

Melanoma



Melanoma

Clinical manifestations

Risk factors

Clinical diagnosis and diagnostic techniques

Staging

Patient care pathway

Prevention



Melanoma

Some figures

2000-2500 new diagnoses of MM per year in Belgium

11 millions: 0,018-0,023%

1/10 patients with MM will develop a 2nd MM in 10 years

50% of MM diagnosis is made by the patient him/herself

Most frequent site:
- males: back
- females: legs

50% of MM develop in pre-existing naevi
50% of MM develop de novo

100-150 MM related deaths per year



Melanoma

Some figures

Lifetime risk for MM: males: 1 in 28

Lifetime risk for MM: females: 1 in 44

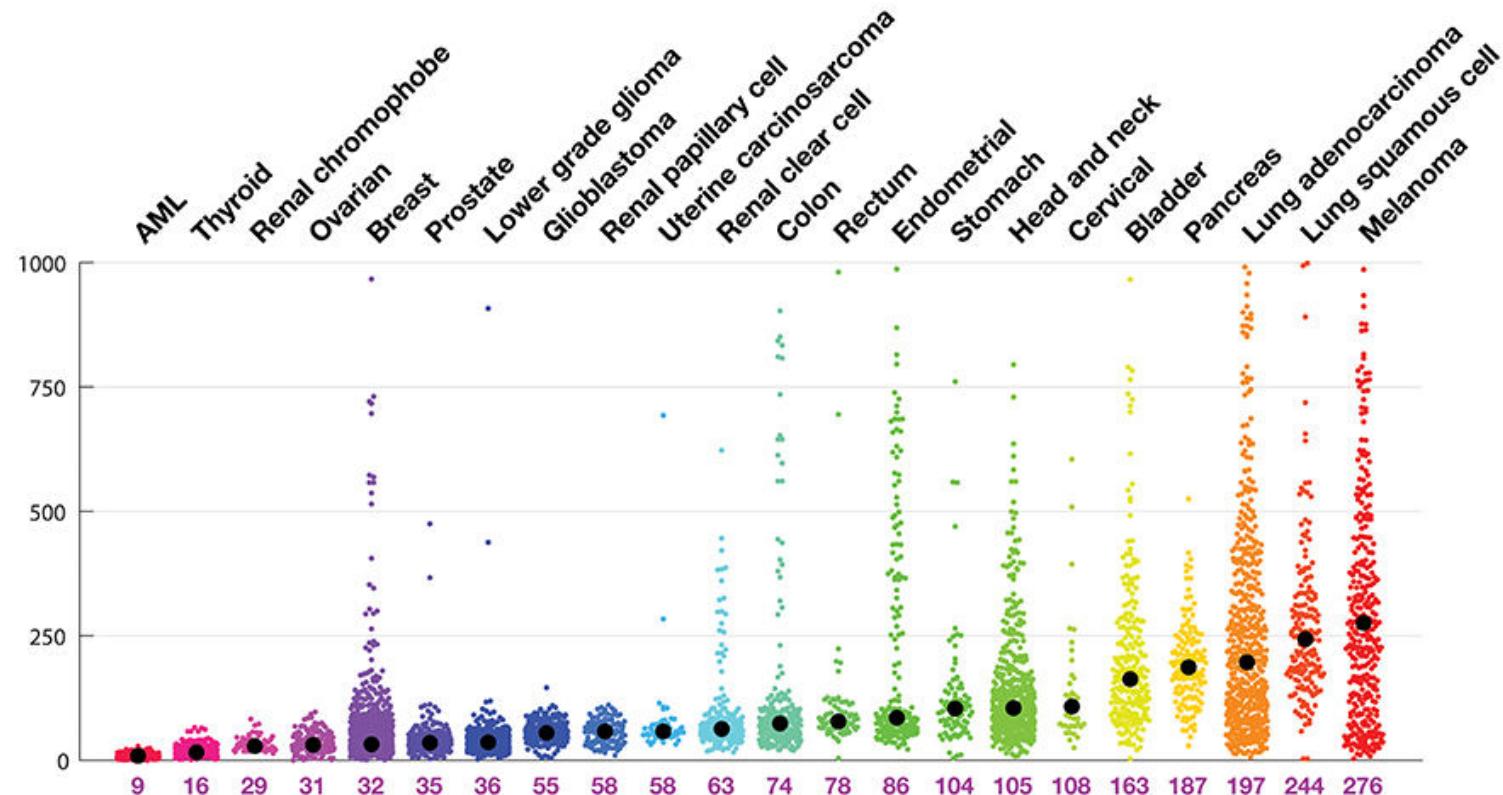
USA: 2017: invasive MM: 87100
in situ MM: 63000

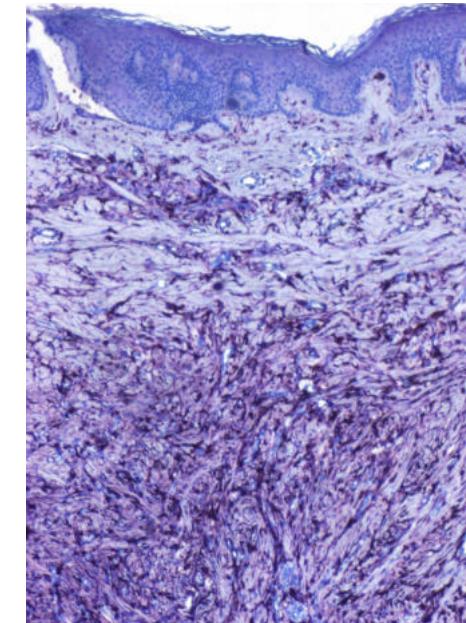
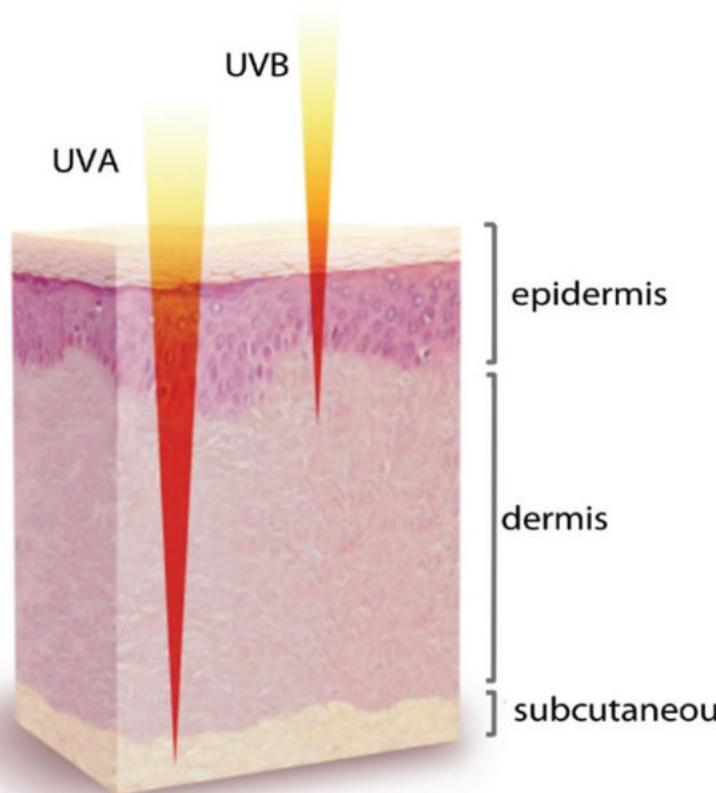
2016: 323 millions: invasive 0,027%
in situ 0,019%
0,046%

Survival rates (5-y) White Black

	White	Black
Localized	98	90
Regional	63	47
Metastatic	18	22

Mutational charge of melanoma





Melanoma Clinical manifestations

Dubreuilh's melanoma



Melanoma Clinical manifestations

Dubreuilh's melanoma

10% of the cases

Difficult clinical margins

Long indolent periods

Hidden deep invasion





Melanoma Clinical manifestations

Dubreuilh's melanoma



Melanoma Clinical manifestations

SSM melanoma

70% of MM cases





Melanoma Clinical manifestations

SSM melanoma





Melanoma Clinical manifestations

SSM melanoma



Melanoma Clinical manifestations

Nodular melanoma



5% of MM cases

Melanoma Clinical manifestations

Nodular melanoma



Melanoma Clinical manifestations

Nodular melanoma





Melanoma Clinical manifestations

Acrolentiginous melanoma

5 % of MM cases





Melanoma Clinical manifestations

Acrolentiginous melanoma



Melanoma Clinical manifestations

Acrolentiginous melanoma





Melanoma Clinical manifestations

Atypical melanoma

10% of MM cases

Melanoma masquerading as nonmelanocytic lesions.

Detrixhe A, Libon F, Mansuy M, Nikkels-Tassoudji N, Rorive A, Arrese JE, Quatresooz P, Reginster MA, Nikkels AF.

Melanoma Res. 2016 Dec;26(6):631-634.





Melanoma Clinical manifestations

Metastatic melanoma





Melanoma Clinical manifestations

Metastatic melanoma





Melanoma Clinical manifestations

Metastatic melanoma





Melanoma

Risk factors

- UV exposure
- Number of moles (both normal and atypical)
- Fair skin, freckling, light hair
- Family history of melanoma
- Personal history of melanoma
- Immunosuppression

OTR: risk increases, prognosis worse

Donor-derived MM:
Screening of donor

- Age: Risk increases with age
- Gender: Males > females
- Xeroderma pigmentosum

Women with MM during pregnancy and 1 st year after:
increased risk for recurrence



Melanoma

Risk factors

Tall stature: increased risk

Coffee:
chemopreventive effect?

PDE-5 inhibitors (Viagra)
Increased risk for MM and BCC

- UV exposure
- Number of moles (both normal and atypical)
- Fair skin, freckling, light hair
- Family history of melanoma
- Personal history of melanoma
- Immunosuppression
- Age: Risk increases with age
- Gender: Males > females
- Xeroderma pigmentosum

Sildenafil: idem (needs confirmation)



Melanoma

Risk factors

Figure 3 – Clinical risk factors for melanoma ⁽¹⁾

	Relative Risk
Genetic Factors	
1. Strong family history	35-70
2. Weak family history	3
Naevi	
1. Multiple benign naevi (>100)	11
2. Multiple atypical naevi	11
Previous skin factors	
1. Previous melanoma	8.5
2. Previous non-melanoma skin cancer	2.9
Immunosuppression	
1. Transplant recipients	3
2. AIDS	1.5
Sun sensitivity	
1. Type 1 skin (sunburn easily)	1.7
2. Freckling	2.5
3. Blue eyes	1.6
4. Red hair	2.4
UV exposure	
1. History of blistering sunburn	2.5

Please remember:

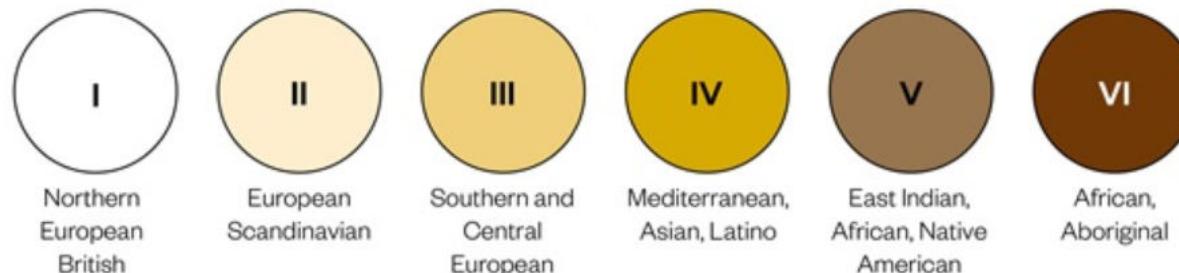
A dysplastic naevus is
NOT a precancerous lesion



Melanoma

Risk factors

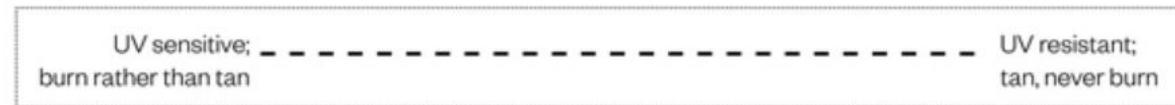
Pigmentary phototype (Fitzpatrick scale)



Epidermal melanin



UV phenotype



Cancer risk





Melanoma

Risk factors



TANNINGBEDS

INCREASE RISK OF MELANOMA BY

75%

WWW.CUREMELANOMA.ORG

Dysplastic naevi with family history of malignant melanoma

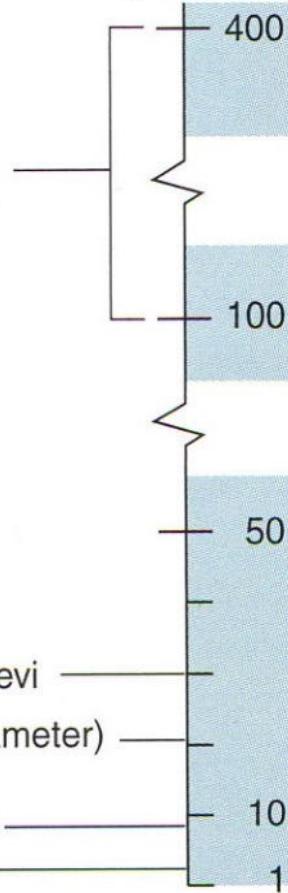
Multiple (>100) melanocytic naevi

Congenital naevus (>1.5 cm diameter)

Previous malignant melanoma

Red hair/blue eyes

burn in sun/tan poorly



Major risk factors

Naevi



Melanoma

Risk of subsequent MM after MM diagnosis

Cumulative lifetime risk of subsequent MM = 20%



Melanoma risk factors

Clinical diagnosis and diagnostic techniques

Clinical recognition - ABCDE Algorithm

NORMAL		CANCEROUS
	A: ASYMMETRY If you draw a line through the centre of the lesion, the two halves of a melanoma won't match.	
	B: BORDER IRREGULARITY The border of a melanoma is irregular, typically geographic; peninsulas, bays, islands.	
	C: COLOUR VARIATION Healthy moles are a uniform colour; A variety of different colours in the same lesion is suspicious.	
	D: DIAMETER > 6 MM Greater than 6 mm is suspicious, although melanomas can be smaller.	
	E: EVOLVING Recent change in size, shape or colour, or bleeding or scabbing are suspicious.	

Dermoscopy

Hand-held dermatoscopes



Oil-immersion

Polarized light





Melanoma risk factors

Clinical diagnosis and diagnostic techniques

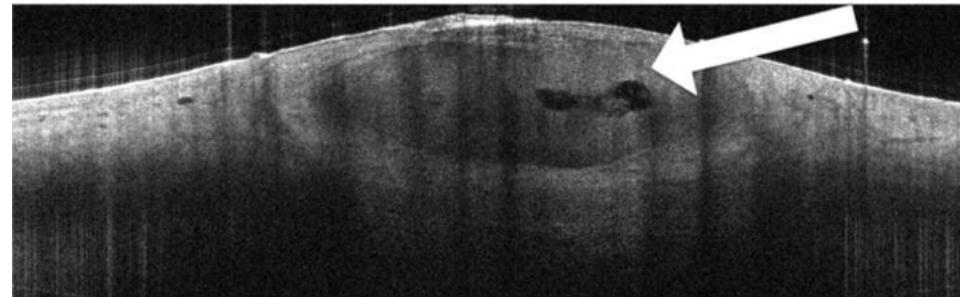
Nevisense



20 Mhz echography



Optical coherence tomography (OCT)



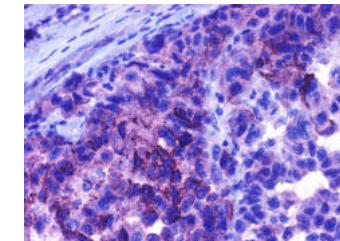
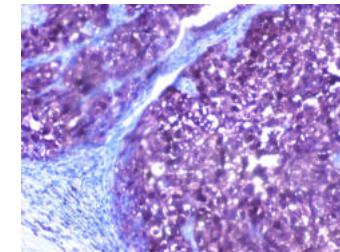
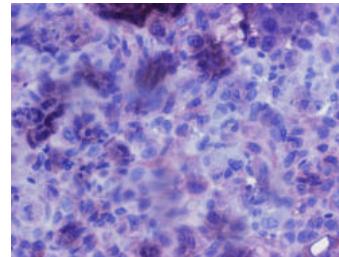
Histology and immunohistology

KI67 rate

Breslow depth

Tyrosinase

S-100a



Melan-A

HMB-45

BRAF

Genetic analysis

Excision

No biopsy



Melanoma risk factors

Clinical diagnosis and diagnostic techniques

GEP: gene expression profile

Melanoma 31-GEP test

Change in clinical approach ????????

MM tissue

RNA isolation

Microfluid PCR gene card

28 discriminant gene targets and 3 control genes

Analysis of GEP with algorithm

Class 1
Low metastatic risk

Class 2
High metastatic risk



Melanoma

MM survival

Diagnostic techniques

Diagnosis of early MM
Lesion depth:



Rapid growing MM
Amelanototic MM
Atypical MM



These MM are not responsible for a lethal outcome



Difficult and delayed diagnosis





Melanoma

Surveillance strategies

50% of MM develop
in pre-existing naevi



Clinical examination: Total body examination
Lesion-directed examination
High-risk patients: DDS

50% of MM develop
de novo



Rapid diagnosis consultation

Lesion-directed diagnosis



Regular
circuit



Rapid
circuit



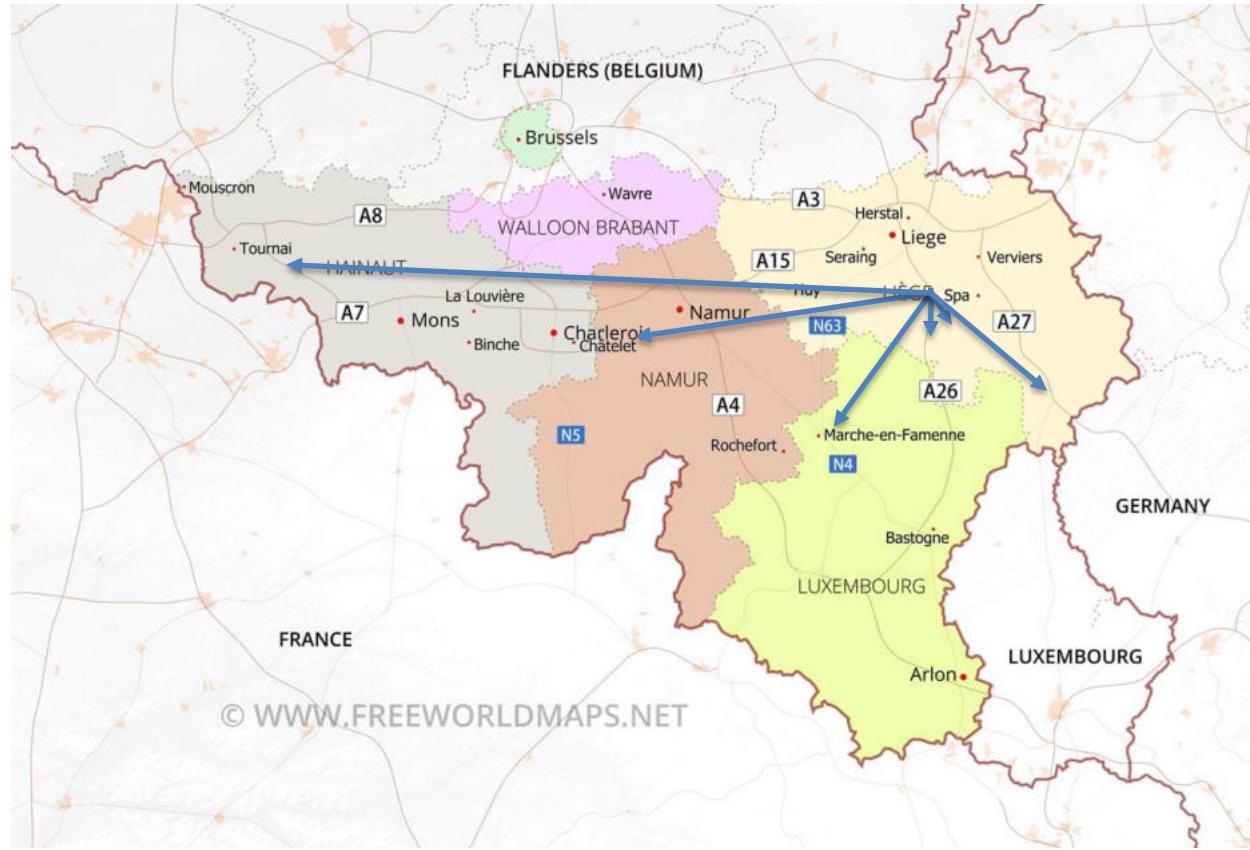
Melanoma

TELESPOT project

7 GP center

Teledermoscopy

Application on Iphone





AJCC Melanoma of the Skin Staging

8th
Edition

Definitions

Primary Tumor (T)

- T_X** Primary tumor cannot be assessed (for example, curretaged or severely regressed melanoma)
- T₀** No evidence of primary tumor
- T_{is}** Melanoma in situ
- T₁** Melanomas 1.0 mm or less in thickness
- T₂** Melanomas 1.1 - 2.0 mm
- T₃** Melanomas 2.1 - 4.0 mm
- T₄** Melanomas more than 4.0 mm

NOTE: a and b subcategories of T are assigned based on ulceration and thickness as shown below:

T CLASSIFICATION	THICKNESS (mm)	ULCERATION STATUS
T₁	≤1.0	a: Breslow < 0.8 mm w/o ulceration b: Breslow 0.8-1.0 mm w/o ulceration or ≤ 1.0 mm w/ ulceration.
T₂	1.1-2.0	a: w/o ulceration b: w/ ulceration
T₃	2.1-4.0	a: w/o ulceration b: w/ ulceration
T₄	>4.0	a: w/o ulceration b: w/ ulceration

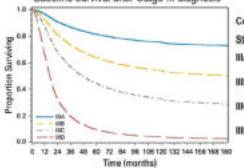
Regional Lymph Nodes (N)

- N_X** Patients in whom the regional nodes cannot be assessed (for example previously removed for another reason)
- N₀** No regional metastases detected
- N₁₋₃** Regional metastases based on the number of metastatic nodes, number of palpable metastatic nodes on clinical exam, and presence or absence of MSI²

NOTE: N₁₋₃ and a c subcategories assigned as shown below:

T CLASSIFICATION	# NODES	CLINICAL DETECTABILITY/STATUS
N₁	0-1 node	a: clinically occult ¹ , no MSI ² b: clinically detected ¹ , no MSI ² c: 0 nodes, MSI present ²
N₂	1-3 nodes	a: 2-3 nodes clinically occult ¹ , no MSI ² b: 2-3 nodes clinically detected ¹ , no MSI ² c: 1 node clinical or occult ¹ , MSI present ²
N₃	>1 nodes	a: >3 nodes, all clinically occult ¹ , no MSI ² b: >3 nodes, ≥1 detected ¹ or matted, no MSI ² c: >1 nodes clinical or occult ¹ , MSI present ²

Baseline survival after Stage III diagnosis³



Notes

¹Nodes are designated as 'clinically detectable' if they can be palpated on physical exam and are confirmed melanoma by pathology following excision biopsy.

²MSI comprise any satellite, locally recurrent, or in transit lesions.

³Clinical staging includes assessment of the primary melanoma and clinical/anthropologic evaluation of metastases. By convention, it must be done after a complete excision of the primary melanoma with clinical assessment for regional and distant metastases.

⁴Pathologic staging includes metastaging of the primary melanoma and pathologic information about the regional lymph nodes after partial or complete lymphadenectomy.

Pathologic Stage III and I patients are the exceptions; they do not necessarily require pathologic evaluation of their lymph nodes. Physicians should 'discuss and consider' SLNB for patients with T1b Stage IA disease; physicians should 'discuss and offer' SLNB for patients with Stage IIIB disease.

From Haydu et al., Journal of Clinical Oncology, 2017.



Department of Dermatology

Centre Hospitalier Universitaire de Liège

Staging



Melanoma



Department of Dermatology

Centre Hospitalier Universitaire de Liège

Patient care pathway

Trajet de soins en dermat-oncologie

Mélanome: algorithme diagnostique

Lésion pigmentaire suspecte



Examen clinique
Et/ou Loupe



Diagnostic clinique évident



Chirurgie



Diagnostic clinique pas évident



Dermoscopie



Origine
mélanocytaire



Lésion bénigne



Abstention
thérapeutique

Lésion douteuse



- DDS
- Nevisense
- Exérèse/biopsie pour diagnostic histopathologique

Lésion maligne

Pigmentation d'autres origines



Mélanome

Lésion pigmentaire hautement suspecte
Mélanome



Echographie 20 Mhz



> 1 mm d'épaisseur



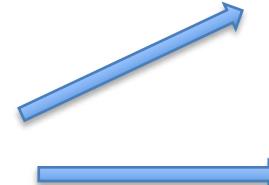
Excision chirurgicale
avec 2 cm de marge
Ganglion sentinelle si
possible



< 1 mm d'épaisseur



Excision chirurgicale avec 1 cm de marge



Histologie < 1mm mais

- Ulcération
- Mitoses: 2 ou >
- Micrométastases

Histologie < 1mm



Suivi clinique dermatologie

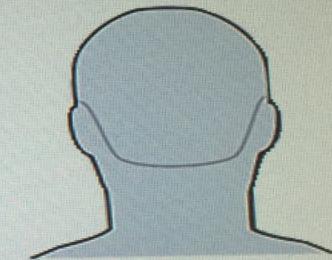
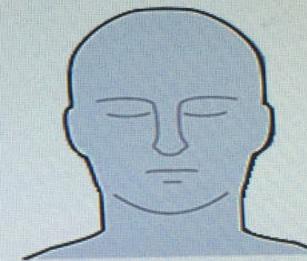
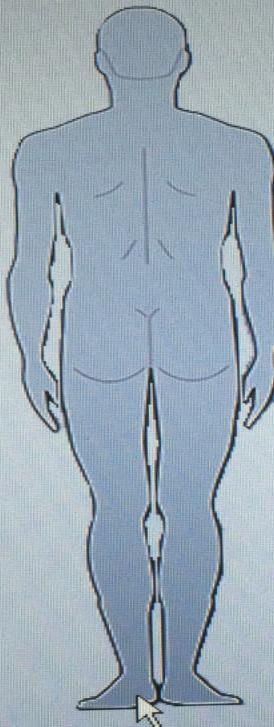




CHAMP CARTE DU CORPS



26/10/2015
11:08



Position de la lésion:

Bras, gauche

Âge:

64

Âge

Mesurer

Annuler



MESURE

ID du patient:
Lésion:

[ID-20151026-1108]
Bras, gauche 01



26/10/2015
11:09



Humidifiez la peau

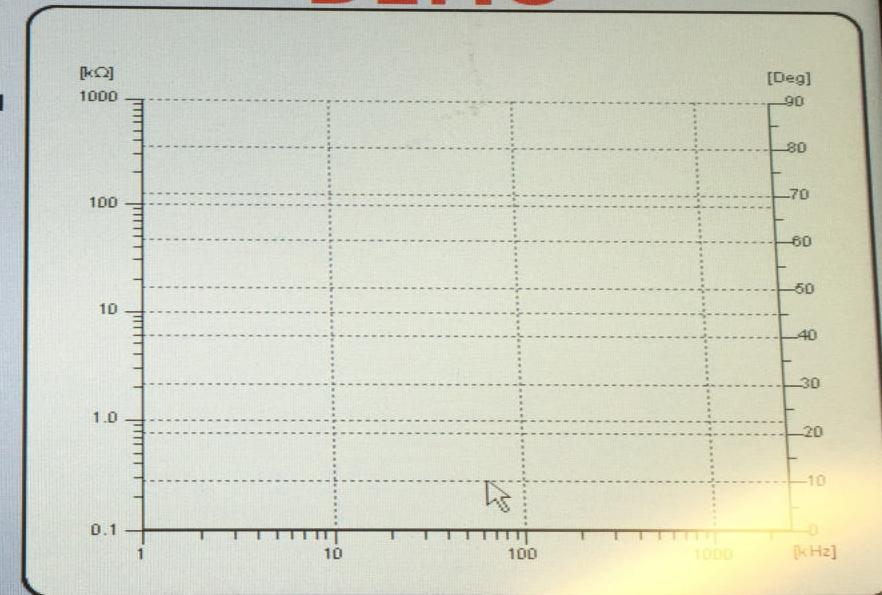


Essuyez



Mesurez

DEMO



Comptage des électrodes: DEMO

Annuler

Rejeter

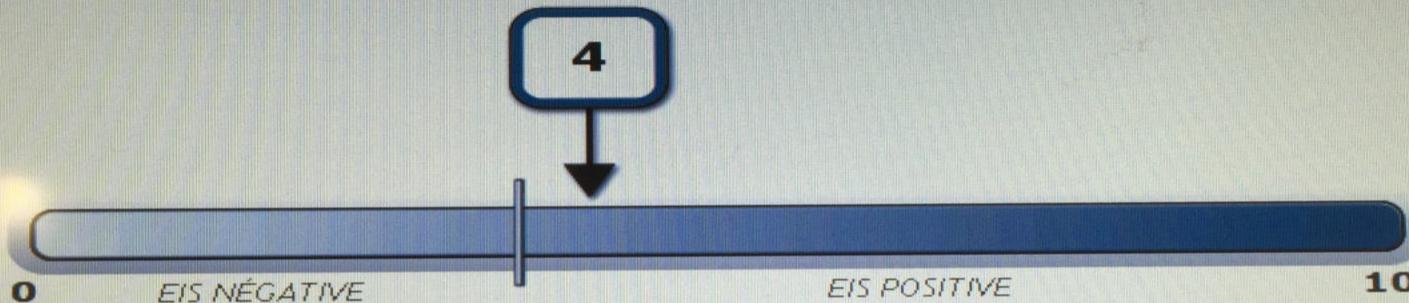
Terminé



SCORE EIS

DEMO

This is a demonstration result only.
The score is not valid and will not be saved.



Visualiser la référence clinique

REMARQUE : Score EIS positif. Un changement d'électrode peut être à recommander avant d'effectuer des mesures supplémentaires. Consultez les Instructions d'utilisation.

Créer rapport

Nouvelle
lésion

Terminé







ontal

Vertical

Arbitrary

Clear

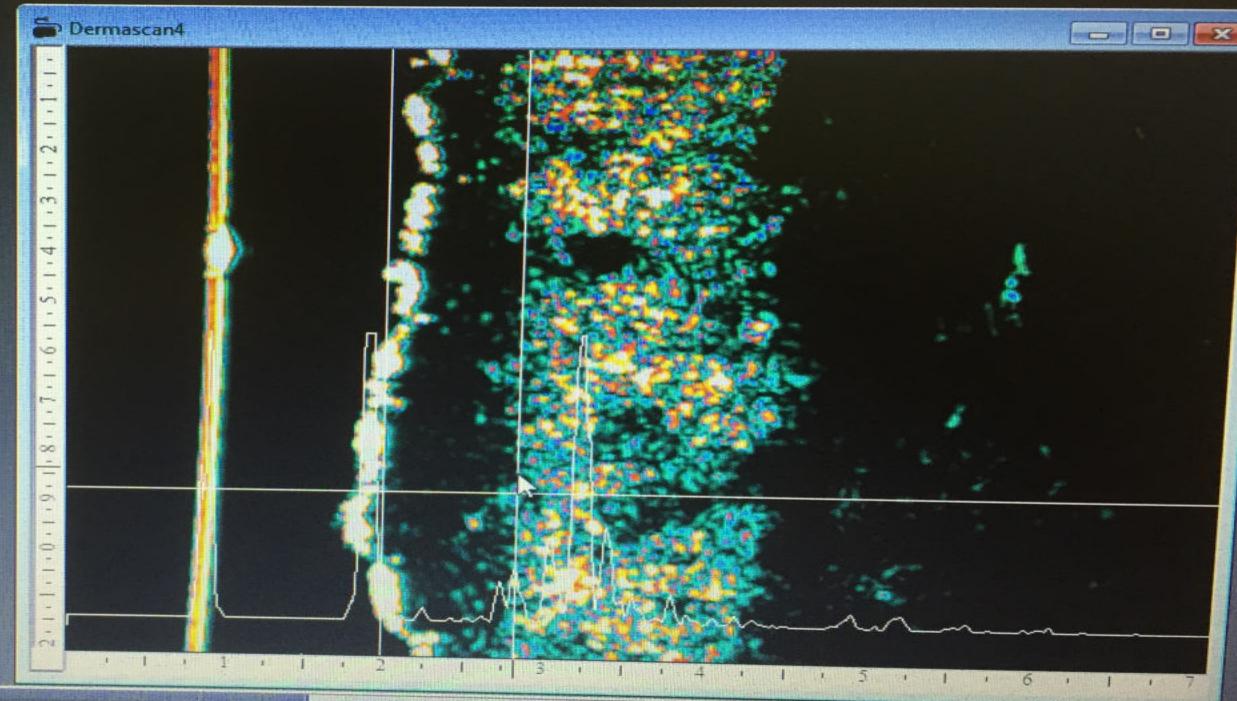
End

Export

A-Meas. A-Meas. Meas.

Measure

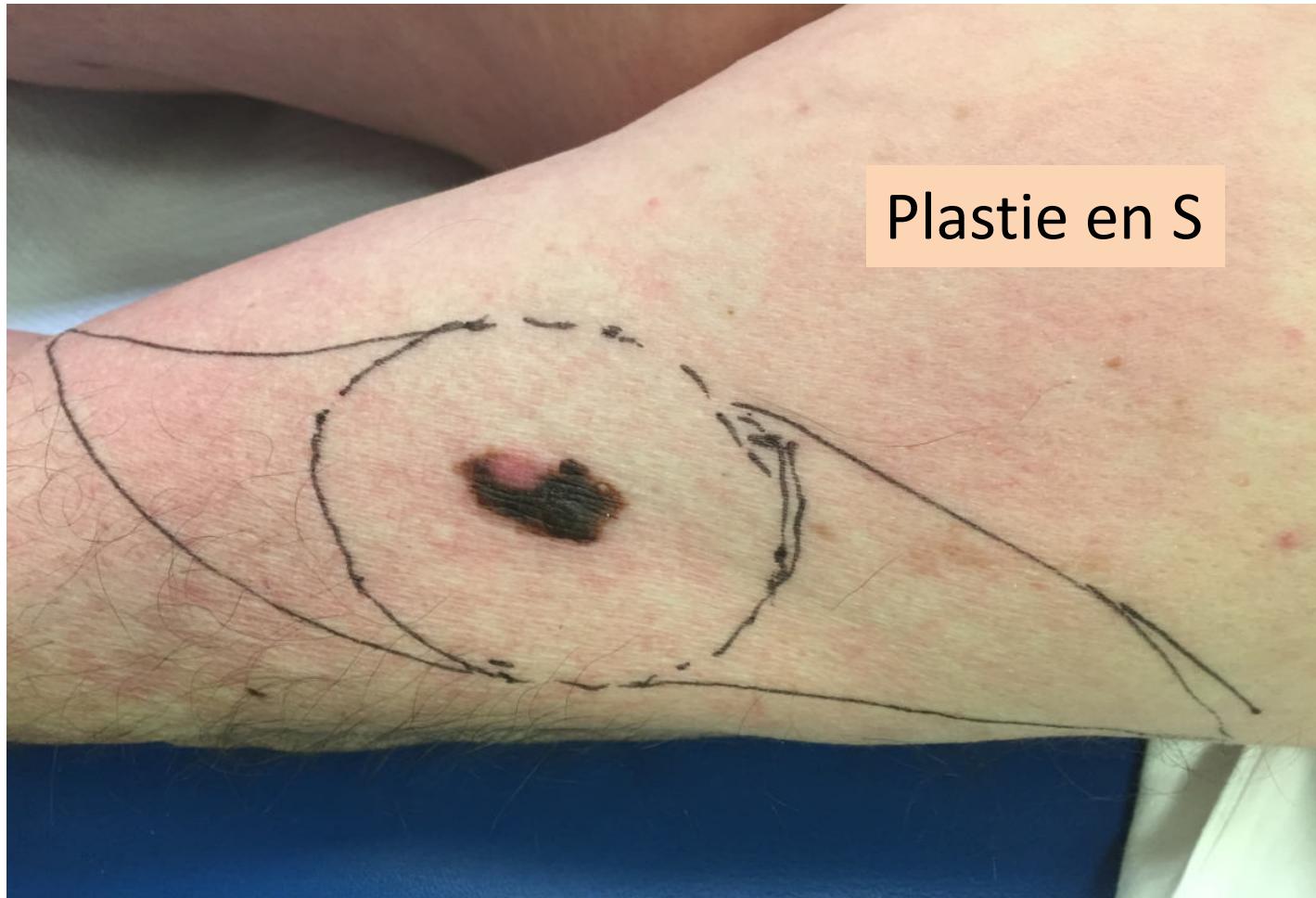
Properties	
Property	Value
+ Current Scan Settings	
- Stored Scan Settings	
Probe	B12.1 20MHz 13mm HS
Velocity in m/s	1580
Colorscale	High Contrast
Gain Profile	5
Stored Gain Profile Level	16
Custom Gain Profile	testprofile.gai
+ A-Scan Measures	
Horizontal Current (mm)	0.831579
Horizontal Average (mm)	0.415789
Vertical Current (mm)	0.000000
Vertical Average (mm)	0.000000
Arbitrary (mm)	0.000000
+ B-Scan Measures	
255	
D	





0,8 mm de profondeur à l'ultrason
20MHz

=> 1 cm de marge chirurgicale







Trajet de soins en dermato-oncologie

Mélanome

Marges d'excision cliniques

- MM in situ extrémité céphalique: 0,9 mm
- MM in situ corps: 0,5 mm
- MM < 1 mm: 1 cm
- MM > 1 mm, et/ou micrométastases, et/ou 2 mitoses ou >, et/ou ulcération: 2cm de marge

CAVE:

Attention: les marges cliniques ne correspondent pas aux marges histologiques
Les tissus rétrécissent dans le formol.



Délais d'interventions dermatologique:

Cas urgents: intervention dans la semaine

- *Mélanome érodé, charnu*

Cas non-urgents:

- *Mélanome de Dubreuilh*
- *Petit mélanome superficiel*

Mélanome



Trajet de soins en dermat-oncologie

Mélanome

Suivi clinique

Années	1, 2	3-5	6 et +
MM in situ (récurrent)	Dermato: 4 x/an	Dermato: 2 x/an	Dermato: 1 x/an
MM < 1 mm (récurrent)	Dermato: 4x/an	Dermato: 2 x/an	Dermato: 1 x/an
MM < 1 mm (récurrent) • Ulcération • Mitoses: 2 ou > • Micrométastases	Dermato: 4 x/an Onco: 4 x/an (Combiner RV)	Dermato: 2 x/an Onco 2 x/an (Alterner RV)	Dermato: 1 x/an Onco 1 x/an (Alterner RV)
MM > 1 mm (récurrent)	Dermato: 4 x/an Onco: 4 x/an (Combiner RV)	Dermato: 2 x/an Onco 2 x/an (Alterner RV)	Dermato: 1 x/an Onco 1 x/an (Alterner RV)

Trajet de soins en dermato-oncologie

Mélanome

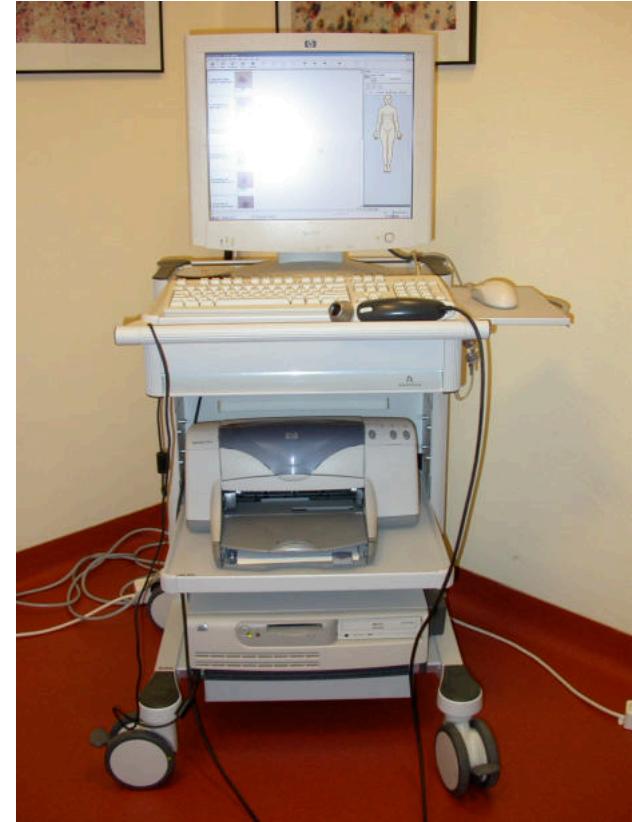
Suivi clinique





Dermogenius

Dermoscopie Digitalisée Séquentielle DDS



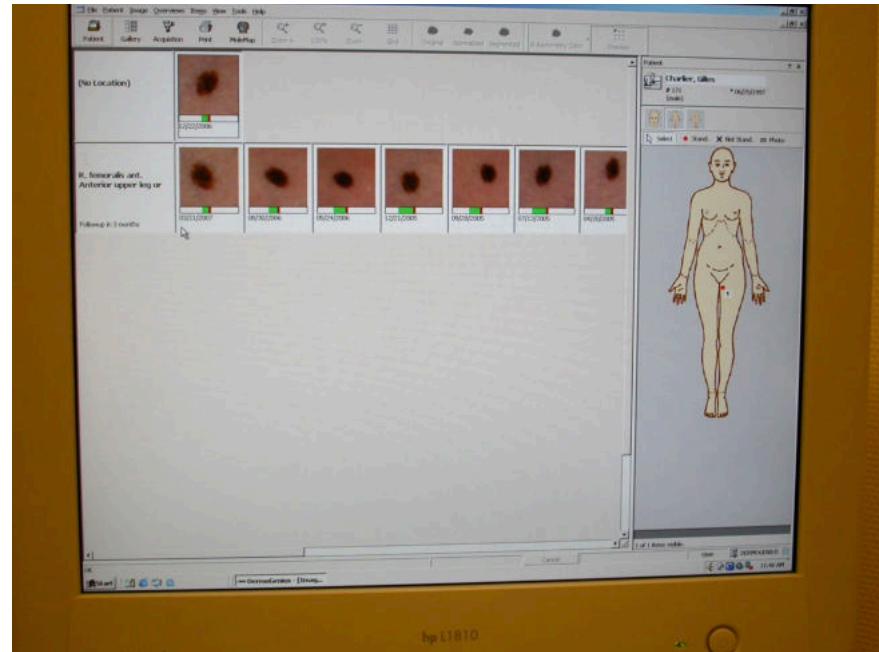
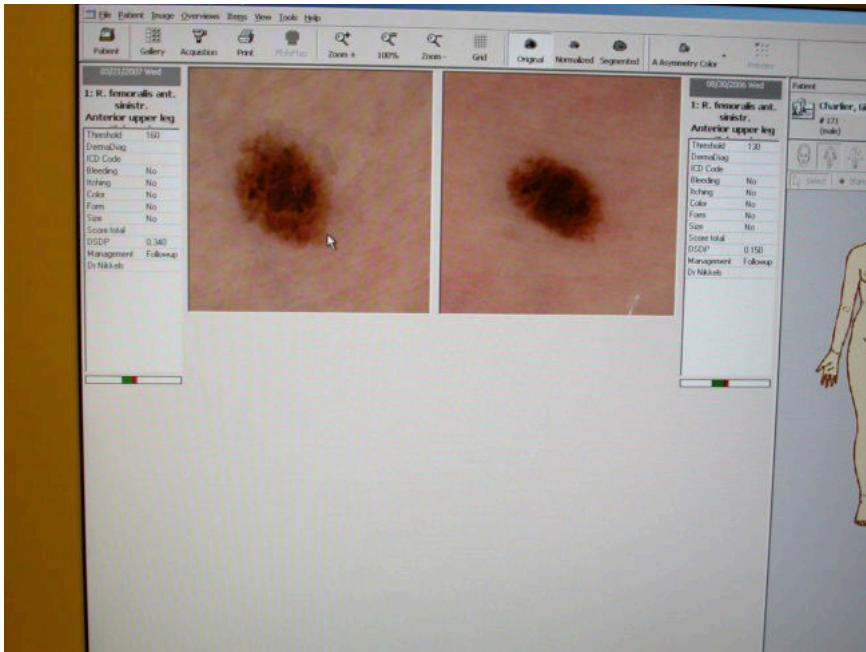


Dermogenius



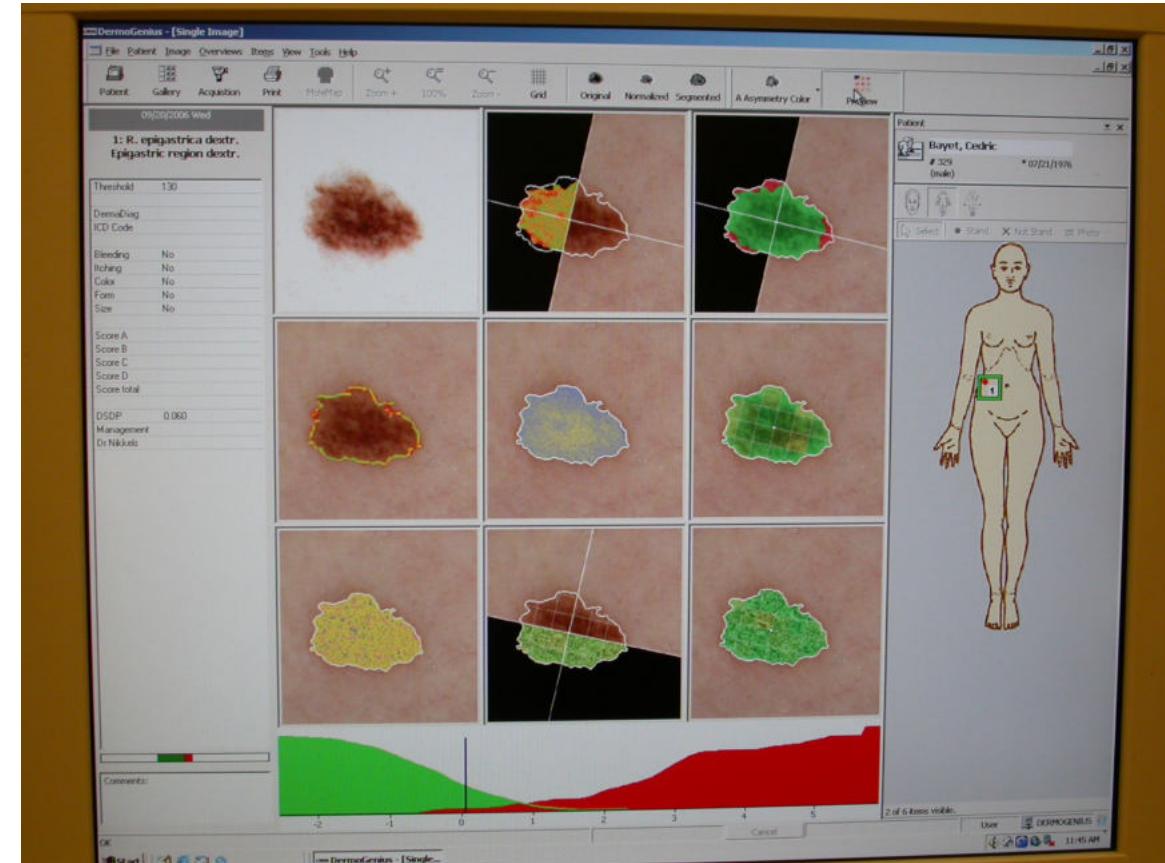
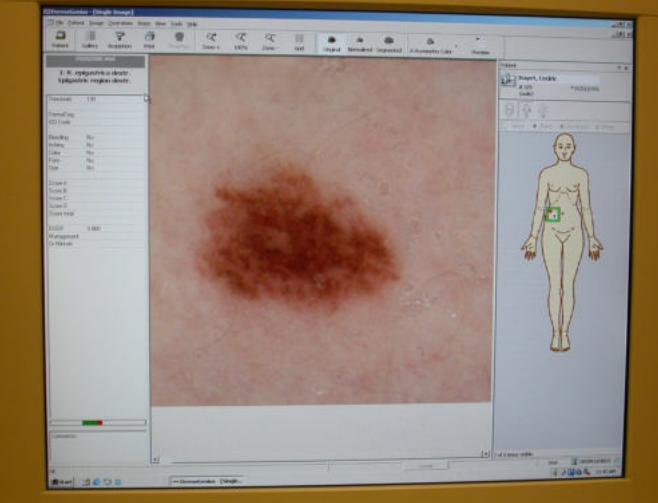
Department of Dermatology

Centre Hospitalier Universitaire de Liège





Dermogenius





Whole body photography



Trajet de soins en dermat-oncologie

Examens lors d'une visite de suivi mélanome

1

1. Anamnèse (apparition nouvelles lésions, signes généraux, etc...)
2. Examen dermatologique (Total body examination)
3. Examen dermatoscopique
4. Examen DDS (Dermoscopie digitalisée séquentielle)
5. Examen clinique:
 - Adenopathies éventuelles
 - Hépatomégalie
 - Splénomégalie

2

Examen et prise en charge dermatologique (recherche des effets indésirables des traitements oncologiques chez le patient métastatique)

3

Rappel de la photoprotection

4

Rappel de la vitamine D

5

Programmer nouveau RV de suivi



Melanoma

Survival according to disease stage

LE MÉLANOME : LE TRAJET DE SOINS DU PATIENT

DU DIAGNOSTIC AU TRAITEMENT

ABSI G (1), DAMSIN T (1), LEVAS E (1), LIBON F (1), SOMJA J (2), COLLINS P (2), REGINSTER MA (2), QUATRESOOZ P (2), RORIVE A (3), MARCHAL N (3), JACQUEMIN D (4), BOUS A (4), PIRET P (5), NIKKELS AF (1)

Résumé : Le traitement du mélanome est un exemple typique de collaboration multidisciplinaire, afin de pouvoir garantir au patient une prise en charge rapide dès le moment de la détection de la lésion. Tant au niveau dermatologique, anatomopathologique et oncologique, d'énormes progrès ont eu lieu ces dernières années. Ils permettent un accès au diagnostic de plus en rapide par la téledermoscopie, une précision diagnostique accrue par la dermatopathologie, l'ultrason à haute fréquence et la tomographie par cohérence optique, une détermination des facteurs de risque immunohistochimiques et génétiques sur les analyses anatomopathologiques ainsi que le recours à des immunothérapies, notamment les anti-PD1, et à des traitements ciblés. Ces nouveaux traitements permettent souvent une plus longue survie sans progression que les protocoles classiques dans certains cas de mélanomes métastatiques. Cet article reprend le trajet de soins du patient, du diagnostic initial et du staging au traitement éventuel avec son survi.

MOTS-CLÉS : Mélanome - Dermoscopie - Télédermatologie - Téledermoscopie - Diagnostic - Traitement - Traitements ciblés - Immunothérapies

INTRODUCTION

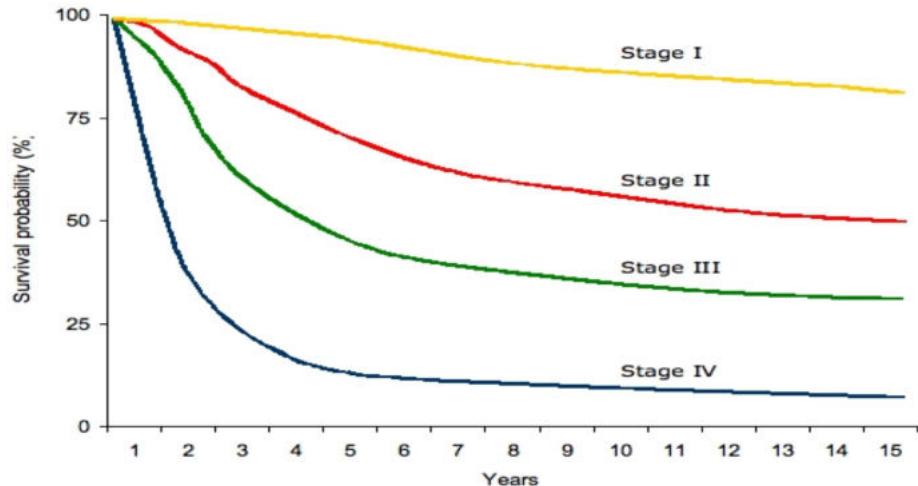
Le mélanome est une prolifération maligne des mélanocytes (1). Environ la moitié des mélanomes se développent au sein d'un naevus pigmentaire préexistant, et l'autre moitié sur une peau saine. Le site le plus fréquent est le dos

MELANOMA : THE PATIENT'S CARE PATHWAY. FROM DIAGNOSIS TO THERAPY

SUMMARY : The management of melanoma is a typical example of a pluridisciplinary approach aiming to provide the patient with a rapid and adequate treatment plan after the initial diagnosis. Both in the domains of dermatology, pathology and oncology, enormous progress has been made. Recent advances permit a rapid access to diagnostic techniques using teledermoscopy, an improved diagnostic accuracy using dermoscopy, pre-interventional high-frequency ultrasound and optical coherence tomography, the determination of immunohistochemical and genetic analyses on the pathology samples. Furthermore, the development of immunotherapies, in particular the anti-PD1 antibodies, and the directed therapies, therapies permitting an increased number of patients to experience an increased survival with an acceptable tolerance profile in the event of metastatic lesions. This article describes the patient's care pathway, from the initial diagnosis, staging, to an eventual treatment and follow-up.

KEYWORDS : Melanoma - Dermoscopy - Teleradiology - Teledermoscopy - Diagnosis - Treatment - Targeted therapies - Immunotherapies

campagne Euromelanoma (4, 5). Une éducation à la photoprotection externe, vestimentaire et comportementale est également fournie chaque année par la presse écrite, télévisée et sur les réseaux sociaux.





Melanoma

Prevention

Slip



Slop



Slap



Seek



Slide



Protect yourself in five ways from skin cancer



Melanoma

Prevention



EVERY DAY
USE SUNSCREEN



EVERY MONTH
CHECK YOUR SKIN



EVERY YEAR
VISIT A DERMATOLOGIST
IF YOU'RE AT HIGHER RISK
WWW.CUREMELANOMA.ORG



Melanoma

Prevention





Melanoma

Prevention



No, sunscreen use does not impair vitamin D production

No, you cannot use the cream from last year

The real-life SPF is about 1/3 of the figure on the package

One single texture is OK for the entire family



Melanoma

Prevention

SPF 50 vs SPF 100 ?

SPF 100/50: 50: 55% more sunburns

ERYTHEMA SCORE:
50: 0,75
100: 0,52

Impact on MM incidence: still ?????



Melanoma

Prevention

UV-index

Smart phone apps
Weather forecast

0-4: nothing to worry
5-8: protection
9-11: stay inside

UV PROTECTION CHART

Low (0-2)	Medium (3-5)	High (6-7)	Very High (7-10)	Extremely High (11+)
Sunscreen	Sunscreen	Sunscreen	Sunscreen	Sunscreen
Sunglasses	Sunglasses	Sunglasses	Sunglasses	Sunglasses
	Hat	Hat	Hat	Hat
		Shade	Shade	Shade
				Staying indoors between 10am-4pm

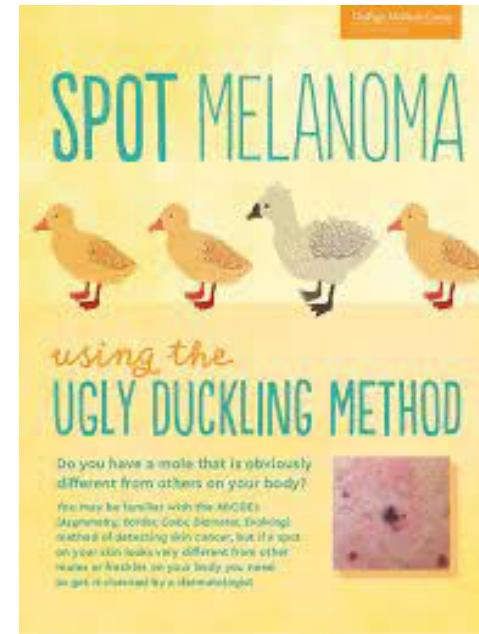


Melanoma

Prevention

Take a shower with your partner

Shadow



Children

Beta-HPV infection during childhood increases keratinocytic photosensitivity

Melanoma ??????



Melanoma

Prevention

Information and awareness campaigns

Only 5% of all melanoma patients arrive at the oncology department

We are fighting to get this figure even lower





Melanoma

Thank you for your attention



LE MÉLANOME : LE TRAJET DE SOINS DU PATIENT

DU DIAGNOSTIC AU TRAITEMENT

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Résumé : Le traitement du mélanome est un exemple type de collaboration multidisciplinaire, afin de pouvoir garantir au patient une prise en charge rapide dès le moment de la détection de la lésion. Tant au niveau dermatologique, anatomopathologique et oncologique, d'énormes progrès ont eu lieu ces dernières années. Ils ont permis un accès au diagnostic plus en rapide par la téledermoscopie, une précision diagnostique accrue par la dermoscopie, l'ultrason à haute fréquence et la tomographie par cohérence optique, une détermination des facteurs de risque immunohistochimiques et génétiques sur les analyses anatomo-pathologiques ainsi qu'un recours à des immunothérapies, notamment les anti-PD1, et à des traitements ciblés. Ces nouveaux développements permettent d'espérer une plus longue survie du patient, avec un profil de tolérance acceptable en cas de lésions métastatiques. Cet article reprend le trajet de soins du patient, du diagnostic initial et du staging au traitement éventuel avec son suivi.

MOTS-CLÉS : Mélanome - Dermoscopie - Téledermatologie - Téledermoscopie - Diagnostic - Traitement - Traitements ciblés - Immunothérapies

INTRODUCTION

Le mélanome est une prolifération maligne des mélanocytes (1). Environ la moitié des mélanomes se développent au sein d'un naevus pigmentaire préexistant, et l'autre moitié sur une peau saine. Le site le plus fréquent est le dos

MELANOMA : THE PATIENT'S CARE PATHWAY. FROM DIAGNOSIS TO THERAPY

SUMMARY : The management of melanoma is a typical example of a pluridisciplinary approach, in order to provide the patient with a rapid and adequate treatment plan after the initial diagnosis. Both in the domains of dermatology, pathology and oncology, enormous progress has been made. Recent advances permit a rapid access to diagnostic techniques using teledermoscopy, an improved diagnostic accuracy by dermoscopy, high-frequency ultrasound and optical coherence tomography, a determination of risk factors using immunohistochemistry and genetic analyses on the pathology samples. Furthermore, the development of immunotherapies, in particular the anti-PD1 antibodies, and the directed therapies, therapies permitting an increased number of patients to experience an increased survival with an acceptable tolerance profile in the event of metastatic lesions. This article describes the patient's care pathway, from the initial diagnosis, staging to an eventual treatment and follow-up.

KEYWORDS : Melanoma - Dermoscopy - Teledermatology - Teledermoscopy - Diagnosis - Treatment - Targeted therapies - Immunotherapies

campagne Euromelanoma (4, 5). Une éducation à la photoprotection externe, vestimentaire et comportementale est également fournie chaque année par la presse écrite, télévisée et sur les réseaux sociaux.