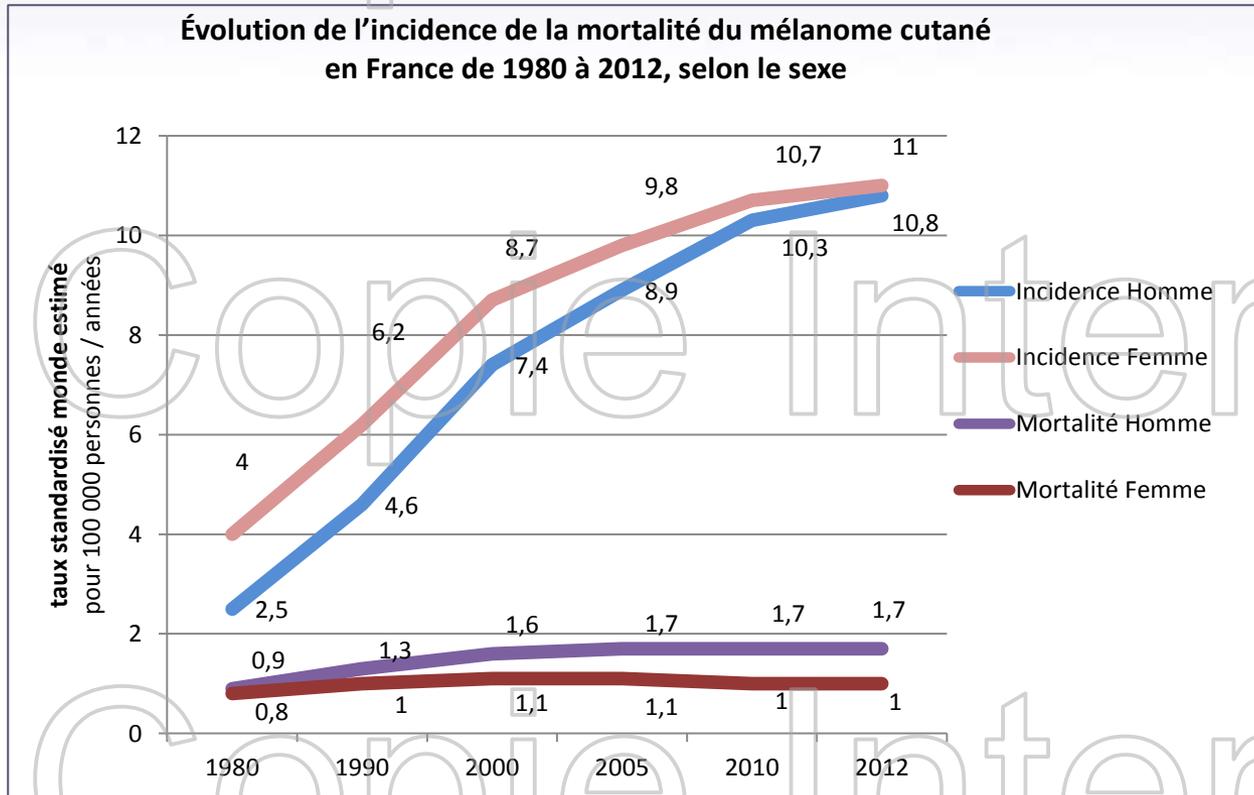
CH Métropole-Savoie, Chambéry

INSERM U1039, UJF, Grenoble

Ganglion sentinelle et mélanome

- **Introduction**
- Indications
- Technique
- Résultats
- Perspectives

Ganglion sentinelle et mélanome



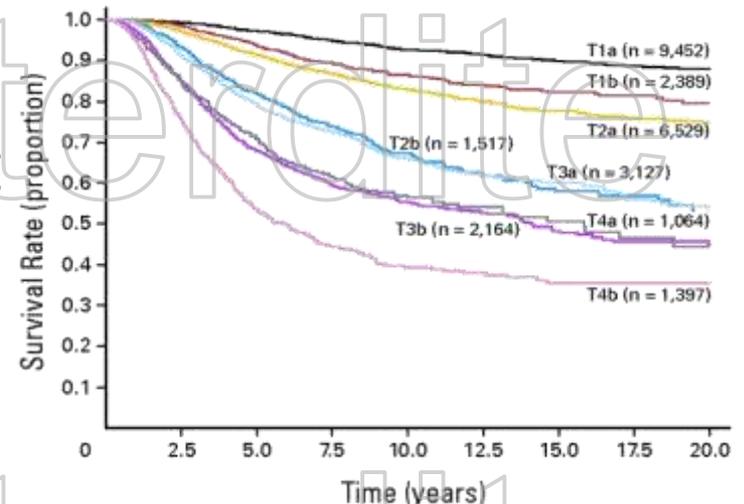
[1] Les cancers en France. Les données. INCa ed.: 2015.

Ganglion sentinelle et mélanome

| Classification | | |
|----------------|--|--|
| T | Epaisseur (mm) | Ulcération / index mitotique |
| Tis | Non applicable | |
| T1 | ≤ 1,00 | a : pas d'ulcération et index mitotique <1/mm ² b : présence d'ulcération ou index mitotique >1/mm ² |
| T2 | 1,01-2,00 | a : pas d'ulcération b : présence d'ulcération |
| T3 | 2,01-4,00 | a : pas d'ulcération b : présence d'ulcération |
| T4 | >4,00 | a : pas d'ulcération b : présence d'ulcération |
| N | Nombre de ganglions envahis | Charge tumorale |
| N0 | 0 | |
| N1 | 1 | a : découvert sur le ganglion sentinelle b : détectable cliniquement |
| N2 | 2-3 | a : découvert sur le ganglion sentinelle b : détectable cliniquement c : métastase satellite ou en transit, sans envahissement ganglionnaire |
| N3 | >3 ou métastase satellite ou en transit, associée à un envahissement ganglionnaire | |
| M | Site | Taux de LDH |
| M0 | Absence de métastase à distance | Non applicable |
| M1a | Métastase cutanée ou sous cutanée à distance, envahissement ganglionnaire à distance | Normal |
| M1b | Métastase pulmonaire | Normal |
| M1c | Toute autre métastase viscérale | Normal |
| | Toute métastase | Élevé |

Stade

I
II
III
IV



VOLUME 27 NUMBER 20 DECEMBER 20 2009
JOURNAL OF CLINICAL ONCOLOGY ORIGINAL REPORT

[2]

Final Version of 2009 AJCC Melanoma Staging and Classification

Charles M. Balch, Jeffrey E. Gershenwald, Sergio Hoon, John F. Thompson, Michael B. Atkins, Darrel R. Byrd, Antonio C. Burdick, Alan J. Cochran, Daniel G. Cote, Shihuan Ding, Alexander M. Eggner, Keith T. Flaherty, Phyllis A. Gimotty, John M. Kirkwood, Kelly M. McMasters, Martin C. Mihov Jr, Donald L. Morton, Merrick I. Ross, Arthur J. Sober, and Vernon K. Sondak

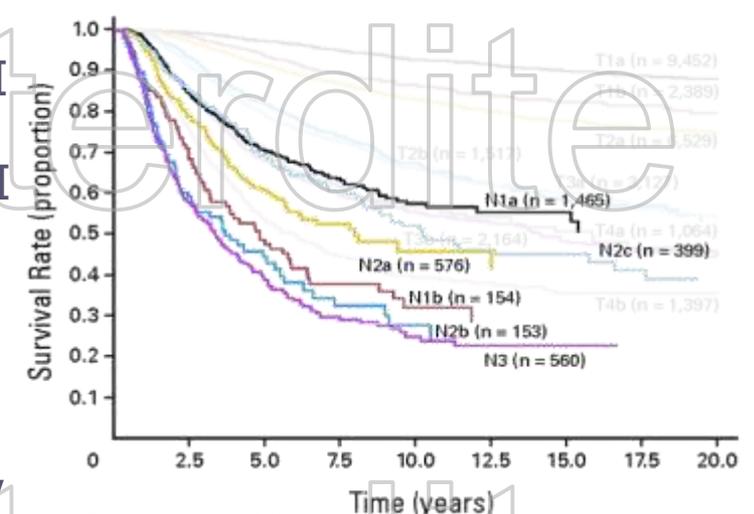
Ganglion sentinelle et mélanome

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Stade



Ganglionnaire



Final Version of 2009 AJCC Melanoma Staging and Classification
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Ganglion sentinelle et mélanome

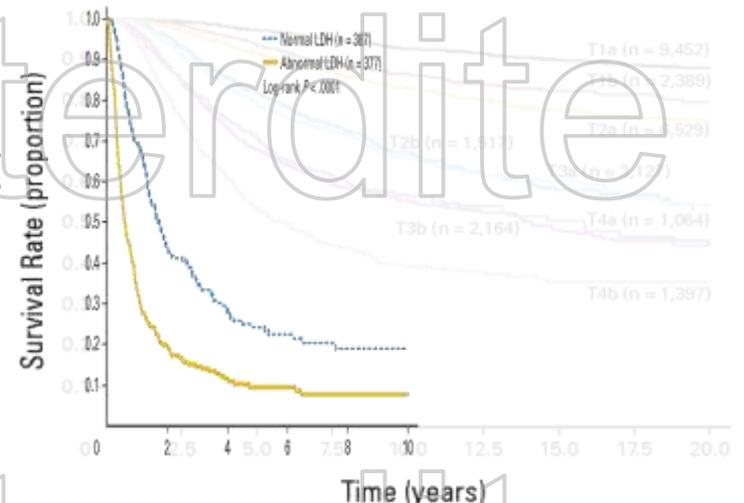
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| T3 | 2,01-4,00 | a : pas d'ulcération b : présence d'ulcération |
| T4 | >4,00 | a : pas d'ulcération b : présence d'ulcération |
| N | Nombre de ganglions envahis | Charge tumorale |
| N0 | 0 | |
| N1 | 1 | a : découvert sur le ganglion sentinelle b : détectable cliniquement |
| N2 | 2-3 | a : découvert sur le ganglion sentinelle b : détectable cliniquement |
| N3 | >3 ou métastase satellite ou en transit, associée à un envahissement ganglionnaire | c : métastase satellite ou en transit, sans envahissement ganglionnaire |
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Stade

I
II
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IV

Extra-Ganglionnaire



VOLUME 27 NUMBER 20 DECEMBER 20 2009
JOURNAL OF CLINICAL ONCOLOGY ORIGINAL REPORT

[2]

Final Version of 2009 AJCC Melanoma Staging and Classification

Charles M. Balch, Jeffrey E. Gershenwald, Sergio A. Soong, John F. Thompson, Michael B. Atkins, Darrel R. Byrd, Antonio C. Gogas, Alan J. Cochran, Daniel G. Coit, Shitanshu Ding, Alexander M. Eggemann, Keith T. Flaherty, Phyllis A. Gimotty, John M. Kirkwood, Kelly M. McMasters, Martin C. Mihov Jr, Donald L. Morton, Merrick I. Ross, Arthur J. Sober, and Vernon K. Sondak

Ganglion sentinelle et mélanome

- Introduction
- **Indications**
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Ganglion sentinelle et mélanome

- SOR 2005

Les Standards, Options et Recommandations

Exérèse du ganglion sentinelle chez les patients atteints d'un mélanome cutané sans envahissement ganglionnaire cliniquement décelable

Standards

L'exérèse du ganglion sentinelle n'est pas recommandée de façon systématique (accord d'experts).

Option

L'exérèse du ganglion sentinelle, réalisée par une équipe entraînée, peut être proposée dans le cadre d'essais thérapeutiques ou de protocoles d'évaluation pour les mélanomes supérieurs à 1 mm d'épaisseur ou ulcérés (accord d'experts).

Recommandations 2005

Prise en charge des patients adultes atteints d'un mélanome cutané MO

Recommandations pour la Pratique Clinique :
Standards, Options et Recommandations 2005
pour la prise en charge des patients adultes
atteints d'un mélanome cutané MO [3]

Texte court

S. Négrier, P. Saiag, B. Guilloeuf, O. Verola, M.-F. Avril, C. Bailly, D. Cupissol, S. Dulac, A. Dunino, B. Dreno, J.-J. Grob, M.-T. Leccia, C. Renaud-Vilmer, L. Boscquet

- EANM 2009
- ASCO 2012

Ganglion sentinelle et mélanome

- SOR 2005
- EANM 2009

In light of the above-mentioned evidence, it is recommended that patients diagnosed with melanoma in clinical stage *T1b-T4b, N0 and M0* should be given an option of SNB.

- ASCO 2012

Int J Nucl Med Mol Imaging
DOI 10.1007/s00295-009-1224-4

GUIDELINES [4]

EANM-EORTC general recommendations for sentinel node diagnostics in melanoma

Annette H. Chakera • Birger Hesse • Zeynep Borak • James R. Ballinger • Allan Britten • Corrado Caracci • Alistair J. Cochran • Martin G. Cook • Krzysztof T. Drzewinski • Richard Essner • Einat Even-Sapir • Alexander M. M. Eggermont • Tanja Ganciner Stopar • Christian Ingvar • Martin C. Mihm Jr. • Stanley W. McCarthy • Nicola Mozillo • Otago E. Nieweg • Richard A. Scolyer • Hans Starz • John F. Thompson • Giuseppe Trifiro • Giuseppe Viale • Sergi Vidal-Sicart • Roger Uren • Wendy Waddington • Arturo Chini • Alan Spatz • Aleksandra Todorci

VOLUME 30 • NUMBER 22 • AUGUST 10, 2012

JOURNAL OF CLINICAL ONCOLOGY

ASCO SPECIAL ARTICLE

Sentinel Lymph Node Biopsy for Melanoma: American Society of Clinical Oncology and Society of Surgical Oncology Joint Clinical Practice Guideline [5]

Patricia Harley, Sejin E. Agarwala, Timothy J. Alberti, Alberto Cochran, Dandoo Y. Kim, Kelly M. McManus, R. David Noyes, Lynn M. Schwab, et al., and Gary H. Lyman

Key Recommendations

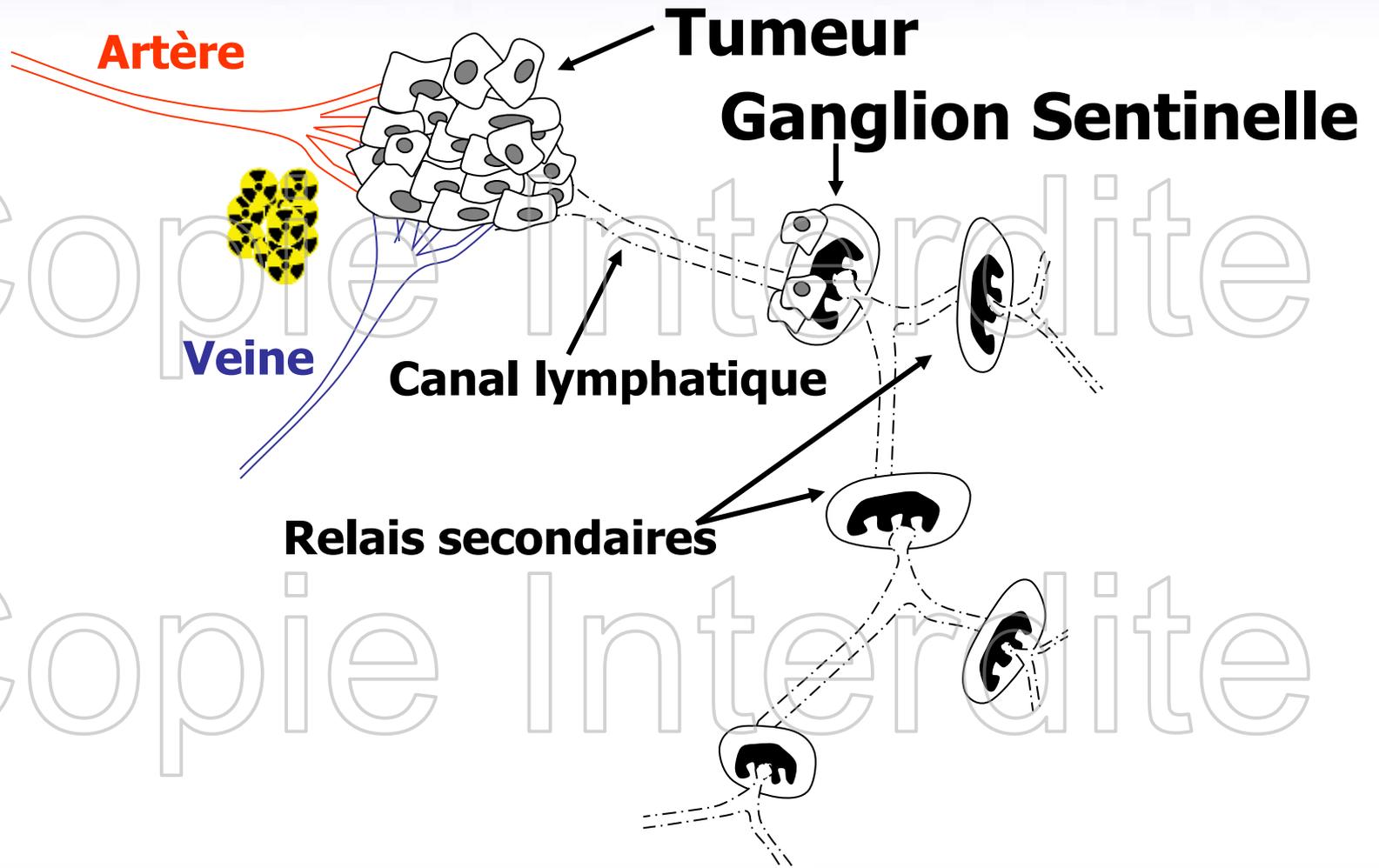
- Intermediate-thickness melanomas: SLN biopsy is recommended for patients with cutaneous melanomas with Breslow thickness of 1 to 4 mm at any anatomic site
- Thick melanomas: SLN biopsy may be recommended for staging purposes and to facilitate regional disease control for patients with melanomas that are T4 or > 4 mm in Breslow thickness
- Thin melanomas: There is insufficient evidence to support routine SLN biopsy for patients with melanomas that are T1 or < 1 mm in Breslow thickness, although it may be considered in selected high-risk patients
- Completion lymph node dissection is recommended for all patients with a positive SLN biopsy

Ulcération ou index mitotique élevé

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Concept



Technique

Exérèse sans marge réalisée « en ville »



Mélanome confirmé par l'anatomie pathologique
+ épaisseur selon Breslow



Reprise chirurgicale avec marge

➔ technique du GS (la veille ou le matin)

| Recommandations 2005 | |
|---|-------------------------------|
| Prise en charge des patients adultes atteints d'un mélanome cutané MO | |
| Epaisseur de Breslow (classification) | Marges d'exérèse recommandées |
| Mélanome <i>in situ</i> (pTis) | => 0,5 cm |
| 0-1 mm (Pt1) | => 1 cm |
| 1,01-2 mm (Pt2) | => 1-2 cm |
| 2,01-4 mm (Pt3) | => 2 cm |
| > 4 mm (Pt4) | => 2-3 cm |

Eur J Nucl Med Mol Imaging (2009) 36:928-937
DOI 10.1007/s00259-008-1036-2

ORIGINAL ARTICLE

[6]

One-day or two-day procedure for sentinel node biopsy in melanoma?

A. H. Chakera · J. Lock-Andersen · U. Hesse ·
B. M. Nürnberg · B. R. Juhl · K. H. Stokholm ·
K. T. Drzewiecki · B. Hesse

Technique

- Injections intradermiques
- En périphérie de la cicatrice d'exérèse
- Images dynamiques précoces (10 min)

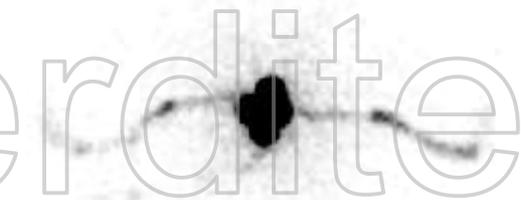
Tableau 3
Images à réaliser obligatoirement (*) ou en option (°) lors de la recherche du GS, en fonction de la topographie du mélanome [4].

Summary of early dynamic imaging or delayed static imaging must () or optionally could (°) be performed according to melanoma's localization.*

| Localisation | Images dynamiques | Images statiques |
|--------------------|-----------------------------------|--|
| Tronc | Centrées sur le site d'injection° | Balayage du cou à l'aîne* Ou* creux axillaire (antérieur) Aîne (antérieur) Tronc (postérieur) |
| Main et avant-bras | Centrées sur le site d'injection° | Creux axillaire (antérieur)* Cou (antérieur)* |
| Bras | Centrées sur le site d'injection° | Creux axillaire (antérieur)* Cou (antérieur)* Creux axillaire (profil)* |
| Membre inférieur | Centrées sur le site d'injection° | Aîne (antérieur)* Creux poplité (postérieur)* |
| Tête et cou | Centrées sur le site d'injection* | Tête et cou (antérieur)* Tête et cou (profil)* |



Face antérieure

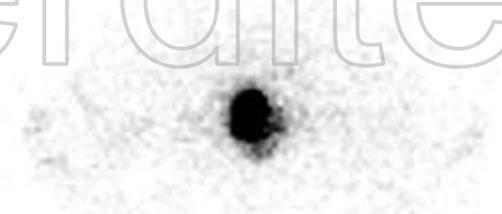


Face postérieure

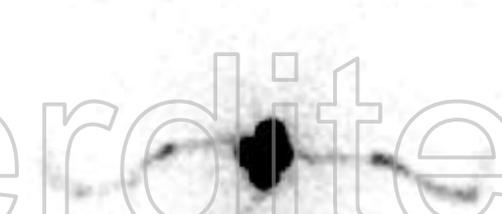
[7] Bourre JC, Brard PY, Charles J, Stella M, Rouet C, Andreani R, Francois-Joubert A. Ganglion sentinelle et mélanome Med Nucl 2013; 39:339-347

Technique

- Images statiques précoces (5 min/image)
 - Dès que le (les) GS est (sont) identifié(s)
 - +/- clichés tardifs

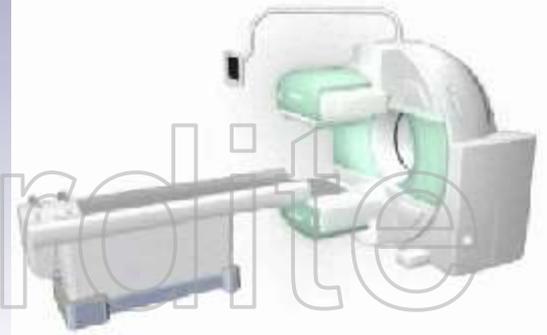


Face antérieure

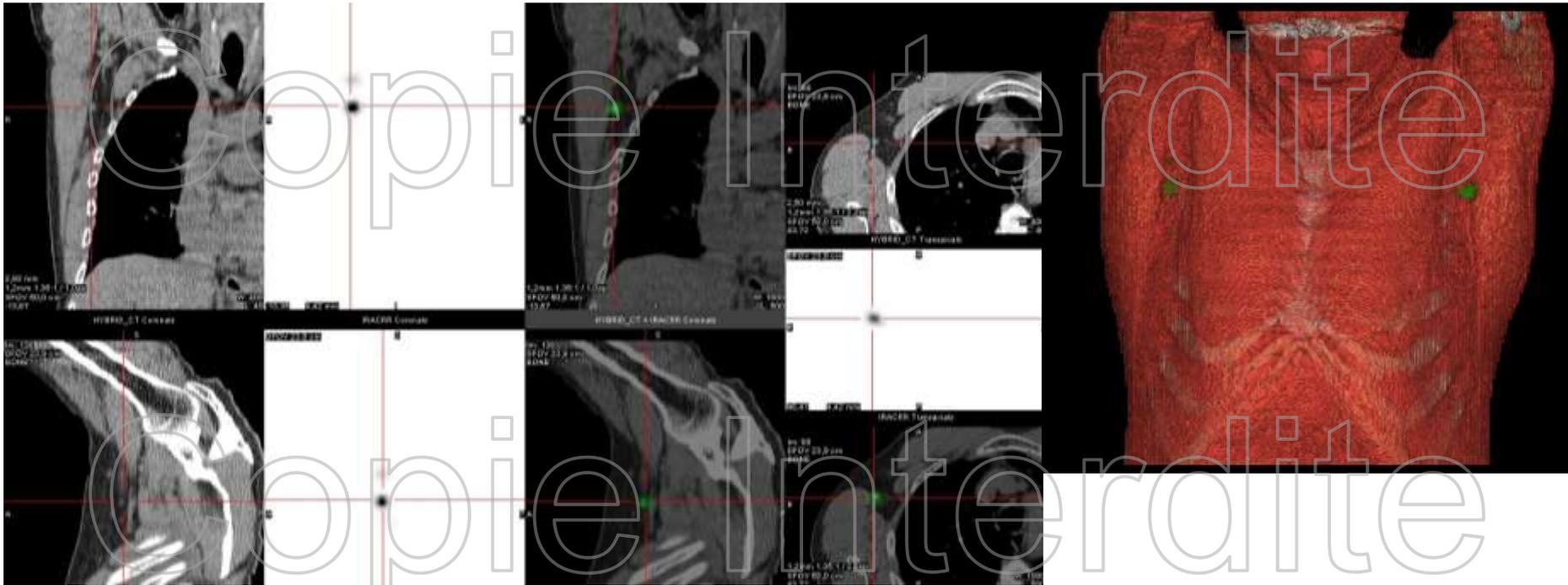


Face postérieure

Technique : TEMP-TDM



- Tomoscintigraphie couplée au scanner X (TEMP-TDM)



Technique : TEMP-TDM

- Intérêt de la TEMP-TDM
 - Augmentation de la sensibilité de détection
 - Augmentation de la spécificité
 - Modifie le geste chirurgical
 - Coût efficace
 - Améliore la survie sans récurrence ?

TEMP-TDM

Augmente la sensibilité

- N=85
 - Membre 41
 - Tronc 31
 - Tête et cou 14
- +8% de GS identifiés (226 vs 214)
 - Correction de l'atténuation
 - Meilleur rapport signal sur bruit
 - Proximité des sites d'injection
- 14% GS **envahis** n'étaient vus qu'en TEMP-TDM
- + séméiologie TDM en cas de GNS suspect

Ann Surg Oncol (2009) 16:1537–1542
DOI 10.1245/s10434-009-0339-2

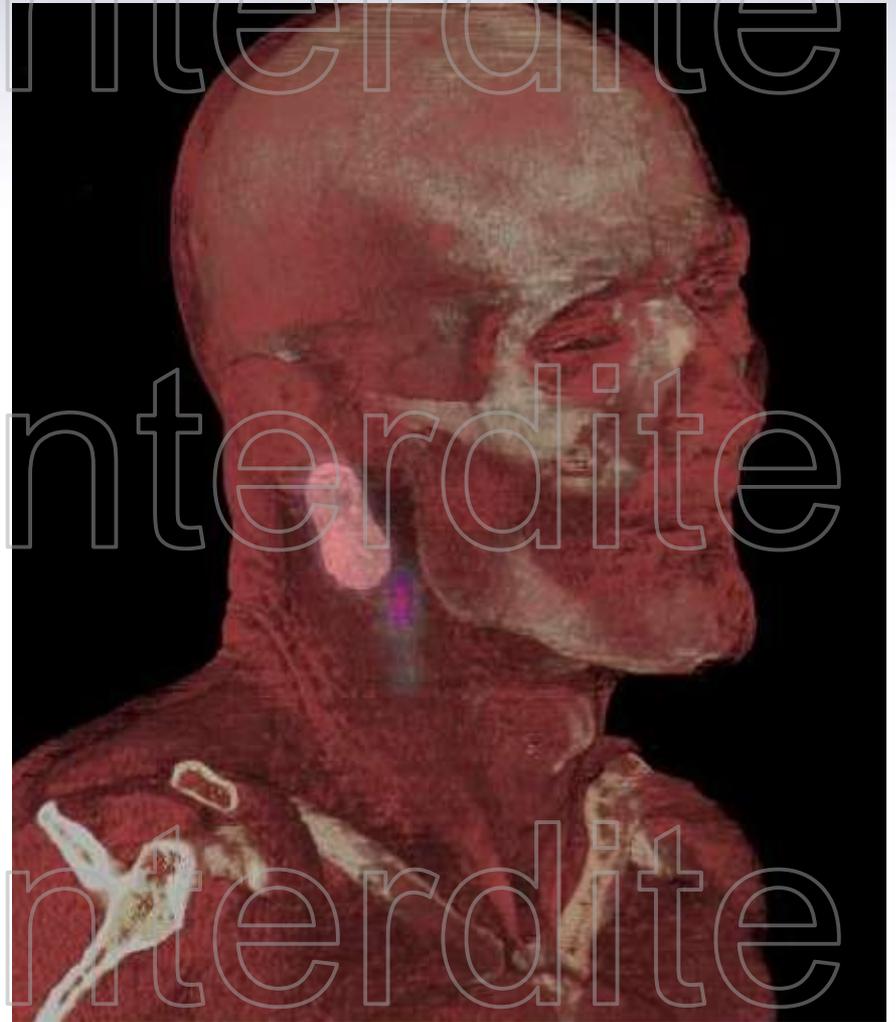
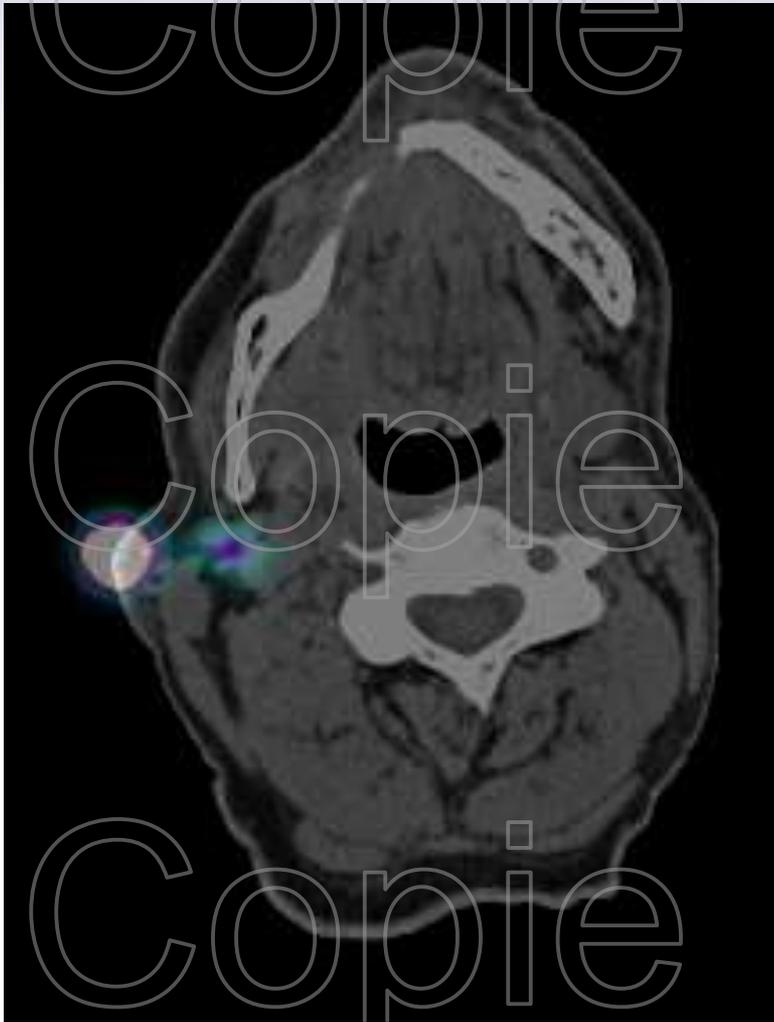
Annals of
SURGICAL ONCOLOGY
OFFICIAL JOURNAL OF THE SOCIETY OF SURGICAL ONCOLOGY

ORIGINAL ARTICLE – MELANOMAS

[8]

The Yield of SPECT/CT for Anatomical Lymphatic Mapping in Patients with Melanoma

Iris M. C. van der Ploeg, MD¹, Renato A. Valdés Olmos, MD, PhD², Bin B. R. Kroon, MD, PhD, FRCS¹, Michael W. J. M. Wouters, MD¹, Michiel W. M. van den Brekel, MD, PhD³, Wouter V. Vogel, MD, PhD², Cornelis A. Hoefnagel, MD, PhD², and Omgo E. Nieweg, MD, PhD¹

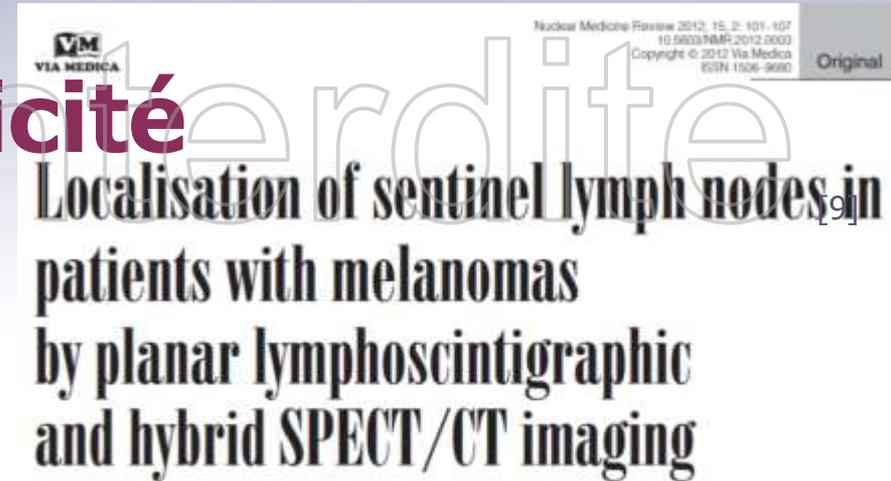


Images F. Gimmarile

TEMP-TDM

Augmente la spécificité

- N=113
 - Membre 54
 - Tronc 55
 - Tête et cou 4
- Spécificité
 - Précision anatomique
 - Faux positifs
 - Contamination
 - Trajet tortueux



Otakar Kozel¹, Martin Havel²
¹Clinic of Nuclear Medicine, University Hospital of Olomouc, Olomouc, Czech Republic;
²Department of Imaging Methods, Faculty of Medicine, University of Olomouc, Olomouc, Czech Republic

ients. Planar images identified 253 SLNs in 100 (88.5%) pts, with a mean of 2.2 ± 1.7 (range 0-9 nodes) per patient. In the remaining 13 (11.5%) patients no SLNs were detected on planar images. On SPECT/CT images, 334 hot nodes were detected in 107 (94.7%) patients with a mean of 3.0 ± 2.1 (range 0-9)

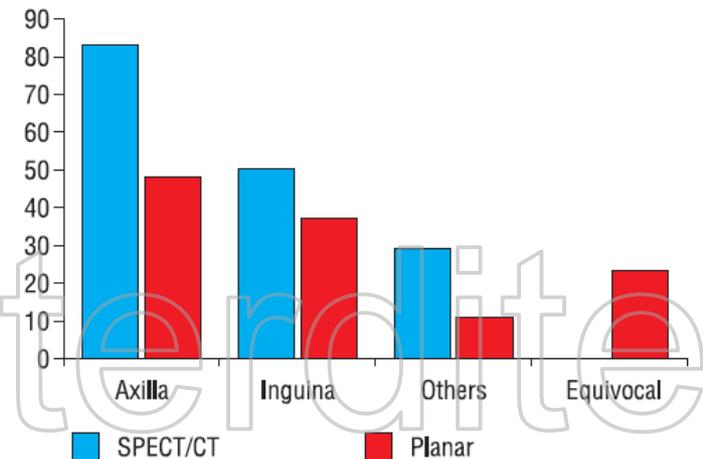


Figure 2. Numbers of found lymph nodes on SPECT/CT and planar imaging in GROUP B

TEMP-TDM

Modifie le geste chirurgical

- 35% chirurgie modifiée
 - 11 incisions plus larges
 - 8 incisions adaptées
 - 5 incisions supplémentaires

Ann Surg Oncol (2009) 16:1537–1542
DOI 10.1245/s10434-009-0339-2

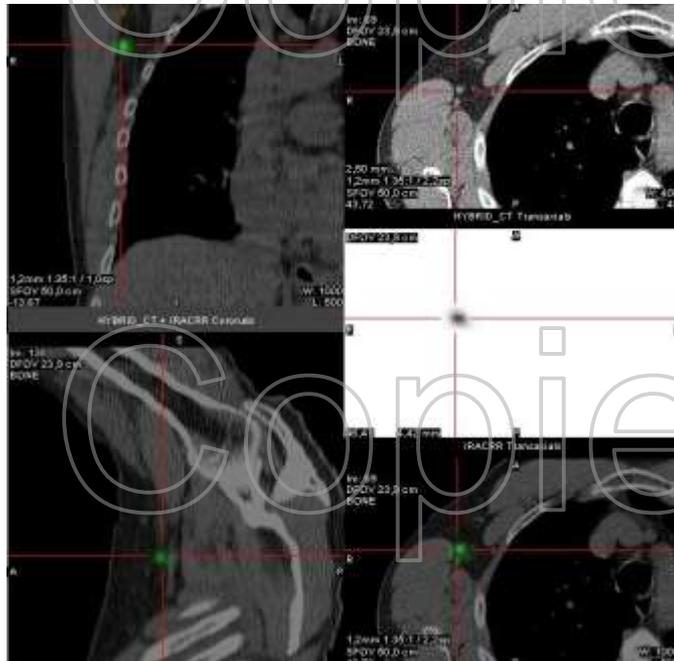
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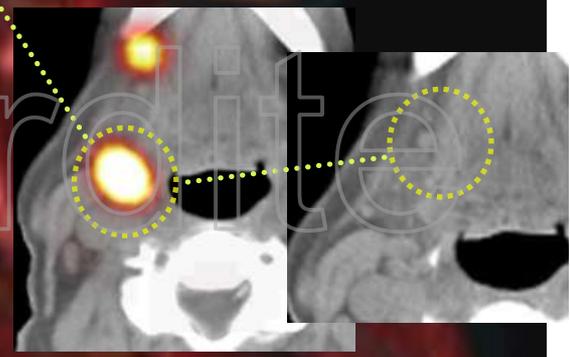
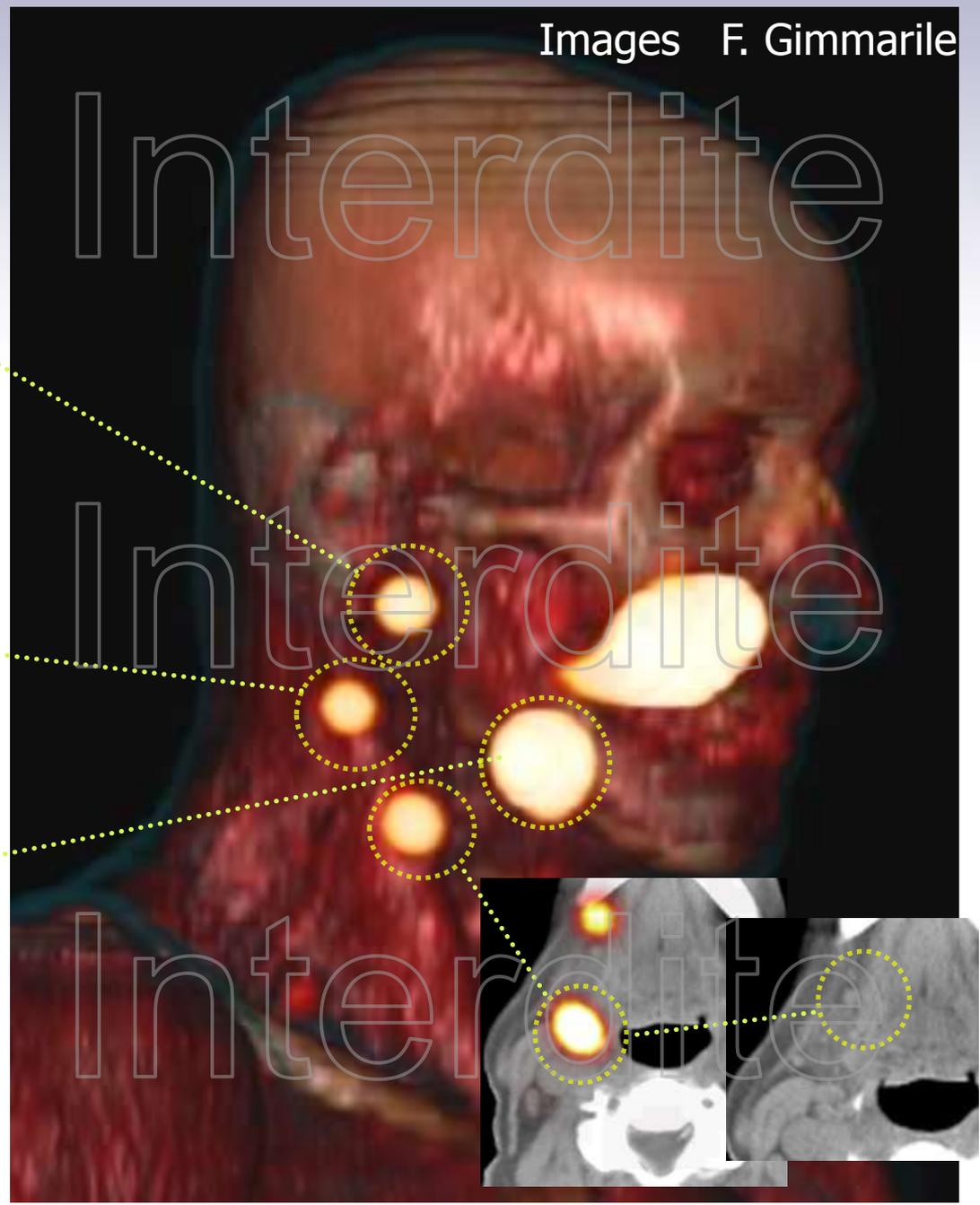
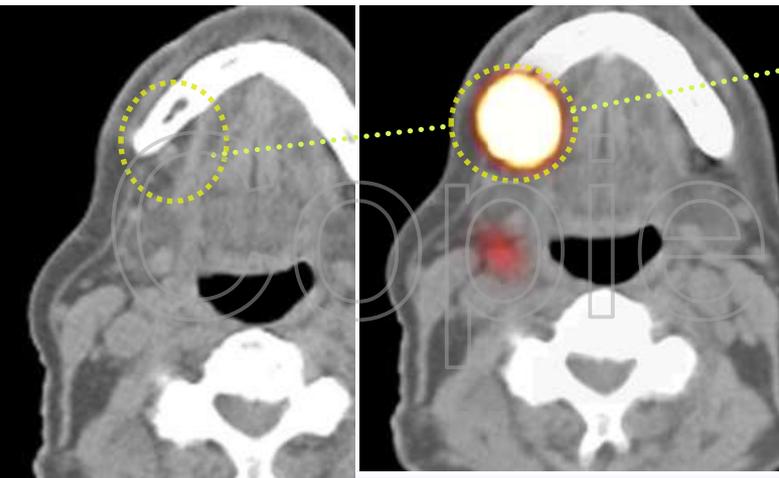
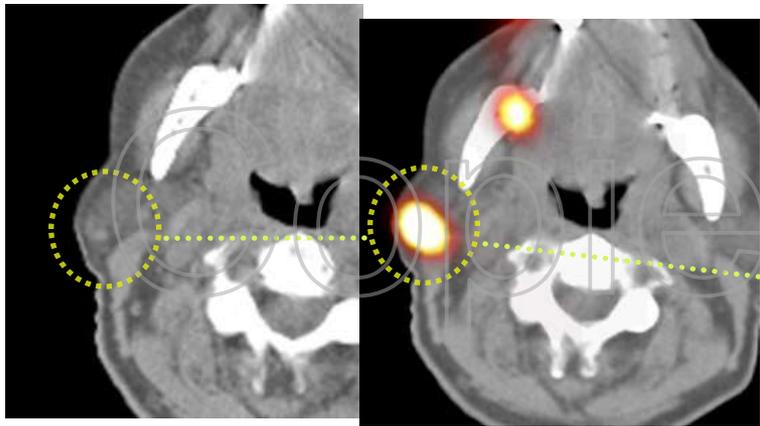
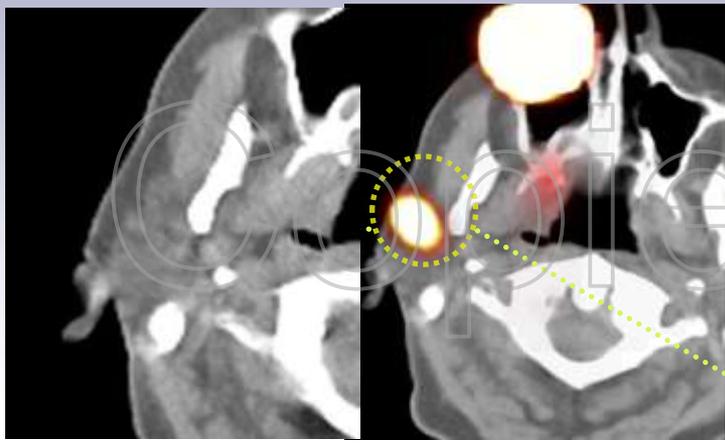
ORIGINAL ARTICLE – MELANOMAS

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| Location | No. | Benefit from SPECT/CT | No. |
|---|-----|---|-----|
| Head and neck | 14 | Incision at another site | |
| | | Submandibular instead of on the cheek | 2 |
| | | Retroauricular instead of at the preauricular region | 3 |
| | | Longer incision | |
| | | Deep in the neck instead of superficial | 1 |
| | | Deep from parotid gland instead of in the gland | 2 |
| | | Smaller incision | |
| | | Superficial in the neck instead of deep | 2 |
| | | Superficial from parotid gland instead of in the gland | 4 |
| | | Trunk | 5 |
| additionally visualized sentinel nodes in the flank | 2 | | |
| additionally visualized sentinel nodes in the scapular region | 2 | | |
| Arm | 3 | additionally visualized sentinel nodes in the supraclavicular fossa | 1 |
| | | Incision at another site | |
| Leg | 8 | Node was in triceps muscle instead of biceps muscle or in between Both muscles instead of in epitrochlear fossa | 3 |
| | | Longer incision | |
| | | Nodes in iliac-obturator zone instead of superficial groin | 8 |





TEMP-TDM

Coût - efficace

- Avant/après
- Allemagne
- 403 patients
 - 254 → planaires
 - 149 → TEMP-TDM
- Economie 30,5% (-710€/pt)
 - Réduction du temps opératoire
 - Réduction des durées d'hospitalisation

Cost-effectiveness of preoperative SPECT/CT combined with lymphoscintigraphy vs. lymphoscintigraphy for sentinel lymph node excision in patients with cutaneous malignant melanoma [10]

Ingo Stoffels • Markus Müller • Marie Henrike Geisel • Julia Leyh • Thorsten Pöppel • Dirk Schadendorf • Joachim Klode

TEMP-TDM

Améliore la survie sans récurrence ?

- Même série que [10]

Association Between Sentinel Lymph Node Excision With or Without Preoperative SPECT/CT and Metastatic Node Detection and Disease-Free Survival in Melanoma [11]

Ingo Stoffels, MD
 Christian Boy, MD
 Thorsten Pöppel, MD
 Jasna Kuhn, MD
 Kerstin Klötgen, MD
 Joachim Dissemund, MD
 Dick Schadendorf, MD
 Joachim Klode, MD

Context Malignant melanoma has become an increasing interdisciplinary public health challenge worldwide. Sentinel lymph node excision (SLNE) is considered the most sensitive and specific staging test for the detection of micrometastatic melanoma in regional lymph nodes.

Objective To compare metastatic node detection and disease-free survival using single-photon emission computed tomography/computed tomography (SPECT/CT)-aided SLNE vs standard SLNE in patients with melanoma.

Design, Setting, and Patients A prospective, computerized melanoma patient database at the University Hospital Essen, Skin Cancer Center, Essen, Germany, was used to identify a cohort of 464 patients eligible for SLNE between March

Table 1. Patient Characteristics of the SPECT/CT Cohort vs Standard Cohort of Lymphoscintigraphy

| Characteristics | Total (N = 403) | Standard Cohort (n = 254) | SPECT/CT Cohort (n = 149) | P Value |
|--------------------------------------|-----------------|---------------------------|---------------------------|---------|
| Age, y | | | | |
| Mean (SD) | 58.62 (15.95) | 58.11 (15.52) | 59.48 (16.70) | .25 |
| Median | 62.00 | 62.00 | 62.00 | |
| Sex, No. (%) | | | | |
| Male | 244 (60.5) | 151 (59.4) | 93 (62.4) | .60 |
| Female | 159 (39.5) | 103 (40.6) | 56 (37.6) | .24 |
| Tumor depth, mm | | | | |
| Mean (SD) | 2.69 (2.18) | 2.71 (2.08) | 2.66 (2.38) | .16 |
| Median | 1.90 | 1.93 | 1.80 | |
| Primary localization, No. (%) | | | | |
| Head or neck | 38 (9.4) | 6 (2.4) | 32 (21.5) | <.001 |
| Torso | 175 (43.4) | 110 (43.3) | 65 (43.6) | >.99 |
| Extremity | 169 (41.9) | 123 (48.4) | 46 (30.9) | <.001 |
| Hand or foot | 21 (5.2) | 15 (5.9) | 6 (4.0) | .49 |
| Ulceration of primary tumor, No. (%) | 114 (28.3) | 83 (32.7) | 30 (20.0) | .005 |
| Localization of SLNs, No. (%) | | | | |
| Head or neck | 41 (10.2) | 6 (2.0) | 35 (23.5) | <.001 |
| Axilla | 191 (47.4) | 123 (48.4) | 68 (45.6) | .61 |
| Groin | 166 (41.2) | 122 (48.0) | 44 (29.5) | <.001 |
| Pectoral | 2 (0.5) | 1 (0.4) | 1 (0.7) | >.99 |
| Popliteal | 3 (0.7) | 2 (0.8) | 1 (0.7) | >.99 |
| SLNs | | | | |
| SLNs per patient | 2.07 (833/403) | 1.87 (475/254) | 2.40 (358/149) | <.001 |
| Median (range) | 2.00 (0-9) | 2.00 (0-9) | 2.00 (0-9) | |
| Positive SLNs | | | | |
| Patients, No. (%) | 99 (24.57) | 48 (18.9) | 41 (27.5) | <.001 |
| SLNs per patient | 0.26 (105/403) | 0.21 (54/254) | 0.34 (51/149) | .04 |
| Median (range) | 0.00 (0-3) | 0.00 (0-2) | 0.00 (0-3) | |
| Obesity (BMI >30) | | | | |
| Patients, No. (%) | 31 (7.7) | 24 (9.4) | 7 (4.7) | .05 |
| SLNs per patient | 2.05 (64/31) | 1.82 (44/24) | 2.86 (20/7) | .07 |
| Positive SLNs per all SLNs | 0.14 (9/64) | 0.09 (4/44) | 0.25 (5/20) | <.001 |
| Positive SLNs per patient | 0.29 (9/31) | 0.16 (4/24) | 0.71 (5/7) | <.001 |

Abbreviations: BMI, body mass index, calculated as weight in kilograms divided by height in meters squared; SLNs, sentinel lymph nodes; SPECT/CT, single-photon emission computed tomography/computed tomography.

TEMP-TDM

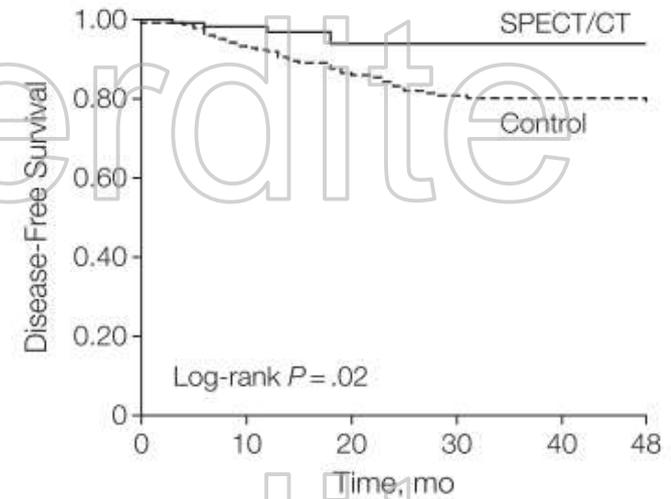
Améliore la survie sans récurrence ?

Table 3. Factors Associated With Disease-Free Survival for Patients With Melanoma Who Underwent SLNE Aided by Preoperative SPECT/CT

| Variable | No. (%) of Patients | Univariate Analysis | | Multivariate Analysis | |
|----------------------------------|---------------------|---------------------|---------|-----------------------|---------|
| | | HR (95% CI) | P Value | HR (95% CI) | P Value |
| SPECT/CT | | | | | |
| Yes | 149 (36.9) | 1 [Reference] | | 1 [Reference] | |
| No | 254 (63.1) | 3.21 (1.14-9.06) | .03 | 4.11 (1.25-13.51) | .02 |
| Histology | | | | | |
| SSM | 165 (40.9) | 1 [Reference] | | 1 [Reference] | |
| NM | 141 (35.0) | 2.88 (1.43-5.81) | .003 | 1.41 (0.03-3.12) | .40 |
| Other | 97 (24.1) | 1.48 (0.57-3.81) | .42 | 1.43 (0.51-4.04) | .49 |
| Sex | | | | | |
| Female | 159 (39.5) | 1 [Reference] | | 1 [Reference] | |
| Male | 244 (60.5) | 0.80 (0.44-1.43) | .45 | 0.44 (0.22-0.92) | .03 |
| Primary localization | | | | | |
| Head or neck | 38 (9.4) | 1 [Reference] | | 1 [Reference] | |
| Hand or foot | 21 (5.2) | 0.51 (0.05-4.95) | .56 | 0.15 (0.13-1.70) | .13 |
| Torso | 175 (43.3) | 1.34 (0.40-4.49) | .63 | 0.91 (0.23-3.51) | .89 |
| Extremity | 169 (41.9) | 1.24 (0.37-4.21) | .73 | 0.34 (0.08-1.41) | .14 |
| Age, per y | | 1.01 (0.99-1.03) | .45 | | >.99 |
| Breslow thickness, per mm | 403 (100) | 1.26 (1.15-1.38) | <.001 | 1.16 (1.02-1.32) | .02 |
| Ulceration | | | | | |
| Absent | 289 (71.7) | 1 [Reference] | | 1 [Reference] | |
| Present | 114 (28.3) | 3.69 (2.04-6.67) | <.001 | 2.07 (1.07-3.99) | .03 |
| Sentinel node status | | | | | |
| Negative | 314 (77.9) | 1 [Reference] | | 1 [Reference] | |
| Positive | 89 (22.1) | 5.02 (2.79-9.03) | <.001 | 4.14 (2.14-7.98) | <.001 |

Abbreviations: HR, hazard ratio; NM, nodular melanoma; SLNE, sentinel lymph node excision; SPECT/CT, single-photon emission computed tomography/computed tomography; SSM, superficial spreading melanoma.

Figure 1. Kaplan-Meier Curve With the SPECT/CT-Specific Disease-Free Survival Over 48 Months



| No. at risk | 0 | 10 | 20 | 30 | 40 | 48 |
|-------------|-----|-----|-----|-----|-----|----|
| SPECT/CT | 149 | 84 | 27 | 17 | 5 | |
| Control | 254 | 203 | 163 | 123 | 102 | |

TEMP-TDM

Améliore la survie sans récurrence ?

- Etude cas-témoin avant/après
 - Biais temporel
 - +++ pour critères de jugements temporels (survie)
 - Mais
 - plus de GS+
 - Analyse multivariée

TEMP-TDM

- Optionnelle (EANM 2009^[4])
- Bénéfice pour le patient
- Infos pertinentes pour le chirurgien
- Coût modique (France)
 - ZZQL013 (313,08 €)
 - FCQL001 + ZZQL007 (410,51 €)



Ganglion sentinelle et mélanome

- Introduction
- Indications
- Technique
- **Résultats**
- Perspectives

Résultats

[12]
Lymphatic Mapping and Sentinel Lymph Node Biopsy in
Patients With Melanoma: A Meta-Analysis

Matias E. Valsecchi, Damian Silbermins, Nicole de Rosa, Sandra L. Wong, and Gary H. Lyman

- Méta-analyse

- 78 articles, 25 240 patients

- Taux détection 98,1 %

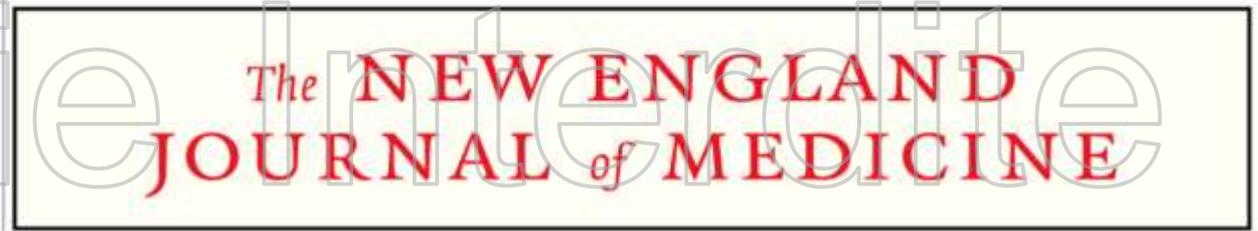
- Taux FN 12,5 % (vs récurrence ganglionnaire)

- Taux Faux Négatifs 0-5% (vs curage)

→ **Le GS permet une stadification
ganglionnaire fiable**

Résultats

- MSLT-I



Final Trial Report of Sentinel-Node Biopsy versus Nodal Observation in Melanoma [13]

D.L. Morton, J.F. Thompson, A.J. Cochran, N. Mozzillo, O.E. Nieweg, D.F. Roses, H.J. Hoekstra, C.P. Karakousis, C.A. Puleo, B.J. Coventry, M. Kashani-Sabet, B.M. Smithers, E. Paul, W.G. Kraybill, J.G. McKinnon, H.-J. Wang, R. Elashoff, and M.B. Faries, for the MSLT Group*

1661 Patients underwent randomization:
10-yr follow-up

1347 Had intermediate-thickness primary melanomas

1,2 – 3,5 mm

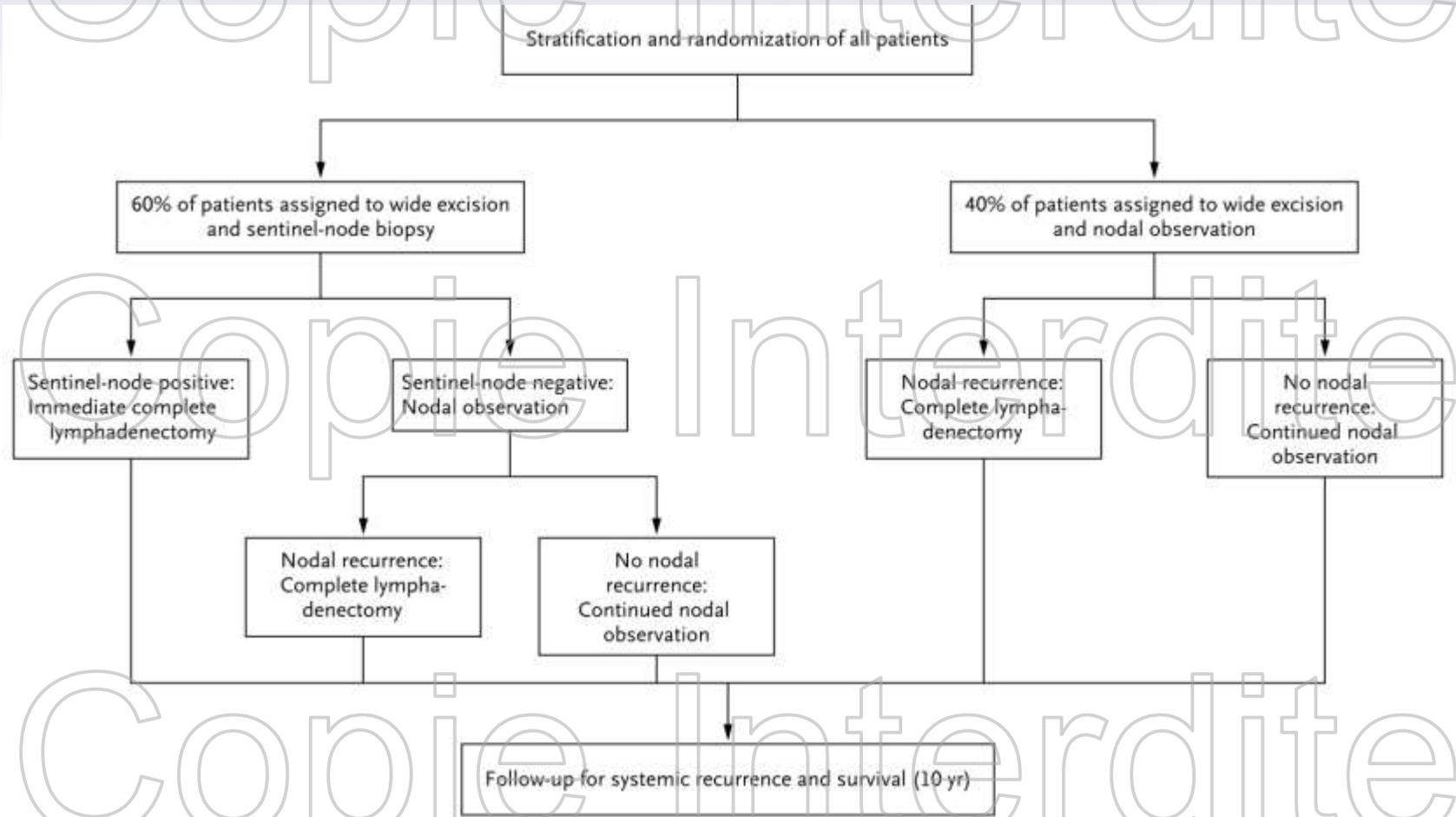
(Analyse post hoc 1-4 mm,

résultats disponibles en ligne similaires)

314 Had thick primary melanomas

>3,5 mm

MSLT-1



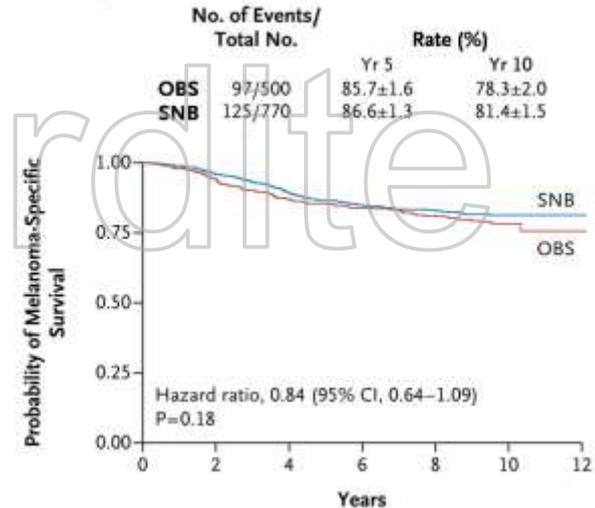
MSLT-1

- Objectif principal

- Survie spécifique à 10 ans

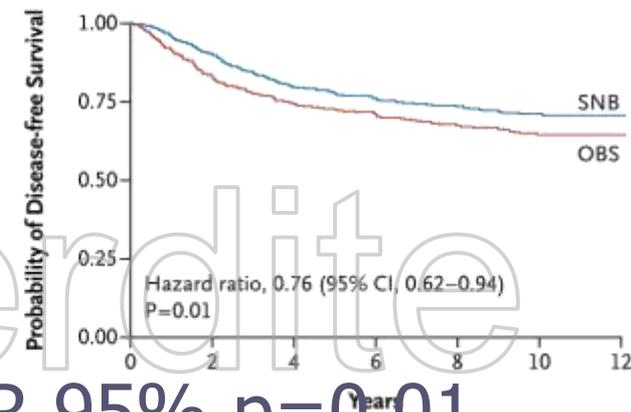
- **Négatif** ($81,4 \pm 1,5\%$ vs $78,3 \pm 2\%$ $p=0,18$)

A Melanoma-Specific Survival, Intermediate-Thickness Melanomas



C Disease-free Survival, Intermediate-Thickness Melanomas

| | No. of Events/ Total No. | Yr 5 Rate (%) | Yr 10 Rate (%) |
|-----|-----------------------------|------------------|-------------------|
| OBS | 161/500 | 72.7 ± 2.1 | 64.7 ± 2.3 |
| SNB | 199/770 | 77.8 ± 1.6 | 71.3 ± 1.8 |



- Objectif secondaire

- Survie sans récurrence

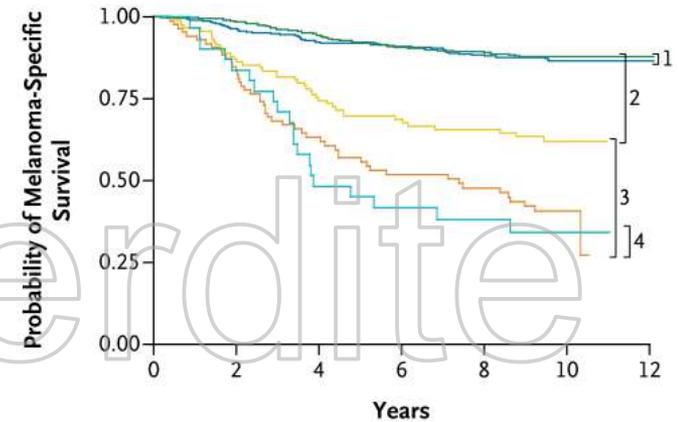
- $71,3 \pm 1,8\%$ vs $64,7 \pm 2,3\%$ HR 95% $p=0,01$

MSLT-1

- Analyse en sous groupe
 - Survie spécifique à 10 ans

C Melanoma-Specific Survival, Intermediate-Thickness Melanomas

| | No. of Events/ Total No. | Rate (%) | |
|--------------------------|-----------------------------|----------|----------|
| | | Yr 5 | Yr 10 |
| OBS, no nodal recurrence | 48/413 | 92.0±1.4 | 86.6±1.8 |
| OBS, nodal recurrence | 49/87 | 57.5±5.4 | 41.5±5.6 |
| SNB, true neg. | 63/612 | 92.3±1.1 | 88.0±1.4 |
| SNB, pos. | 41/122 | 69.8±4.4 | 62.1±4.8 |
| SNB, false neg. | 20/31 | 45.2±8.9 | 34.4±8.7 |

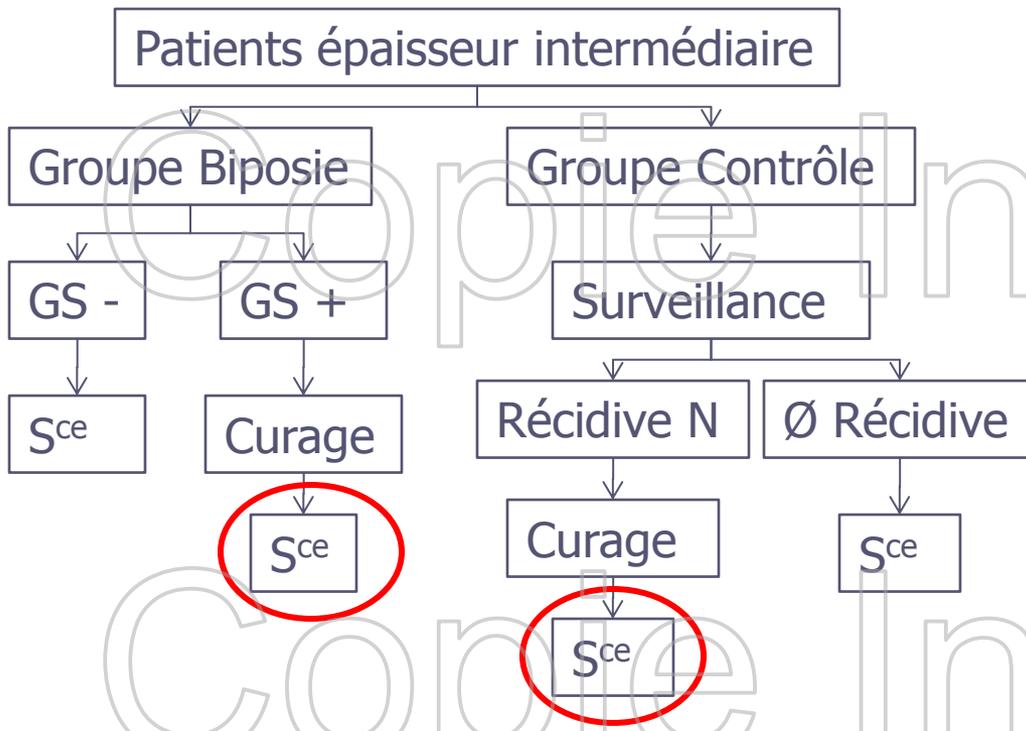


| No. at Risk | 0 | 2 | 4 | 6 | 8 | 10 | 12 |
|--------------------------|-----|-----|-----|-----|-----|-----|----|
| OBS, no nodal recurrence | 413 | 375 | 339 | 311 | 282 | 175 | 4 |
| OBS, nodal recurrence | 87 | 73 | 51 | 40 | 36 | 16 | 0 |
| SNB, true neg. | 612 | 570 | 511 | 448 | 395 | 243 | 5 |
| SNB, pos. | 122 | 100 | 81 | 68 | 60 | 31 | 0 |
| SNB, false neg. | 31 | 26 | 15 | 12 | 10 | 6 | 0 |

1. SNB, neg. vs. OBS, no nodal recurrence: HR, 0.89 (95% CI, 0.61–1.29); P=0.54
2. SNB, pos. vs. SNB, neg.: HR, 3.93 (95% CI, 2.65–5.83); P<0.001
3. SNB, pos. vs. OBS, nodal recurrence: HR, 0.56 (95% CI, 0.37–0.84); P=0.006
4. SNB, false neg. vs. OBS, nodal recurrence: HR, 1.15 (95% CI, 0.68–1.94); P=0.60

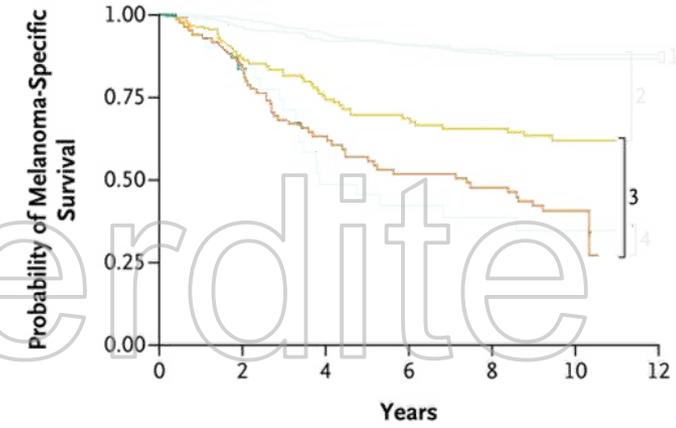
MSLT-1

• Analyse en sous groupe



C Melanoma-Specific Survival, Intermediate-Thickness Melanomas

| | No. of Events / Total No. | Rate (%) | |
|----------------------------|---------------------------|----------|----------|
| | | Yr 5 | Yr 10 |
| — OBS, no nodal recurrence | 48/413 | 92.0±1.4 | 86.6±1.8 |
| — OBS, nodal recurrence | 49/87 | 57.5±5.4 | 41.5±5.6 |
| — SNB, true neg. | 63/612 | 92.3±1.1 | 88.0±1.4 |
| — SNB, pos. | 41/122 | 69.8±4.4 | 62.1±4.8 |
| — SNB, false neg. | 20/31 | 45.2±8.9 | 34.4±8.7 |



| No. at Risk | 413 | 375 | 339 | 311 | 282 | 175 | 4 |
|--------------------------|-----|-----|-----|-----|-----|-----|---|
| OBS, no nodal recurrence | 413 | 375 | 339 | 311 | 282 | 175 | 4 |
| OBS, nodal recurrence | 87 | 73 | 51 | 40 | 36 | 16 | 0 |
| SNB, true neg. | 612 | 570 | 511 | 448 | 395 | 243 | 5 |
| SNB, pos. | 122 | 100 | 81 | 68 | 60 | 31 | 0 |
| SNB, false neg. | 31 | 26 | 15 | 12 | 10 | 6 | 0 |

1. SNB, neg. vs. OBS, no nodal recurrence: HR, 0.61 (95% CI, 0.46–1.29); P=0.54

3. SNB, pos. vs. OBS, nodal recurrence: HR, 0.56 (95% CI, 0.37–0.84); P=0.006

SNB, pos. vs. OBS, nodal recurrence: HR, 0.56 (95% CI, 0.37–0.84); P=0.006

62,1 vs 41,5 %

Résultats

- GS performant pour connaître le statut N
- Élément pronostic majeur
- Intérêt sur la survie plus incertain

Ganglion sentinelle et mélanome

- Introduction
- Indications
- Technique
- Résultats
- **Perspectives**

Perspectives

- Intérêt du curage complémentaire ?

- 80 à 85 % de curage « blanc »

- Nombreux facteurs

prédictifs

Taille >2mm,
nombre,
envahissement capsulaire

.....

- Essai randomisé MSLT II (clos, résultats 2022)
- Essai observationnel EORTC 1208



Available online at www.sciencedirect.com

SciVerse ScienceDirect

EJSO 39 (2013) 669–680

Review

[14]

Is complete lymph node dissection after a positive sentinel lymph node biopsy for cutaneous melanoma always necessary? A meta-analysis

V. Nagaraja, G.D. Eslick*

The Whiteley-Martin Research Centre, Discipline of Surgery, The University of Sydney, Nepean Hospital, Penrith, New South Wales, Australia

Accepted 20 February 2013
Available online 6 April 2013

EJSO

the Journal of Cancer Surgery

www.ejso.com

Perspectives

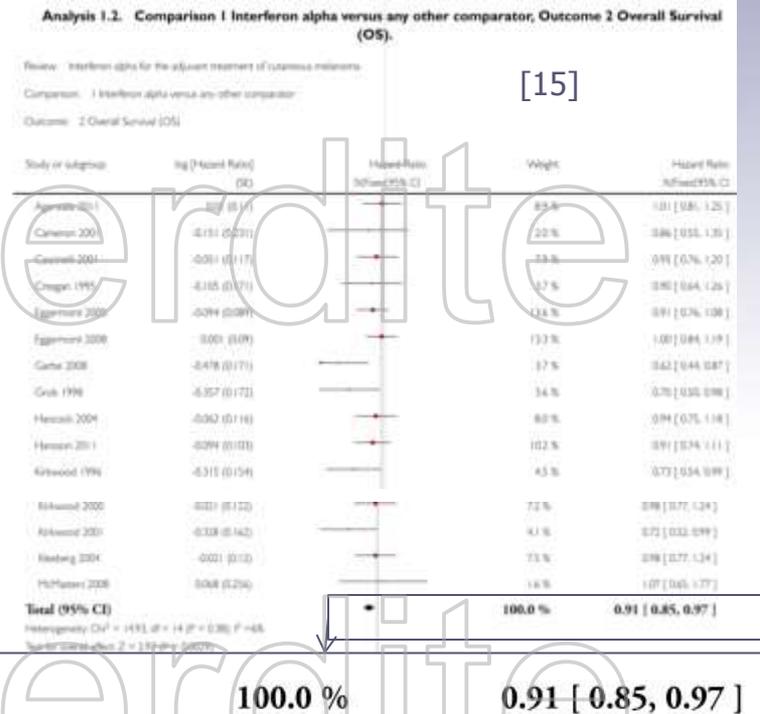
- Traitements adjuvants ?

- Interféron alpha

- Méta-analyse Mocellin
 - 54 études, 8388 patients
 - Patients à risque (II-III)
(>T2b : >1mm +Ulcération)

- Nouvelles stratégies ?

- Ac Anti CLTA4 (Ipilimumab)
 - Ac anti PD1 (nivolumab et pembrolizumab)
 - Inhibiteur de B-RAF



Perspectives ?



Adjuvant ipilimumab versus placebo after complete resection of high-risk stage III melanoma (EORTC 18071): a randomised, double-blind, phase 3 trial

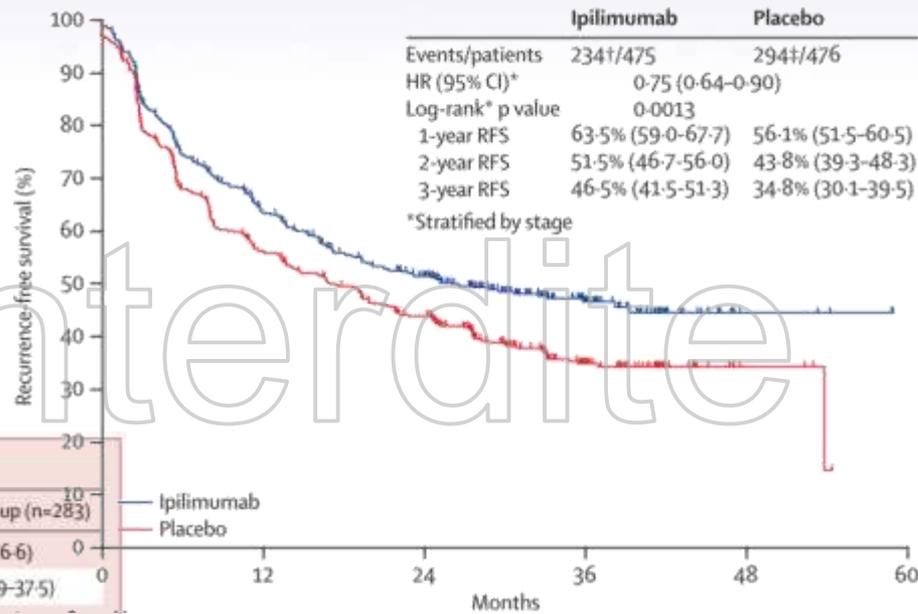
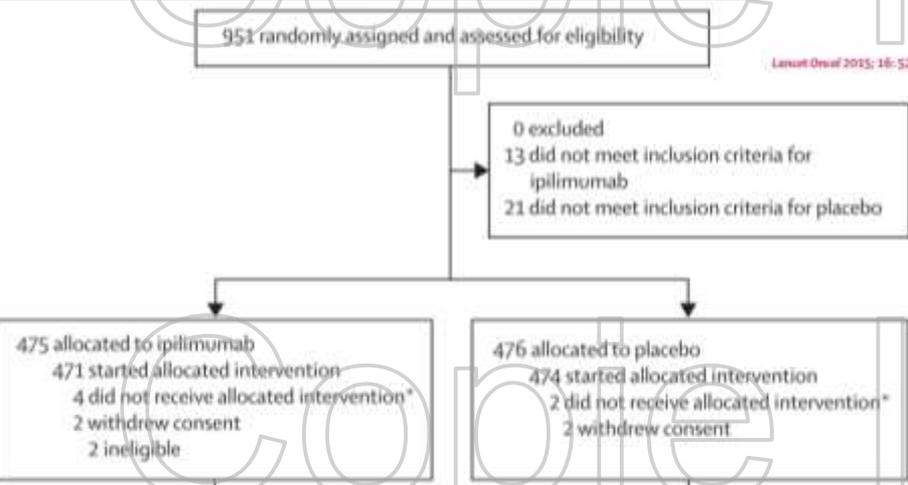
[16]

Alexander M M Eggermont, Vanna Chiarion-Gilani, Jean-Jacques Grob, Reinhard Dummer, Jedd D Wolchok, Henrik Schmidt, Ornid Hamid, Caroline Robert, Paolo A Ascierto, Jon M Richards, Celeste Lebbé, Virginia Ferraresi, Michael Smylie, Jeffrey S Weber, Michele Maio, Cynl Kanto, Axel Hoos, Verle de Prie, Ravichandra Karra Gurusath, Goeran de Scharetzen, Stefani Socio, Alessandro Testori

Summary

Background Ipilimumab is an approved treatment for patients with advanced melanoma. We aimed to assess

Lancet Oncol 2015; 16: 522-30



| | Microscopic stage III* (positive sentinel nodes) | | Macroscopic stage III* (palpable nodes) | |
|---------------------|--|-----------------------|---|-----------------------|
| | Ipilimumab group (n=210) | Placebo group (n=193) | Ipilimumab group (n=265) | Placebo group (n=283) |
| Median RFS (months) | NR (38.4-NR) | 26.9 (19.3-32.9) | 15.4 (11.3-22.9) | 11.3 (8.1-16.6) |
| 3-year RFS | 57.6% (50.0-64.4) | 39.2% (31.4-47.0) | 37.8% (31.3-44.2) | 31.7% (25.9-37.5) |
| HR | 0.65 (0.45-0.96)† | - | 0.81 (0.61-1.08)† | - |
| p value | 0.004† | - | 0.06† | - |

In the ipilimumab group, 465 (99%) of 471 patients had an adverse event of any grade, with grade 3 or 4 adverse events in 254 (54%) patients; 432 (91%) of 474 patients in

Of 471 patients who started ipilimumab, 245 (52%) discontinued treatment because of an adverse event, of which 230 (49%) were drug-related; in 182 (39%) patients

Five (1%) participants died because of drug-related adverse events in the ipilimumab group; three patients died because of colitis (two with gastrointestinal perforation), one patient because of myocarditis, and one

Perspectives

- Nouveaux traceurs ?

Copie Interdite

Copie Interdite

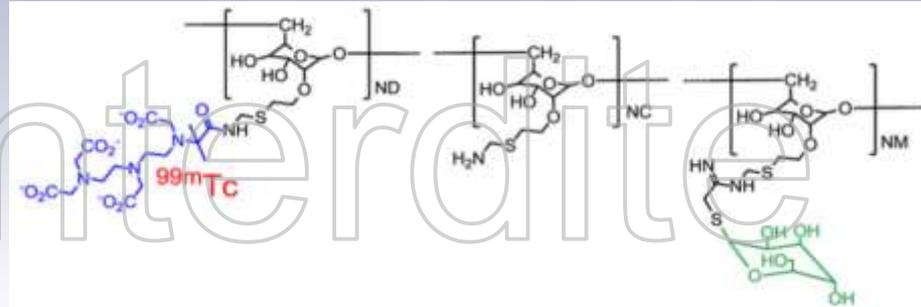
Perspectives

- ^{99m}Tc -Tilmanocept

- mélanome : comparé au bleu patenté

- Meilleur spécificité dans le cancer du sein

- Moins de GS identifiés donc enlevés, mais autant de GS envahis



Ann Surg Oncol (2013) 20:680–688
DOI 10.1245/s10434-012-2612-z

Annals of
SURGICAL ONCOLOGY
OFFICIAL JOURNAL OF THE SOCIETY OF SURGICAL ONCOLOGISTS

ORIGINAL ARTICLE - MELANOMAS

[17]

Combined Analysis of Phase III Trials Evaluating [^{99m}Tc]Tilmanocept and Vital Blue Dye for Identification of Sentinel Lymph Nodes in Clinically Node-Negative Cutaneous Melanoma

Vernon K. Sondak, MD¹, Dennis W. King, PhD², Jonathan S. Zager, MD¹, Schlomo Schneebaum, MD³, Julian Kim, MD⁴, Stanley P. L. Leong, MD⁵, Mark B. Faries, MD⁶, Bruce J. Averbook, MD⁷, Steve R. Martinez, MD⁸, Christopher A. Puleo, PA-C¹, Jane L. Messina, MD¹, Lori Christman, PhD², and Anne M. Wallace, MD⁹

Clin Exp Metastasis (2012) 29:681–686
DOI 10.1007/s10585-012-9497-x

RESEARCH PAPER

[18]

The efficacy of Tilmanocept in sentinel lymph node mapping and identification in breast cancer patients: a comparative review and meta-analysis of the ^{99m}Tc -labeled nanocolloid human serum albumin standard of care

Christopher A. Tokin · Frederick O. Cope · Wendy L. Metz · Michael S. Blue · Beth M. Potter · Bonnie C. Abbruzzese · Richard D. Hartman · Marcus T. Joy · Dennis W. King · Lori A. Christman · David R. Vera · Anne M. Wallace

Perspectives

- Oxyde de fer super-paramagnétique

- Non évalué dans le mélanome

- Un seul cas clinique publié : drainage inhabituel

- Etude SentiMAG



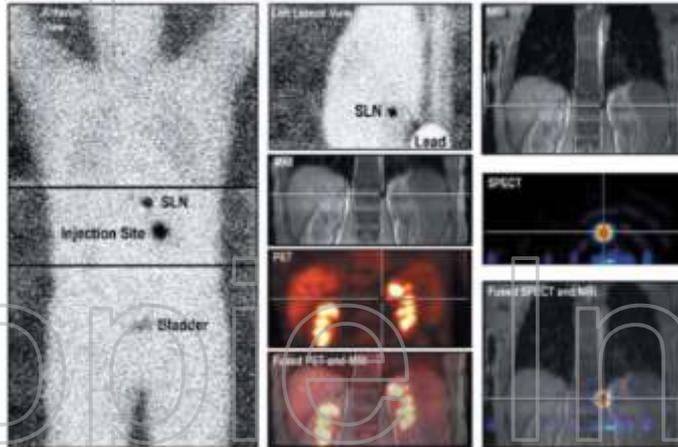
Precise localisation of a sentinel lymph node in a rare drainage region with ^{99m}Tc -nanocolloid a

Sofiane Maza¹, Matthias Taupitz¹, Thors
¹ Clinic for Nuclear Medicine, Charité University

Published online: 4 September 2004
 © Springer-Verlag 2004

[19]

Eur J Nucl Med Mol Imaging (2005) 30
 DOI 10.1007/s00259-004-1660-4



- Non inférieur dans le cancer du sein

- Etude SentiMAG

Ann Surg Oncol (2014) 21:1237–1245
 DOI 10.1245/s10434-013-3379-6

Annals of
SURGICAL ONCOLOGY
 OFFICIAL JOURNAL OF THE SOCIETY OF SURGICAL ONCOLOGISTS

ORIGINAL ARTICLE – BREAST ONCOLOGY

Sentinel Node Biopsy Using a Magnetic Tracer Versus Standard Technique: The SentiMAG Multicentre Trial

[20]

Michael Douek, MD^{1,2}, Joost Klaase, MD³, Ian Monypenny, MD⁴, Ashutosh Kothari, MS⁵, Katalin Zechmeister, MD⁶, Douglas Brown, MD⁸, Lynda Wyld, PhD⁷, Philip Drew, MD⁹, Hans Garino, PhD⁷, Olorunsola Agbaje, PhD⁸, Quentin Pankhurst, PhD⁷, Brooke Annings, MSc^{1,10}, Maarten Grootendorst, MSc^{1,10}, Bennie ten Haken, PhD¹⁰, Margaret A. Hall-Craggs, MD¹¹, Arnie Purushotham, MD^{1,2}, Sarah Pinder, MD^{1,2} and On behalf of the SentiMAG Trialists Group

Perspectives

- Imagerie fluorescence (Vert d'Indocyanine)



Intraoperative SN Detection

| | |
|---|-------------------|
| Clinical Data | All, 104 Patients |
| No. of excised SNs | 301 |
| No. of SNs not excised | 4 |
| All SNs | 305 |
| Intraoperative detection of SNs | |
| With gamma tracing | 286/305 (93.8) |
| With FI guidance | 295/305 (96.7) |
| With blue-dye visualization | 116/188 (61.7) |

Preoperative SN Mapping Results

| Clinical Data | Location Overall |
|------------------------------------|------------------|
| No. of patients | 104 |
| Total no. of SNs visualized* | 246 (2.4, 1–6) |
| Lymphoscintigraphy [†] | 232/246 (94.3) |
| SPECT/CT [†] | 245/246 (99.6) |
| Portable gamma camera [†] | 1/246 (0.4) |

Multimodal Surgical Guidance during Sentinel Node Biopsy for Melanoma: Combined Gamma Tracing and Fluorescence Imaging of the Sentinel Node through Use of the Hybrid Tracer Indocyanine Green-^{99m}Tc-Nanocolloid¹

Purpose: To evaluate the hybrid approach in a large population of

Implications for Patient Care

- FI of the hybrid tracer, compared with blue dye, provided improved SN visualization.
- Accurate guidance during SN biopsy has the potential to reduce morbidity associated with the SN biopsy procedure.

Perspectives



Groupe Hospitalier Universitaire
SAINT-LOUIS
LARIBOISIÈRE
FERNAND-WIDAL

https://www.youtube.com/watch?v=Hahkwe_uoqI

Résultats

- GS performant pour connaître le statut N
- Intérêt de la TEMP-TDM
- Intérêt « direct » sur la survie plus incertain
- Intérêt probable pour sélectionner les patients devant bénéficier d'un traitement adjuvant

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