General Practice Liaison Officer Program

SKIN DEEP – Multidisciplinary Skin Cancer Workshop

Saturday 9 November 2024 Clinical Skills Development Service





An Australian Government Initiative

Metro North Health

Acknowledgement

Metro North Hospital and Health Service and Brisbane North PHN respectfully acknowledge the Traditional Owners of the land on which our services and events are located. We pay our respects to all Elders past, present and future and acknowledge Aboriginal and Torres Strait Islander people across the State.

The plan for today...

555

8.30am

10.10am

10.35am

Welcome address

First session

- A diagnostic approach to skin lesions ٠
- NMSC identification and management •

Morning tea

Second session

- Clinical considerations for referrals •
- Multidisciplinary management of complex patients ٠
- Multidisciplinary panel •

Lunch

1.30pm

3.10pm

3.50pm

12.45pm

Interactive skills stations/case studies

- Biopsy considerations and suturing challenges ٠
- Wound management and aftercare in General Practice •
- Non-surgical treatment of NMSC ٠
- Medical Oncology case study •

Last session

Update on melanoma and immunotherapy

Closing address & evaluations

A diagnostic approach to skin lesions

Adj. A/Prof David B. Francis Clinical Lead, Dermatology | RBWH





An Australian Government Initiative

Metro North Health

Skin Cancer

ADJ A/PROF DAVID B FRANCIS

MBBS FACD FACMS GAICD

CLINICAL LEAD RBWH DERMATOLOGY

VMO PAH



The Royal Brisbane and Women's Hospital and Metro North Health acknowledges the Turrbal and **Jagera Traditional Custodians of the** land upon which we live, work and walk, and pay our respects to Elders past, present and emerging.

Metro North Health



Why do we care ?

 1455 deaths Melanoma 2021 (958 males, 497 females)
 \$1.7 billion per year treating skin cancer (about 2 hospitals)

- 223,000 BCC per year
- ▶ 128,500 SCC per year
- 4180 Qld melanoma per year
- Skin cancer 1 in 3 by 50 years age, 2 in 3 by 70 years
- 840,000 GP skin check encounters per year
- Apart from pain, suffering, loss productivity etc.

BCC



BCC

- Nodular Ulcerative (Rodent Ulcer)
- Multifocal Superficial
- ► Infiltrating
- Micronodular
- Morphoeic, Fibrosing
- Pigmented
- BCC w Squamous differentiation
- BCC with PNI
- Crust vs Scale

Nodulo ulcerative



Superficial BCC



Multifocal Superficial





Morphoeic BCC

Infiltrating



Infiltrating











Micronodular/mixed















Multifocal etc

BCC with PNI







BCC

IEC:

Intrepidermal Squamous cell Carcinoma

- ► In situ SCC
- Bowen's Disease

IEC





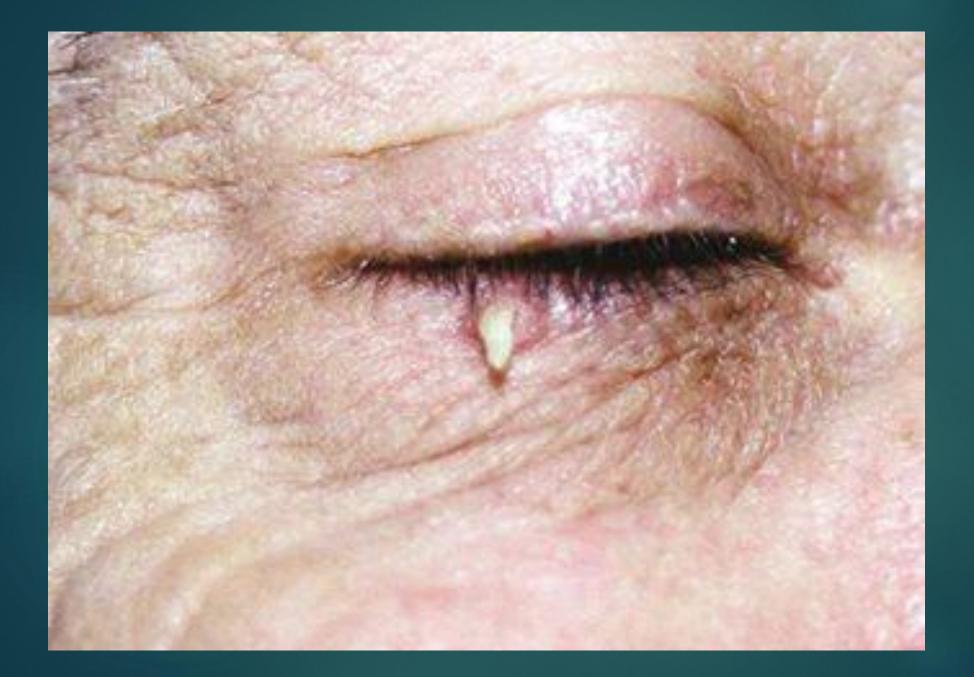






Squamous Cell Carcinoma : SCC







SCC vs Keratoacanthoma



SCC





SCC with PNI



31/01/2022



Melanoma

- ► In Situ, Level !,
- Hutchinson,s Melanotic Freckle (HMF) =
- Lentigo Maligna
- Lentigo Maligna Melanoma
- Superficial Spreading Melanoma
- Nodular melanoma
- Desmoplastic melanoma
- Neurotropic Melanoma
- Acral Lentiginous melanoma
- Subungual Melanoma
- ► Amelanotic





Acral Lentiginous Melanoma





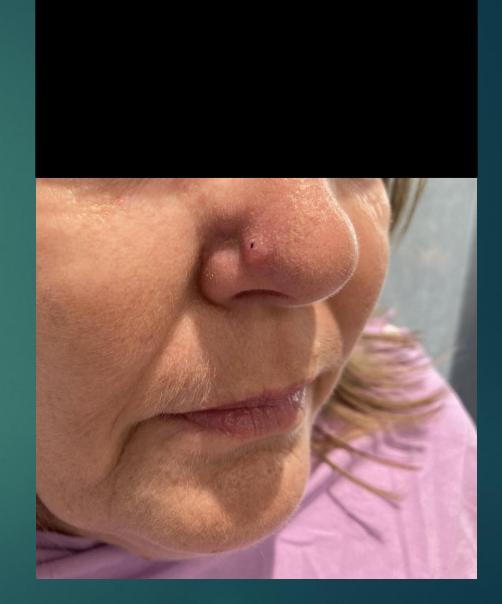


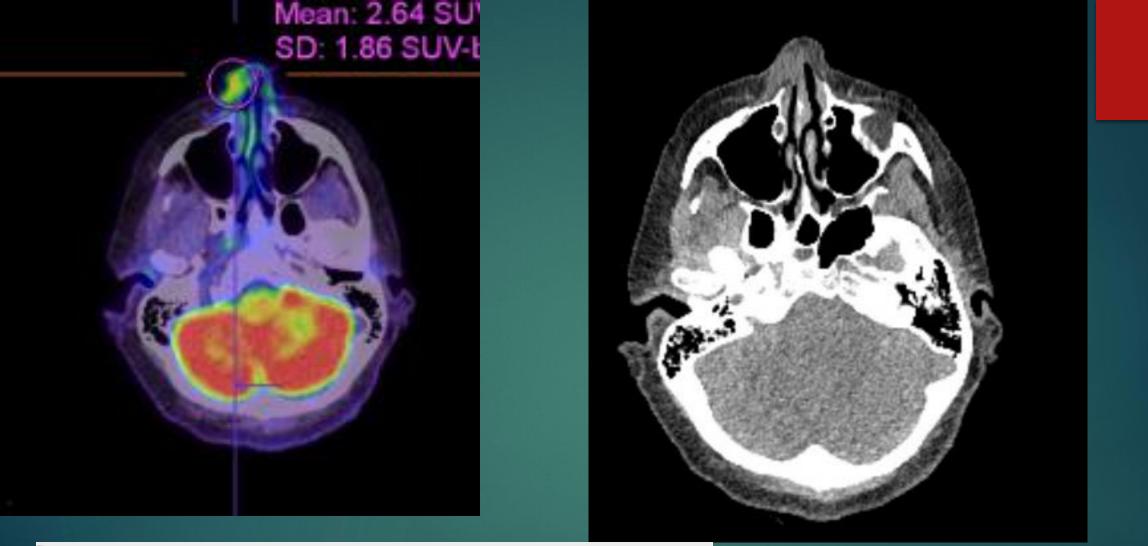












Comment:

 Intense FDG avidity of the right nasal alar biopsy confirmed melanoma. FDG avidity of the bilateral nasolabial folds and nasal septum is likely physiological, although cannot be discerned from the metabolically active right nasal alar lesion. Local disease extent is better characterised on MR face 03/10/2024.

2. No FDG avid nodal or metastatic disease.

 Intense FDG avidity of the larynx and upper cervical oesophagus is nonspecific and may be inflammatory/infective. Clinical correlation is suggested. Direct endoscopy may be indicated.

MICROSCOPIC

R nose: The punch biopsy of skin incorporates epidermis to deep reticular dermis. Almost the entire dermis is infiltrated by a poorly differentiated malignancy. The tumour is composed of pleomorphic cells with enlarged, hyperchromatic nuclei with occasional distinct nucleoli. Binucleate and multinucleate tumour cells are present. Two (2) atypical mitoses per square millimetre is seen.

The lesional cells show strong diffuse positivity for SOX10. They show a variable pattern of staining for S100 protein; with strong nuclear positivity in the papillary dermis nests, and weak patchy granular cytoplasmic reactivit in the deep dermal area. There is focal and patchy 1+ staining with PRAME. SMA, CD10 and CK34 are negative.

INI-1 and H3K27me3 are retained in the nuclei with a normal pattern. No surface ulceration, regression, neurotropism, desmoplasia, perineural invasion, lymphovascular space permeation, tumour infiltrating lymphocytes, associated benign melanocytic lesion or microsatellite nodule can be identified.

The lesion is transected at the peripheral and deep aspects of the biopsy. BRAF immunohistochemistry is being obtained and a supplementary report will follow.

Histo

SOFFARI

R nose: Dermal melanoma (at least 1.5mm thick); BRAF immunohistochemistry pending.

Metastatic breast cancer



Metastatic Skin Cancer..after Rx



Referral

Pre biopsy or excision photo...also post photo if flap etc

- ► Good light , IN FOCUS
- ▶ 5cm (What is it),
- 10 cm, 30 cm (Where is it)..and send most helpful ?all
- Diagram if need
- History..speed of growth etc...timeline?
- What do you want?..lesion for treatment, solar damage for assessment etc
- ► Why refer...eg you have already done etc etc
- Helpful info..eg immunosuppressed, on blood thinner, immobile, lives alone etc etc

Key points

Very common

- Be alert..history may mislead
- Good light and magnification (Dermatoscope)
- ABCDE (Assymetry, Border, Colour variation, Diameter (6mm), <u>Evolving</u>...size/ shape /colour)
- Suspect until proven otherwise : Refer/Biopsy..does it look like it shouldn't be there!
- PLS take photo pre biopsy..5,10, 30 cm..good light, check in focus!
- Lymph nodes

Nonsurgical Management of NMSC

Adj A/Prof David Francis

MBBS FACD FACMS GAICD Accredited in Mohs Micrographic Surgery

Clinical Lead Dept of Dermatology RBWH



Acknowledging:

Nonsurgical Treatments

- ▶ Topical 5Fluorouracil
- (Salicylic Acid, Chemo wraps, Calcipotriol, topical retinoid)
- Topical Imiquimod
- Photodynamic therapy
- Cryotherapy? Curette and cautery
- Radiotherapy
- Systemic drugs immunotherapy, retinoids, vitamin B3, interferon
- Laser Therapy
- etc

Suitable for nonsurgical treatment

- Solar keratosis
- IEC (Bowens disease, SCC in situ)
- Superficial BCC
- Some nodular BCC
- Mixed presentation/pre-operative cleanup when delay in surgical treatment is acceptable
- Avoidance of surgical morbidity
- No inappropriate delay in inevitable surgical treatment eg. sclerotic BCC
- (? HMF..Imiquimod)

Patient Assessment

Age

- Co-morbidity/general health/immunosuppression
- Anatomical site
- Histology
- Clinicopathological correlation
- Consideration of down-side of non-surgical treatment against down-side of surgical treatment
- Treatment failure of non-surgical treatments..What are outcomes of recurrence
- Likely simple or complicated surgical plan
- Unnecessary morbidity associated with surgical treatment
- How much trouble can I get myself into if it doesn't work!

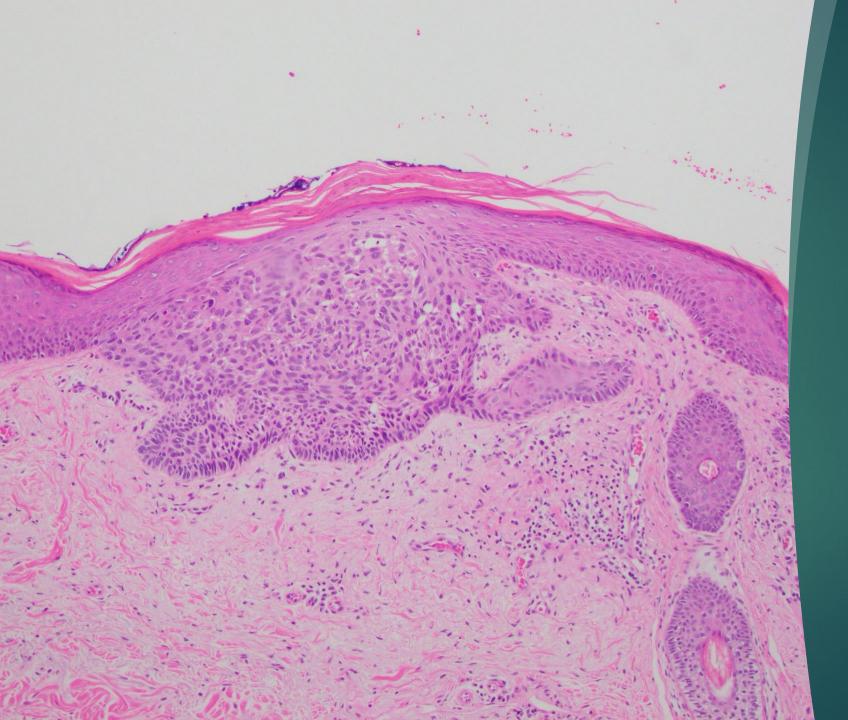
Solar Keratosis

- Approximately 1 in 100 per year become SCC
- Not a skin cancer but frequently associated with BCC/SCC/IEC
- ?masking more important lesions
- A few or Field issue...

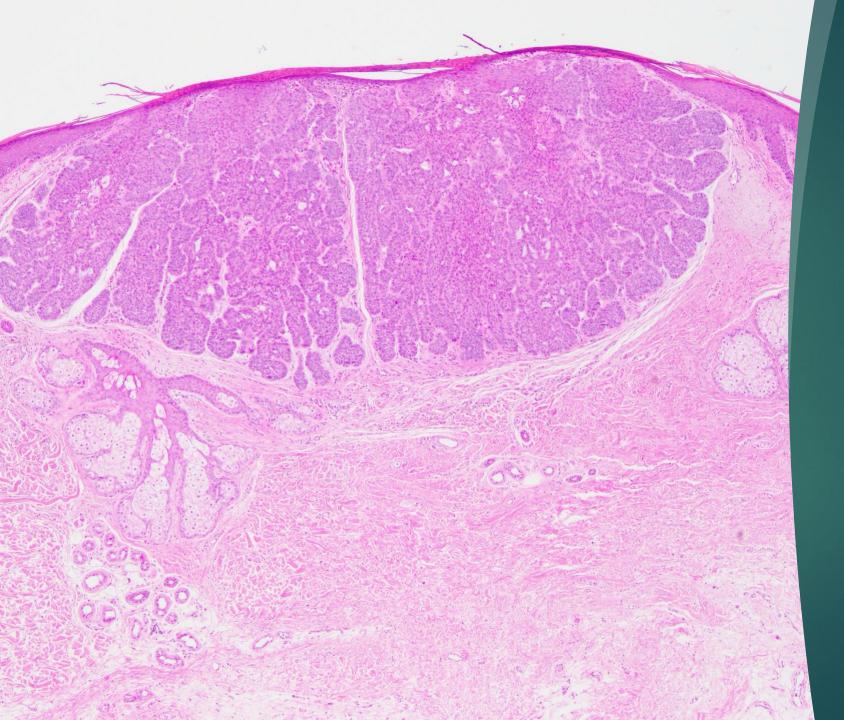
Basal Cell Carcinoma

▶ 70% of skin tumours

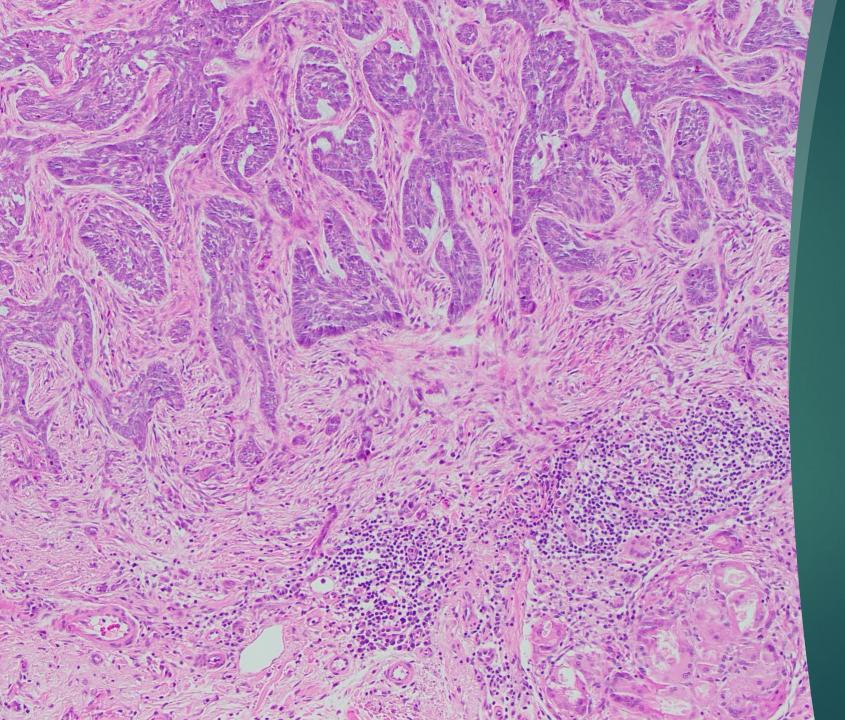
- Approximately half of all Australians will develop a BCC before the age of 70(?1 in three by age 50)
- Superficial, nodular, cystic, micro-nodular, pigmented, adenoid, infiltrating, sclerosing, keratotic, infundibulocystic, basisquamous, appendageal variants
- Rarely metastatic (0.05%)
- Epidermal/superficial dermal involvement = clinical changes
- Excision margin involvement likely recurrence
- Remember eyelids, behind ears, scalp etc



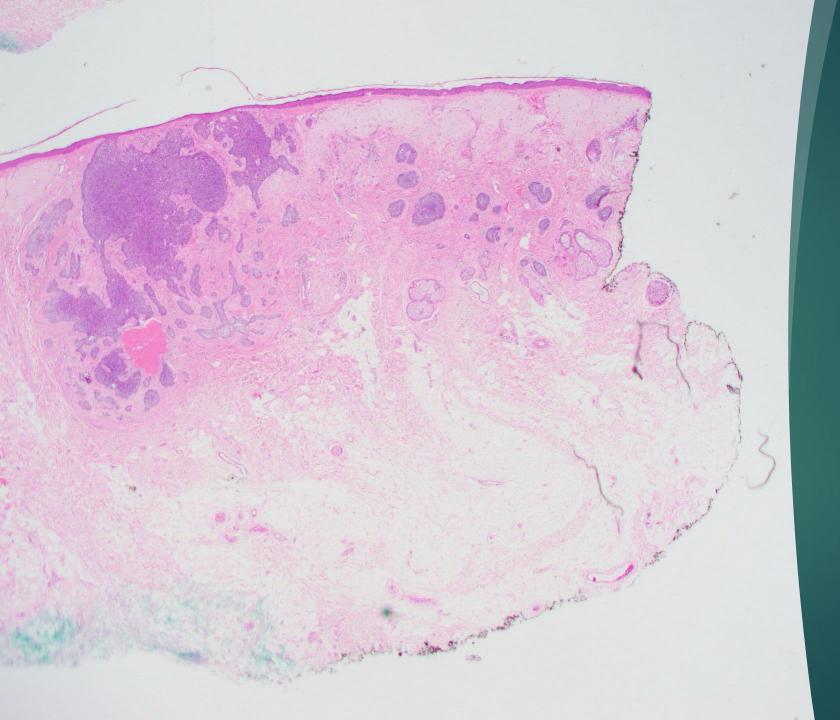
Superficial BCC



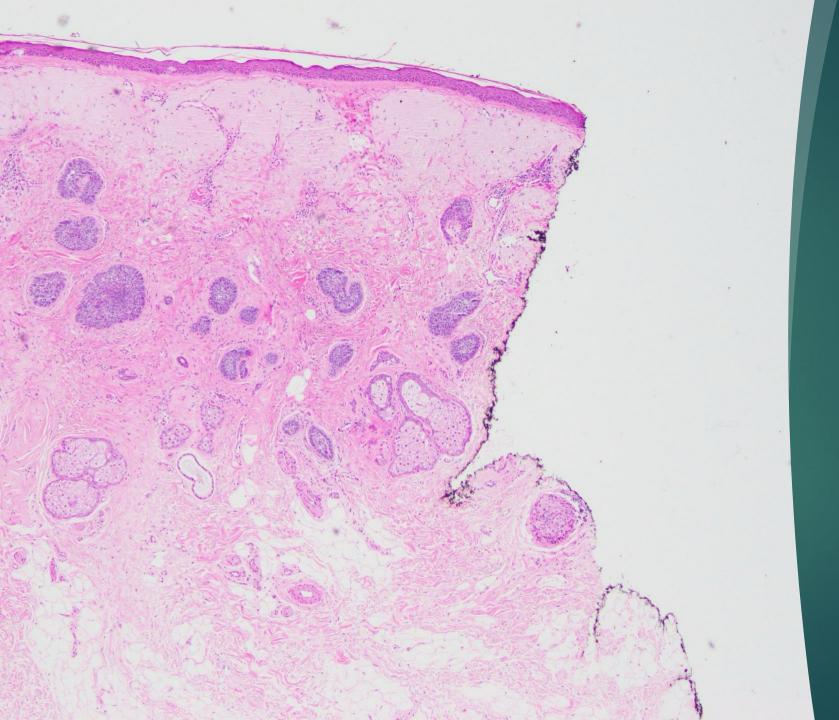
Nodular BCC



Infiltrative BCC



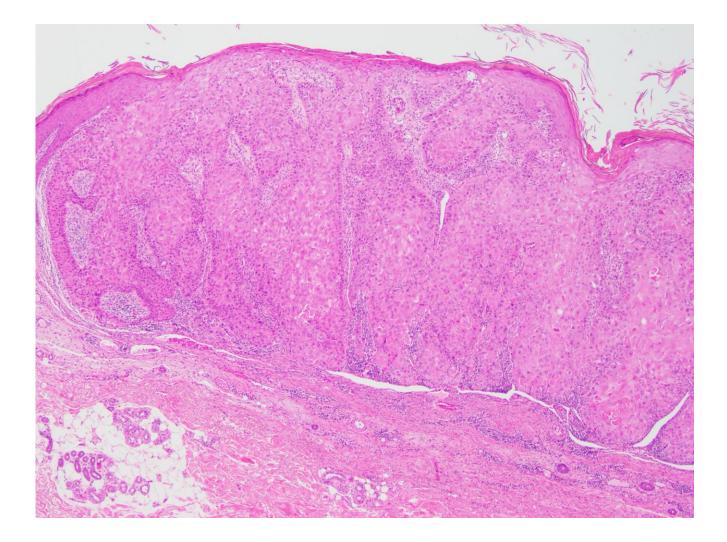
Solid and micronodular BCC



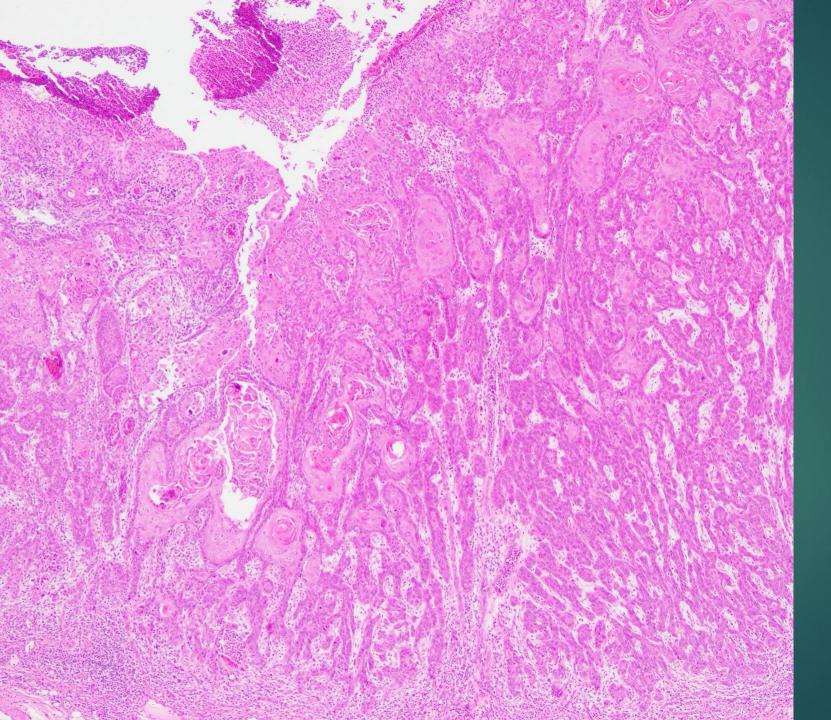
Margin involvement

Squamous Cell Carcinoma

- Second most common form of skin cancer in white people
- UVR, carcinogens (eg. Arsenic), HPV, radiation, burns/scarring
- Australia has highest incidence approximately 170 per 100,000/population
- Metastatic rate varies between 0.5% and 16%; higher rates in poor differentiation, thickness of the primary lesion (>4-5mm), PNI, arising in Bowen's Disease, head and neck, (especially lip), non-sun exposed areas, incomplete excision, immunosuppression – transplant patients
- In-situ (Bowen's, IEC)
- Well-differentiated
- Poorly-differentiated
- Keratocanthoma



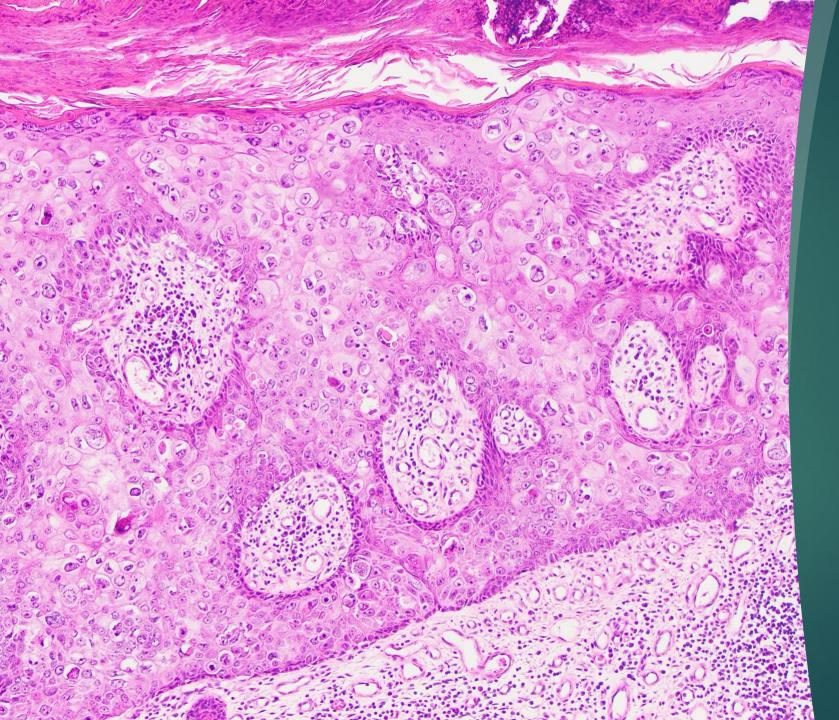
Well differentiated SCC



Moderately differentiated SCC

Intraepidermal Carcinoma (Bowen's, SCC in Situ)

- Superficial, accessible
- Follicular involvement (proven or likely/hairbearing site)...adnexal
- Clinical presentation flat, thick, indurated, hyperkeratotic
- Progress to invasive SCC, around 3-5% incidence
- ▶ If invasive disease develops, metastasis 2-5%



IEC

Topical 5FU 80% Effective

- Pyrimidine analogue of uracil that is incorporated into RNA and disrupts RNA synthesis.
- Binds within a cell to thymidylate synthetase.
- Adjuvant agents cryotherapy, trichloroacetic acid (25-45%), salicylic acid (10%), calcipotriol (Daivonex 0.005%)
- Regimen varies..twice daily, thin smear, avoid grooves, not at bedtime or if going to smear into eyes, not pregnancy...check weekly, stop if not coping and call...its not a test of your character!..its a medical treatment not a torture treatment!..handout and followup...NOT a set and forget treatment
- ▶ "healing ointment"..for a week ...Hydrocortisone or stronger
- Variable response dependent on site eg. face responds more quickly than dorsal hands
- ▶ Thickness of lesion and hyperkeratosis important
- ▶ Inflammatory effects begin day 5-10
- Inflammation lasts 4-6 weeks
- Photosensitivity
- ► Allergy < 1% (5FU and vehicle)
- Secondary infection (staph, HSV)
- Can get textural change and persistent erythema

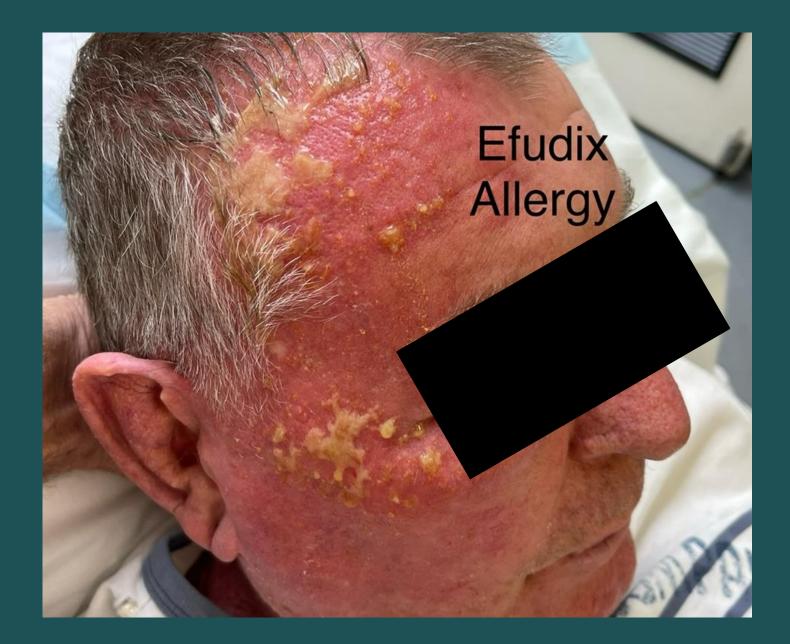








IEC :5FU -10 days



Topical Imiquimod (5% Aldara)

- Anti-tumour and anti-viral (first used for HPV)
- Acts on the innate and the adaptive immune responses both directly and indirectly.
- Direct action by binding toll-like receptors (TLR) on white blood cells and by inducing apoptosis
- Indirect action by inducing cytokines (IL-12, TNF α , INF γ) which increase cytotoxic T-cells and NK cells
- May block hedgehog signaling pathway which plays a role in BCC development
- ▶ Solar keratoses up to 25cm² 2-3 times weekly, up to 16 weeks
- ► In-situ SCC daily, up to 16 weeks
- ► Superficial BCC 5 days/week for 6-12 weeks
- Case reports of clearance of infiltrative and sclerotic BCCs isolated cases and follow up is poor..more work required
- New and potent Imiquimod analogue Resimiquimod

Photodynamic Therapy

(PDT)

Methylaminolevulinate (MAL) (Metvix 16%)

5-aminolevulinic acid (ALA) (Compounded)

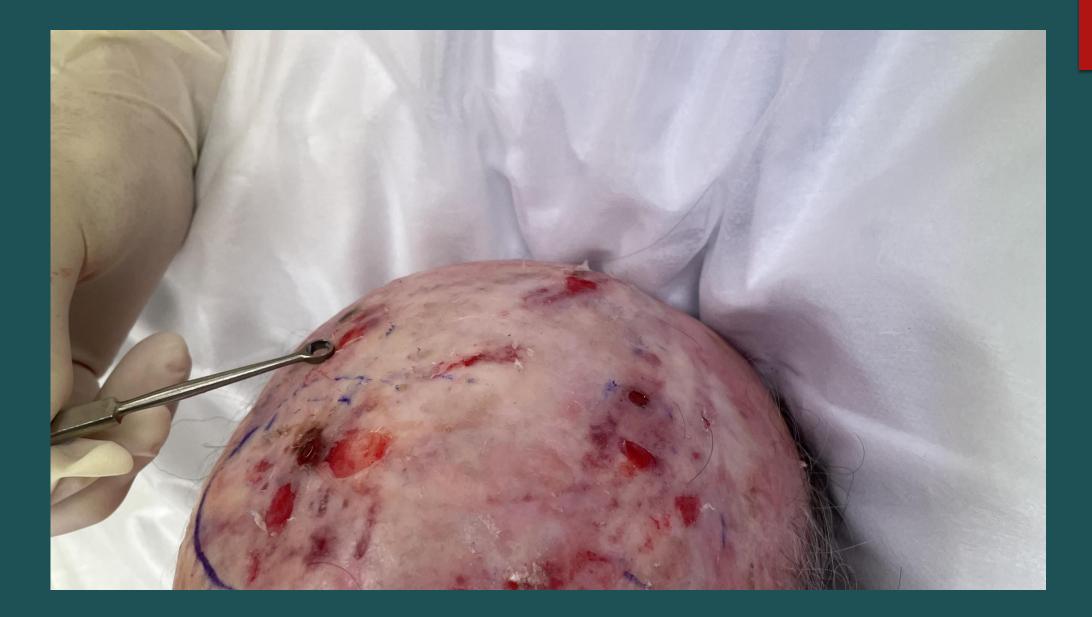
- Dermatologists have been treating a photosensitive condition called porphyria for a long time, protophorphyrin IX (PpIX) responsible
- PDT takes advantage of the photosensitive properties of PpIX
- ALA is a pre-cursor in the haem synthesis pathway and is the rate-limiting step when ALA is administered, it is taken up in cells and metabolized into PpIX
- Accumulation of PpIX in tumour cells may occur because of increased activity of a deaminase which increases conversion of ALA to PpIX and a decreased activity of ferrochelatase which converts PpIX to haem
- ALA penetration of tumour cells is also enhanced if the tumour has poorly formed stratum corneum or if the stratum corneum is altered by the proceduralist.

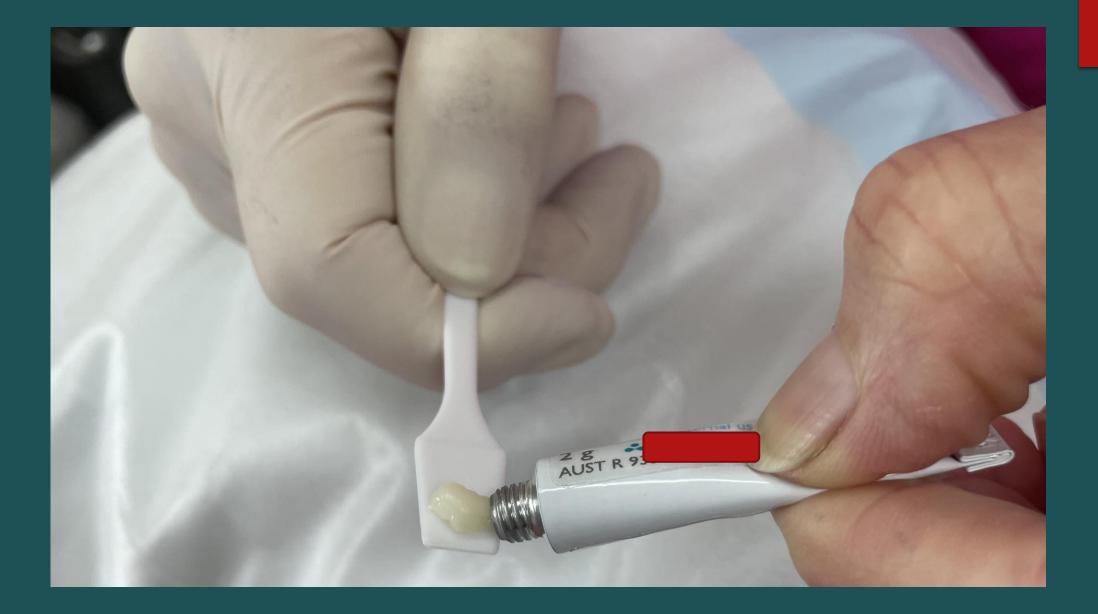
PDT Light Source

- Absorbed well by the photosensitiser corresponds to a peak of the porphyrin excitation spectrum in tissues
- PpIX excitation peaks 410nm (Violet) and 635nm (Red)
- Lights available emits heat-free visible red light at a peak wavelength of 630nm or close to
- Penetrate a desirable depth 630nm light penetrates up to 5mm

PDT Technique

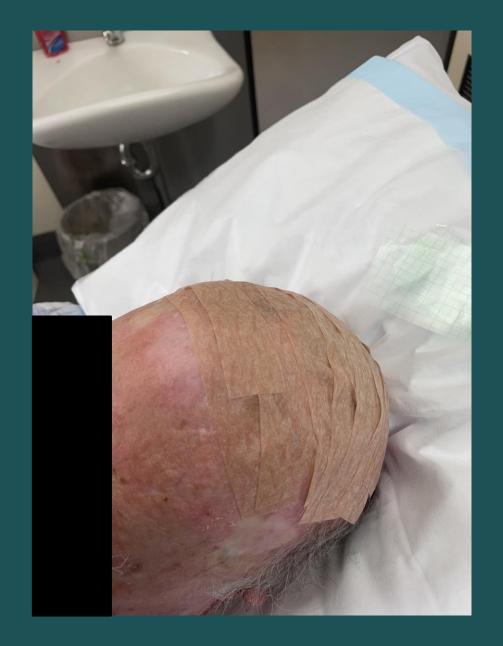
- Lesion prepared gentle curettage, sandpaper, peeling agent
- Local anaesthetic usually required (painful procedure)...in some pain seems disproportionate
- ALA cream applied under opaque occlusive dressing
- After 3 hours, dressing and cream removed. Local anaesthetic repeated
- Light source (630nm) red light approximately 8 minutes
- ► Emollient applied
- Sometimes oral pain relief prescribed
- Post procedure instructions are given expected healing time 7-10 days
- Daylight PDT mild to moderate solar keratoses??

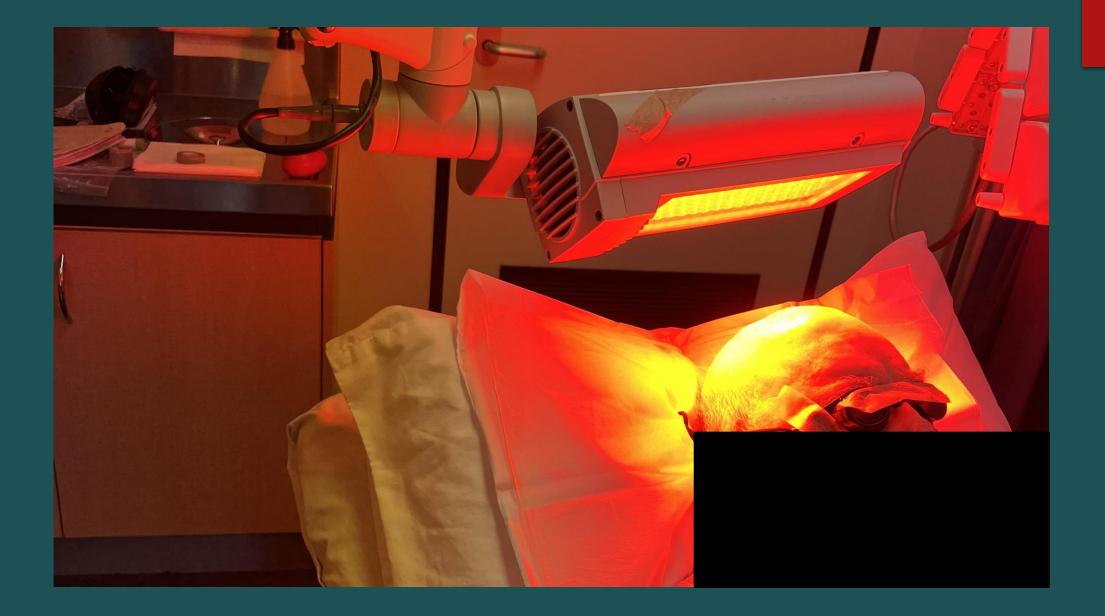












Practical Consent Issues - Specific

- INFORMATION about PDT and what is going to happen including Anaesthesia
- Pre care and after care
- Side effects : usual and less usual
- Outcomes : likely and less likely
- Comparison and cost
- Recurrence and consequences of them
- ► Follow-up...emphasize
- Consent form..in writing and info
- Don't guild the lily
- cure rate not 100%



PDT Complications

Pain

- esp large areas
- group of patients 'disproportional' pain..be aware!
- Pustular inflammation
 - ► Face..upper lip and nose
 - Culture negative
 - Within 48 hrs, settles over 48 hours
- Swelling (esp periorbital) , erythema, erosion lower legs

Recurrence – further treatment including surgery may be needed













Any Questions?

Clinical considerations for referrals

Dr Andrew Lewandowski Plastic & Reconstructive Surgeon | RBWH

Dr Srishti Dutta GPLO





Metro North Health

General Practice Liaison Officer Program

Events 🛛 🖀 Resize font

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Refer your patient

Refer your patient

Information for GPs and health professionals to help refer patients and find services available at Metro North Health.

Latest updates

Multilingual translated videos are now available for Gastroenterology patients explaining about colonoscopy & endoscopy procedures for the following locations:

- RBWH Gastroenterology and Hepatology Caboolture Hospital Gastroenterology
- STARS Gastroenterology & Endoscopy Services
- Redcliffe Hospital Gastroenterology & Hepatology

Rapid Access Services

Rapid Access Clinics and Services - Local GPs can refer patients requiring escalation of care to these services for urgent assessment and treatment within a few days to provide an alternative to an emergency presentation.

Specialist outpatient services

Specialist outpatient referrals are coordinated through the Metro North Health Central Patient Intake Unit for hospitals in the region.

Plastic and Reconst Surgery	GP Referrals Enquiry Line: 1300 364 9	cer/Skin Lesion Go
Community Health Services	Clinical advice	Rapid Access Services ->
Select a service	Virtual Emergency Care	Voluntary Assisted Dying ->
Enquiry hotline:	Service 1300 847 833	Mental Health services ->
1300 658 252	Monday to Sunday 8am-10pm	Oral Health services
Fax: 3360 4822	Metro North Clinical	Sexual Health & HIV Service ->
	Advice Line 1800 569 099	Alcohol & Drug Service ->
	Monday to Friday 8.30am-4pm	Residential Aged Care District Assessment and Referral Service (RADAR)
	Residential Aged Care District Assessment and Referral Service (RADAR)	Behavioural Emergency — Response Team (BERT)
	1300 072 327 Monday to Sunday 8.00am - 8.00pm	Children's Health Queensland 🔿

(PDF) Central Patient Intake FAQ's (PDF)

Chronic Wounds Directory

Home / Refer your patient / Plastic and Reconstructive Surgery Plastic and Reconstructive Surgery

Conditions

Queensland Government

Metro North Health

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unless specifically stipulated in the out of scope section. Dupuytren's Contracture

- Facial Fractures
- General Plastic Surgery
- Head and Neck Mass
- Lower Limb Reconstruction

Paediatric services

Referrals for children and young people should follow the Children's Health Queensland referral guidelines.

Management

Emergency department referrals

All urgent cases must be discussed with the on call Plastic and Reconstructive Surgery Registrar. Contact through Royal Brisbane and Women's Hospital (07) 3646 8111 to obtain appropriate prioritisation and treatment.

Urgent cases accepted via phone must be accompanied with a written referral and a copy faxed immediately to the Central Patient Intake Unit: 1300 364 952.

If any of the following are present or suspected arrange immediate transfer to the emergency department.

- Hand fractures (open or closed)
- Acute fingertip injuries
- Tendon Injuries
- Airway compromise
- Uncontrolled sepsis including hand infections
- Threat to limb viability
- Acute burns
 - Lacerations and wounds not suitable for primary health management e.g. lin lacerations, large facial lacerations

Send referral

Hotline: 1300 364 938

Electronic:

About us

GP Smart Referrals (preferred) eReferral system templates Medical Objects ID: MQ40290004P HealthLink EDI: gldmnhhs

Mail:

Metro North Central Patient Intake Aspley Community Centre 776 Zillmere Road ASPLEY QLD 4034

Health pathways 😮

Access to Health Pathways is free for clinicians in Metro North Brisbane.

For login details email:

healthpathways@brisbanenorthphn. org.au

Login to Brisbane North Health Pathways:

brisbanenorth.healthpathwayscomm

unity.org

Locations

Royal Brisbane and Women's Hospital

Please note this is not an exhaustive list of all conditions for outpatient services and does not exclude consideration for referral

Refer your patient Hospitals & services Health professionals Research Get involved

- Peripheral entrapment
 - neuropathies including CTS

Reconstructive Breast Surgery

- Skin Cancer/Skin Lesion Post-Burn Reconstruction and Scar

Reconstructive Hand Surgery

Home / Refer your patient / Plastic and Reconstructive Surgery / Skin Cancer/Skin Lesion

Skin Cancer/Skin Lesion

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If any of the following are present or suspected, refer the patient to the emergency department (via ambulance if necessary) or seek emergent medical advice if in a remote region.

· Lacerations and wounds not suitable for primary health management e.g. lip lacerations, large facial lacerations, lacerations with altered sensation, large skin defects.

Does your patient wish to be referred? 🚱

Minimum referral criteria

Does your patient meet the minimum referral criteria?

Category 1

Appointment within 30 days is desirable

- skin lesion highly suspicious for melanoma or excision biopsy proven melanoma
- rapidly growing skin lesions especially on the face
- complex non-melanoma skin malignancies and any of the following:
 - ulceration and bleeding
 - rapidly enlarging
 - neurological involvement
 - lymphadenopathy
 - poorly differentiated or infiltrative tumour on biopsy
- other subcutaneous and deep tissue malignancies e.g. Merkel cell carcinoma, sarcoma
- skin lesion causing substantial obstruction to vision suspicion of malignant liposarcoma
- Confirmed SCC
- Prior malignancy at the same site

+ Other Plastic and Reconstructive Surgery conditions

Send referral

Hotline: 1300 364 938

Electronic:

GP Smart Referrals (preferred)

eReferral system templates Medical Objects ID: MQ40290004P

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unity.org

Locations

Royal Brisbane and Women's

Hospital

Category 2

Appointment within 90 days is

desirable

- Uncomplicated non melanoma skin malignancies (BCC/SCC/IEC)
- Skin lesions with any of the following:
 - o causing functional problems or significant disfigurement
 - diameter exceeds ≥ 5cm in size or rapid growth over short period of time
- significant persistent pain that is not solely pressure related
 - fixed to deep tissues, i.e. muscle or fascia
 - recurring after a previous excision
 - prone to recurrent infection
 - · diagnosis in doubt or needs confirmation

Category 3

Appointment within 365 days is desirable

- Benign soft tissue lesions e.g. lipoma, ganglion not suitable for primary health management
- Clinically significant benign lesion

If your patient does not meet the minimum referral criteria

Consider other treatment pathways or an alternative diagnosis.

If you still need to refer your patient:

- Please explain why (e.g. warning signs or symptoms, clinical modifiers, uncertain about diagnosis, etc.)
- Please note that your referral may not be accepted or may be redirected to another service

Other important information for referring practitioners

Not an exhaustive list

- Refer to HealthPathways for assessment and management information if available
- Advise patient regarding sun avoidance and appropriate use of sun screens
- Educate patient on skin cancer surveillance and arrange annual skin checks

Referral requirements

A referral may be rejected without the following information.

- Essential referral information
- Features of pigmented lesions: size, shape, colour, inflammation, oozing, change in sensation
- Biopsy results unless clinically contraindicated excision biopsy is the preferred method for suspected melanoma
- Smoking status
- History of anticoagulant therapy

Additional referral information (useful for processing the referral)

- Photograph with patient's consent, where secure image transfer, identification and storage is possible
- USS lesion result (for a suspicious lipoma)

General Practice Liaison Officer Program

😑 💥 Brisbane North



Brisbane North

Home	
COVID-19	~
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Brisbane North Localised Pathways	
Acute Services	\sim
Allied Health	~
Child and Youth Health	~
End of Life	~
nvestigations	~
Lifestyle and Preventive Care	\sim
Medical	\sim
Mental Health	\sim
Older Adults' Health	v
Pharmacology	~
Public Health	\sim
Reproductive Health	~
Specific Populations	~
Surgical	^
Cardiothoracic Surgery	~
Dentistry	\sim
ENT Head and Neck Surgery	v
General Surgery	~
Neurosurgery	\sim
Ophthalmology	~
Oral and Maxillofacial Surgery	~
Orthopaedics / Musculoskeletal	\sim
Surgery - Child	~
Plastic and Reconstructive Surgery	^
Breast Surgery	
Burn Injuries	
E	

Ear Anomalies

Excess Skin Removal Surgery

Subcutaneous Foreign Bodies

Hand and Wrist (Plastics)

Skin Cancer





() Health Alert

7 August: 4 new confirmed cases of Mpox in Queensland. Clinicians are advised to be alert and test for Mpox in patients with compatible signs and symptoms, particularly in MSM, with a low threshold for testing.

There is currently an outbreak of dengue fever in the Torres Strait and there is an ongoing risk of dengue to travellers in Indonesia. Notify your local public health unit immediately on suspicion of dengue infection (6 June 2024).

Latest News

21 October

GP News Link - 17 October

See the latest GP Link update from your PHN. Read more... [2] To receive the newsletter in your email inbox, subscribe here [2].

18 October

Notifications for internal Queensland Health referrals

GPs may receive notifications via ghRefer when their patient is referred. These are delivered by SWT to practice management software. Practices not using STS may receive a letter. Update your details [2] to receive the correspondence electronically.

18 October

Health Provider Portal (HPP) changes

The QGov system used to log in to the Health Provider Portal (HPP) will be transitioned to a new digital identity provider called Queensland Digital Identity (QDI) in early 2025. Read more ... CI

11 October

GP News Link - 10 October

See the latest GP Link update from your PHN. Read more... D To receive the newsletter in your email inbox, subscribe here D.

4 October

Pathway Updates

Updated – 18 October Osteoporosis

Updated - 11 October Lactation Support Services

Updated – 9 October Sick Day Management in Diabetes

Updated – 8 October Abnormal Vaginal Discharge

NEW - 27 September

Aboriginal and Torres Strait Islander Mental Health

VIEW MORE UPDATES ...

About HealthPathways

What is HealthPathways? >

How do I use HealthPathways? >

How do I send feedback on a pathway?

HEALTH PROVIDER PORTAL

METRO NORTH HHS

@ PHN

LOCAL RESOURCES

CLINICAL RESOURCES

A PATIENT RESOURCES

GP EDUCATION







General Practice Liaison Officer Program

\equiv 💥 Brisbane North



Brisbane North

Investigations	\sim	
Lifestyle and Preventive Care	~	
Medical	\sim	
Mental Health	\sim	
Older Adults' Health	\sim	
Pharmacology	\sim	
Public Health	\sim	
Reproductive Health	\sim	
Specific Populations	\sim	
Surgical	~	
Cardiothoracic Surgery	\sim	
Dentistry	\sim	
ENT Head and Neck Surgery	\sim	
General Surgery	\sim	
Neurosurgery	\sim	
Ophthalmology	\sim	
Oral and Maxillofacial Surgery	\sim	
Orthopaedics / Musculoskeletal	\sim	
Surgery - Child	\sim	
Plastic and Reconstructive Surgery	^	
Breast Surgery		
Burn Injuries		
Far Anomalies		

Excess Skin Removal Surgery Subcutaneous Foreign Bodies

Hand and Wrist (Plastics)

- Skin Cancer
- Pigmented Skin Lesions
- Punch Biopsy

Skin Lesion Excision

Skin Lesion Excision Presentation

Soft Tissue Lumps and Sarcoma

Plastic and Reconstructive Surgery Requests

~

- Urology
- Vascular Surgery
- Wound Care

Q Search HealthPathways

A / Surgical / Plastic and Reconstructive Surgery / Skin Cancer

Skin Cancer

Background

About skin cancer 🗸

Assessment

- any lesion with suspicious features ➤.
- EFG rule for melanoma ∨.
- ABCDE criteria for melanoma Y.

2. If patient with personal history of melanoma, check for signs or symptoms of recurrent or metastatic melanoma V.

3. Examination:

- Perform a thorough examination of the skin, including dermoscopy 2. Follow suggested protocol for conducting skin checks v and consider a chaperone.
- · Check each lesion for features of malignancy (e.g., enlarging, changing in shape, size, colour, non-healing.)
- · If patient with a lesion suspicious for malignancy, consider checking for regional lymphadenopathy (particularly for large or long-standing squamous cell carcinomas (SCCs), SCCs of the head or neck, or melanomas.)
- 4. Consider possible diagnosis based on skin lesions' features:
- Basal cell carcinoma (BCC) V
- Squamous cell carcinoma (SCC) V
- Melanoma, including amelanotic and hypomelanotic melanoma. See Dermnet NZ Dermoscopy of Malignant Melanoma [2],
- Benign or premalignant skin lesions ➤

5. Arrange investigations as required:

- If any suspicion of lymph node metastasis, request:
- CT or ultrasound of relevant lymph node basin, and
- Imaging-guided fine needle aspiration biopsy.
- If suspected brain metastasis, request MRI or CT brain.

Management

- 1. If not confident obtaining an adequate biopsy or managing skin cancer, arrange skin cancer specialist assessment (e.g., general practitioner with special interest, general surgeon, plastic surgeon, dermatologist). 2. If metastatic disease suspected, request non-acute oncology assessment and mark as urgent.
 - 3. If non-melanoma skin cancer with high-risk clinical features of a BCC or SCC 🗸 or high-risk histological features of a BCC or SCC 🗸, consider requesting skin cancer specialist assessment (e.g., GP with special interest, general surgeon, plastic surgeon, dermatologist.
 - 4. Manage other non-melanoma skin cancer according to histology:
 - Intraepidermal carcinoma (i.e., SCC in situ or Bowen's disease) ¥
 - SCC and keratocanthoma V
 - BCC ➤

Q Search HealthPathways

Skin Cancer

- Manage melanoma skin cancer according to histology:
- Melanoma in situ, including lentigo maligna V
- Invasive melanoma V
- 6. For all patients diagnosed and treated for skin cancer, provide:
- counselling regarding recurrence rates and risk v.
- 7. Arrange regular follow-up V.
- 8. If patient with chronic immunosuppression V, consider arranging skin cancer specialist assessment (e.g., general practitioner with special interest, general surgeon, plastic surgeon, dermatologist) for closer monitoring.
- 9. If any concerns, seek skin cancer specialist advice (e.g., general practitioner with special interest, general surgeon, plastic surgeon, dermatologist),

Request

- If metastatic disease suspected, request non-acute oncology assessment and mark as urgent.
- If non-melanoma skin cancer with high-risk clinical features of a BCC or SCC ∨ or high-risk histological features of a BCC or SCC V, consider requesting skin cancer specialist assessment (e.g., GP with special interest, general surgeon, plastic surgeon, dermatologist).
- If high-risk melanoma, request non-acute plastic surgery assessment (and mark as urgent), non-acute dermatology assessment (and mark as urgent), or non-acute general surgery assessment and mark as urgent for consideration of:
 - · further management.
 - sentinel lymph node (SLN) biopsy I before wider excision.
- · If treating intraepidermal carcinoma with fluorouracil:
- consider arranging punch biopsy if in doubt about possible invasive component.
- arrange surgical excision if lesion/s persist at 3 months.
- · Seek radiation oncology advice for consideration of radiotherapy if SCC, keratoacanthoma, or BCC, and surgery cannot be performed.
- · If patient with chronic immunosuppression, consider arranging skin cancer specialist assessment (e.g., general practitioner with special interest, general surgeon, plastic surgeon, dermatologist) for closer monitoring.
- · If not confident obtaining an adequate biopsy or managing skin cancer, arrange skin cancer specialist assessment (e.g., general practitioner with special interest, general surgeon, plastic surgeon, dermatologist).
- If any concerns, seek skin cancer specialist advice (e.g., general practitioner with special interest, general surgeon, plastic surgeon, dermatologist).

Information

■ For health professionals ∨

Ť For patients V

Smart Referrals

Why should I use Smart Referrals?

- 1. Allows you to <u>attach test results, imaging reports and other</u> <u>clinical documents (e.g. ECGs, photos)</u> from the patient's clinical record or your PC
- 2. Supports you to provide essential referral information
- 3. Integrated service directory identifies the speciality closest to home
- 4. Can be used for *Request for Advice (RFA)*
- 5. Shows where the referral is in the system (received, accepted, not accepted)
- 6. Shows appointment date linked to the referral
- 7. Includes an increasing number of allied health and community services



GP Smart Referrals features



- Integrated with Best Practice and Medical Director
- Aligned with state-wide referral guidelines to prompt essential referral information required for triage, decreasing the number of referrals returned requesting additional clinical information.

Brisbane North PHN Digital Health Support Officers GPSR@brisbanenorthphn.org.au

	p						
8	Smart Referrals					- 🗆	×
O.B.: ione: insion	Queensland Government	art Referrals				Dr Srishti Dutta	
bacc	Patient name: Mr Test Test DoB: 16 Oct 1959						
ations	Request information						-
	Request date	7 Nov 2024					
anding	* Request type	New referral	Update	Continuation	Request for advice		
ntive ntive ntive ntive	* Reason for referral	New condition requiring spe Deterioration in condition, r Other		outpatients < 12 months			
	* Priority	Urgent Routine					
П	* Provider	QHSR Private					
	Consents						
	* Date patient consented to request	07 Nov 2024					
D	* Patient is willing to have surgery if required?	Yes No	Not applicable]			
	* Condition and Specialty	skin cancer		HealthPathwa	i <u>vs</u> >		
	Suitable for Telehealth?	Plastic and Reconstructive Surg	lery		Plastic and Reconstructive		
	* Are you the patient's usual GP?	General Surgery Ophthalmology		Lid lesions (Ophthalmolog	ology - benign and maligna ay) (Adult)		
	Request recipient	Dermatology					-
	* Service/Location	General Medicine General Surgery					
	Specialist name	Ophthalmology					
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	Investigations and imaging						+
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	Insurance information						+
	Referring GP's information						+
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Smart Referrals		- □ >
Queensland Government	Smart Referrals	🛓 Dr Srishti Dutta
Patient name: Mr Test Test DoB: 16 Oct 1959		
E Condition specific clinical information		-
Minimum Referral Criteria		
Show 'skin lesions with other features'	Show Hide	
 Minimum referral criteria 	Skin lesion highly suspicious for melanoma or excision biopsy proven melanoma Complex non-melanoma skin malignancies Other subcutaneous and deep tissue malignancies e.g. Merkel cell carcinoma, s Poorly differentiated SCC Uncomplicated non-melanoma skin malignancies (BCC/SCC/IEC) Suspicion of malignant liposarcoma Prior malignancy at the same site Rapidly growing skin lesions especially on the face Skin lesion causing substantial obstruction to vision Skin lesion suth other features (see above) Benign soft tissue lesions e.g. lipoma, ganglion not suitable for primary health m Clinically significant benign lesions Request clinical override of minimum referral criteria	sarcoma
History and Examination Essential referral information: The smoking history recorded in the practice softy	are will automatically be included in the referral, please ensure that this is up to date	
* History	0	
* History of anticoagulant therapy	Yes No	<i>h</i>
Referral Letter		
Referral letter	0	
Pathology and Test Results Essential referral information:		
	ision biopsy is the preferred method for suspected melanoma. Please select manually.	
Click link to manually select investigations	A Go to Investigations	

V[XX] Effective: [MM/YYYY] Review: [MM/YYYY]

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e smoking history recorded in the practice software will automatically be included in the referral, please ensure that this is up to date		ndine Pathology and Test Results	
History 0		tive two • Biopsy results unless clinically contraindicated - excision biopsy is the preferred method for suspected melanoma. Please select manually.	
		the Click link to manually select investigations (¹⁴ Go to Investigations	
History of anticoagulant therapy Yes No		Click link to manually attach investigations (Pago to Attachments	
ferral Letter		Request to override essential referral information requirement Yes No	
Referral letter		Imaging and Reports Additional referral information:	
athology and Test Results		Photograph - with patient's consent USS lesion result (for suspicious lipoma)	
sential referral information:		Imaging performed Verbotograph USS	
Biopsy results unless clinically contraindicated - excision biopsy is the preferred method for suspected melanoma. Please select manually.		Other	
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Health Provider Portal

- Provides *eligible Queensland health practitioners (HPs) secure online access to their patients' Queensland Health records.
- Read-only online access allows HPs to view public hospital information including appointments, clinic letters, inpatient & ED discharge summaries, radiology & pathology reports, and medications.

* Queensland AHPRA registered GPs, nurses, midwives, optometrists, paramedics & pharmacists

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Prev Page 1 of 1	Next		Encounters			AR/Alerts	Pathology	Medical Imaging	Procedures	Care Plans
© 08-Oct to 08-Oct-2015 TNH: 2015035963 LEE, PATRICK		Event Sumn	naries	B My Health I	Record					
a <mark>l 16-Jul to 20-Jul-2011, 4 day</mark> GCH: 760000-6 DR Donald George Kardux PITCHF	<u>_</u>	•	care date	Outpatient : 12-Jan-2016	17 medie	cation(s) + 2 co	eased	The Townsville Hosp	ital	ģ
- ≝16-Jul-2011, ? TNH: 800801-1 DR ROBERTA MCFARLANE		Authorised Source Sys Authorised Medication	stem by	: 12-Jan-2016 : eLMS : Langdon, Co patient Profile						
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DR Donald George Kardux PITCHR		Spironolae Tablets	ctone (Alda	ictone) 25mg	Take 1 tablet in	the MORNING		Unchanged	Remove exce Improve hear	
01-Apr to 01-Apr-2011, 0 da PAH: 429999-1 DR MARK DONALDSON	ays ⊫⇒ 	Aspirin (A	strix) 100m	ig Tablets	Take 1 tablet in	the MORNING	with food	Unchanged	Prevent heart strokes, blood	
a18-Feb to 23-Feb-2011, 5 d	days ⇒	Esomepra Tablets	zole (Nexi	um) 40mg	Swallow whole	1 tablet once ea	ach day	Unchanged	Treat reflux di Treat/prevent	
GCH: 760000-4 DR Peter Michael DAVOREN		Ramipril - 5mg-5mg	Felodipine Tablets	(Triasyn)	Take 1 tablet in	the MORNING		Unchanged	Treat high blo Improve hear	
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IN Peter Michael DAVOREN	days ⊨⇒	Rosuvasta Tablets	atin (Cresto	or) 10mg	Take 1 tablet in	the MORNING		Unchanged	Prevent heart strokes, lower	
GCH: 760000-2 DR Peter Michael DAVOREN	<u> </u>	Venlafaxin CAPS	ie (Altven)	75mg MR	Swallow whole	1 capsule in the	MORNING	Unchanged	Improve moo	ł
02-Nov to 09-Nov-2010, 7 (GCH: 760000-1	days ⇒ ▲	Vitamin Co Tablets (C		vith Minerals	Take 2 tablets in	the MORNING	3	Unchanged	Multivitamin	
DR Peter Michael DAVOREN		Mega Cal	cium Table	ts (Cenovis)	Take 2 tablets in	the MORNING	3	Unchanged	Calcium and supplement	Vitamin D
		Magnesiu (elementa	m Forte Ta I Magnesiu	blets ım ~350	Take 1 tablet in	the MORNING		Unchanged	Magnesium S	upplement
		Paracetan MR TABS		l SR) 665mg	Swallow whole 2 Maximum of 6 p hours.				Treat pain	

Clinical Photography considerations

Key points

General considerations

- Know your obligations under the law with regard to the collection, disclosure and storage of clinical photos in the jurisdiction of your practice (refer to Section 1)
- Check your practice's policy with regard to the use of personal mobile devices for clinical photos (refer to Section 2)
- Obtain informed consent from the patient prior to collecting photos (refer to Section 3)
- · Report data breaches if they occur (refer to Section 4)

Collecting photos (refer to Section 5)

- Take care when using clinical photography apps
- Ensure the device has a high-resolution camera
- Capture only what is required
- · Remove metadata when de-identifying photos, where applicable

Storing photos (refer to Section 6)

- Use strict privacy settings on the device
- Store photos in the patient's health record
- · Always delete the photos from the device
- · Avoid third-party storage options and prevent automated back-ups
- Treat photos sent by others as if you took them yourself

Disseminating photos (refer to Section 7)

- · Take measures to transmit photos securely where possible
- · Never share photos outside of a professional context
- Be wary of social media sharing

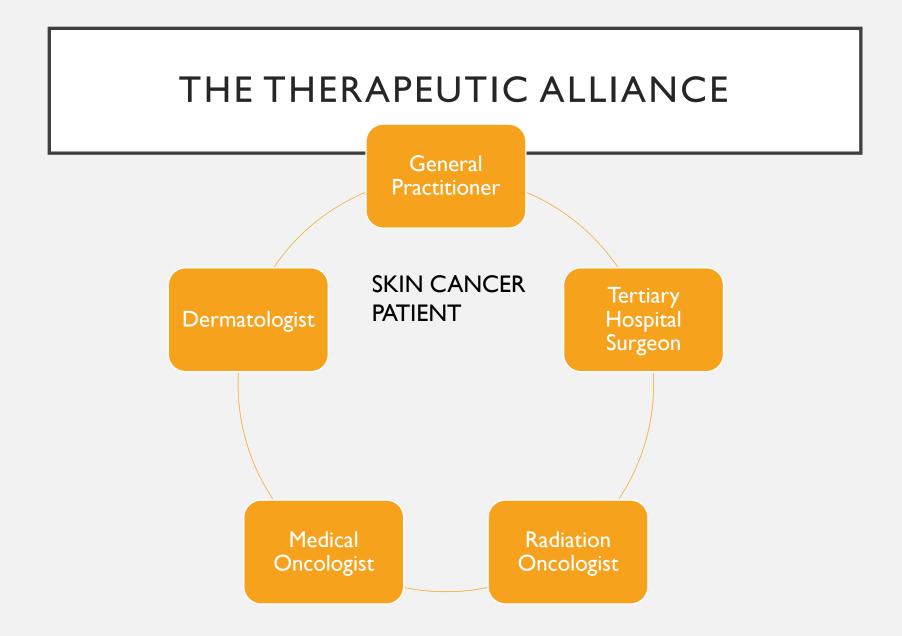
Clinical photography has become a boon to the world of medicine.¹ Table 1 outlines the numerous uses of clinical photography in the medical field. Regard must be had for the legal and ethical obligations of practitioners to uphold a patient's right to privacy, confidentiality and autonomy to consent to the use of photographs.² Current research focuses on the use of clinical photography by dermatologists.^{3–5} However, clinical photography is also a valuable tool for general practitioners (GPs), who are often the first to assess skin conditions. It is therefore important for GPs to be aware of their ethical and legal obligations.

Table 1. Primary categories and uses of clinical photography of skin lesions						
Category	Uses					
Clinical	 Documentation of condition Teledermatology Tracking patient progress Evidence in case of future legal action 					
Academic	 Visual aid that supports verbal clinical descriptions to benefit medical students Assists professionals to obtain a better understanding of skin conditions and lesions 					
Research	Photographs are mandatory for most research work as they help to substantiate findings; this contributes to the spread of knowledge and improvement in healthcare provision					
Commercial	Advertisements to generate public awareness about various conditions and the treatment options available					

SKIN CANCER REFERRALS

Dr Andrew Lewandowski Plastic and Reconstructive Surgeon Royal Brisbane and Womens Hospital





HOW DO WE IMPROVE ALLIANCE

Communication via referral / correspondence

- Urgent vs non-urgent
- Finite resources to manage everything need to stratify

URGENT VS NON-URGENT

Iceberg

Difficult to differentiate sometimes

Patients stoic

Others not...stoic

Small lesion may have iceberg extension into deeper tissues

Large fungating tumours may actually be pedunculated and not significantly invasive



ICEBERG VS UMBRELLA



KEY TERMS / INFORMATION

- Symptoms suggestive of adverse features
 - Formication (latin *formica* "ants" crawling under skin)
 - Lymphadenopathy
 - B symptoms weight loss, fevers
 - Rapid growth
- Biopsy features
 - Clinical size of lesion
 - Proximity to important structures (eyes, nose, lips)
 - PNI / LVI
 - IEC adnexal growth vs not (suitability for non-operative)
- Radiology
 - CT / USS reporting lymphadenopathy suspicious for metastasis

Relevant Medical Information

- Necessary for triaging into correct service
 - Main clinic
 - See and Treat Rapid Access Clinics
- Comorbidities
 - Major cardiac / respiratory
 - Blood thinners
 - T2DM
 - Immunosuppression (higher risk)
- Accessibility
 - Mobile
 - Wheelchair bound
 - Nursing home in a bed requiring Ambulance transport

WHAT HAPPENS – ROYAL SURGERY

- Melanoma generally standalone surgical clinic
 - Staffed by Jason Miller (general surgery), Brendan Louie & Andrew Lewandowski (PRS)
 - See patients ideally within 2 weeks
- Melanoma causes significant stress and anxiety given its social stigma
- Also see Merkel Cell, Dermal Sarcoma (AFX/PDS) urgently
- Thinner MIS or TIa melanomas where sentinel node not generally indicated go out to other consultants still relatively urgently (2-4 weeks)

WHAT HAPPENS – ROYAL SURGERY

- Main plastic surgical clinic
 - Average half day clinic ~ 30 patients of varying complexity, generally 5-10 new per consultant
 - See variety of presentations for NMSC, general reconstruction (some sub-specialty areas)
 - Makes up to ~ 30 new patients per week
- Lesions on limbs / trunk offered to General Surgery as well unless significant size / complexity
- Utilising SuperClinics getting every consultant in unit and every registrar in on a Saturday – see 150 patients
- Newly setup NewCase clinics seeing 25/week to get on top of waitlist

THE WAITLIST IS GROWING

- I'm not here to say we don't want your patients, we DO want to see them
- I'm not here to preach
- I think its important for you all to hear DIRECTLY from the source we WANT to be contacted and annoyed if your patient comes back with increased size, worsening features/symptoms
- If something doesn't feel right call us and escalate
- GP are primary point of contact and manage vast majority of skin cancer in community
 - When it is in difficult areas, has a significant surface area, has escalated to a point where not treatable in community OR multiple involved margins good reason to send to us

General Practice Liaison Officer Program

Update on melanoma and immunotherapy

Dr Melissa Eastgate Operations Director & Senior Medical Oncologist, Cancer Care Services | RBWH





Metro North Health

Update on Melanoma and immunotherapy

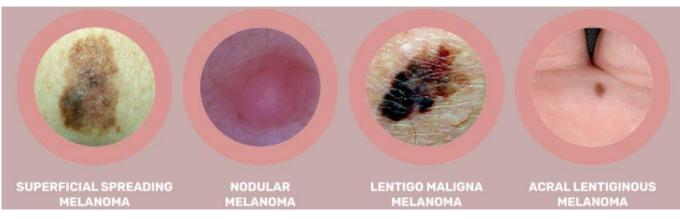
A/Prof Melissa Eastgate

Operations Director, CCS

Senior Medical Oncologist

Melanoma – Australian Stats

- Australia and NZ have the highest rates of melanoma in the world
- 4180 new cases in Qld each year
- 1 Australian Dx every 30 minutes
- 2nd most common cancer in men and women



55-60% of cases age < 40 yrs Can start as new brown/black spot or an existing spot that demonstrates change Trunk, slow growing

10-15% of cases age > 65 yrs Round, raised firm lump that is pink /red / brown / black. Can have crusty surface w/ tendency to bleed Head and neck, fast growing

10-15 % of cases age > 40 yrs Large coloured spot in sun-damaged skin. Face, ears, neck, head. Can grow slowly and superficially 1-2% of cases age > 40 yrs Palms, soles, fingernails/toenails. Colourless or lightly coloured, long streak of pigment in nails.

Metastatic melanoma

up to 10% have unknown 1°

M1a: mets to distant skin, subcutaneous or LN sites

M1b: lung mets

M1c: non-CNS visceral mets

M1d: CNS mets with/without other sites involved

Prognosis:

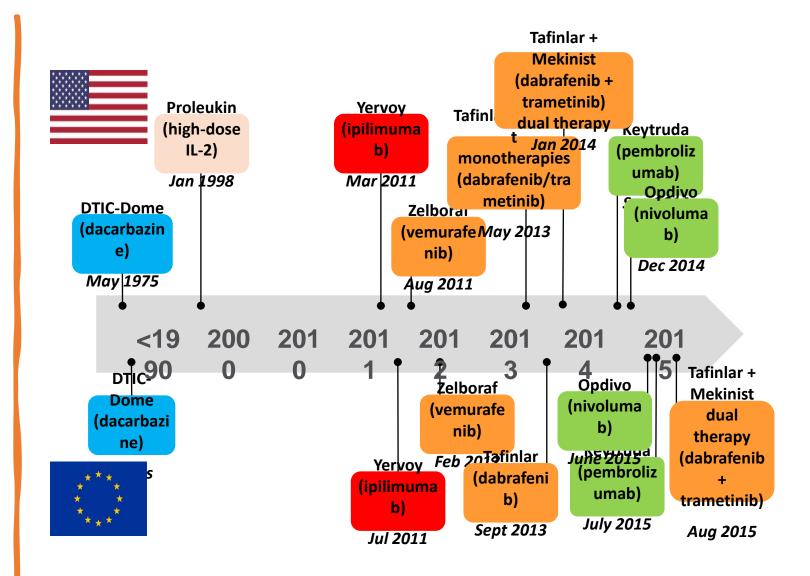
- mOS of untreated stage IV melanoma: <12 mo, 5yr OS: 10%
- M1a Prognosis: up to 15 mo
- M1c Prognosis: 6-9 mo
- M1d Prognosis: ~ 3 months
- Poor prognostic features: high tumour burden, elevated LDH, poor PS

1. DeVita et al

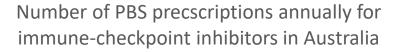
2. Davies et al. Prognostic factors for survival in melanoma patients with brain mets. Cancer 2011; 117(8):1678-96

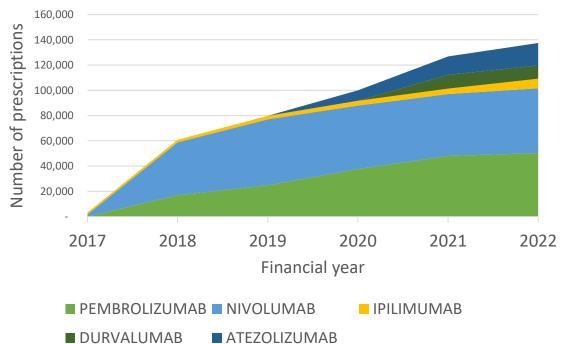
3. Vecchio et al. The treatment of melanoma brain mets before the advent of targeted therapies. Melanoma Res. 2014;24(1):61-7.

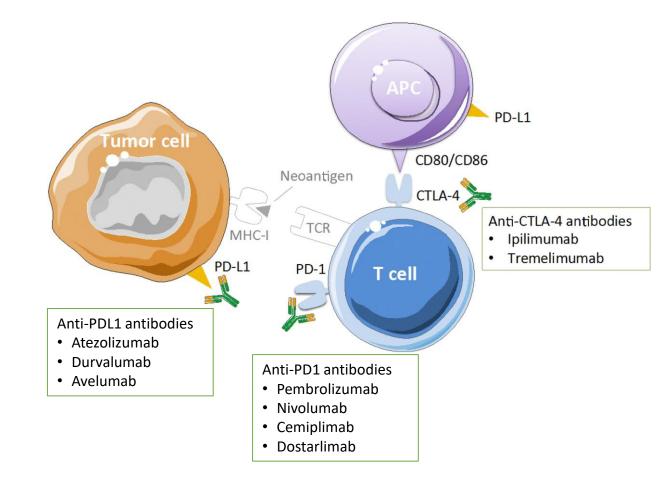
Metastatic melanoma treatments



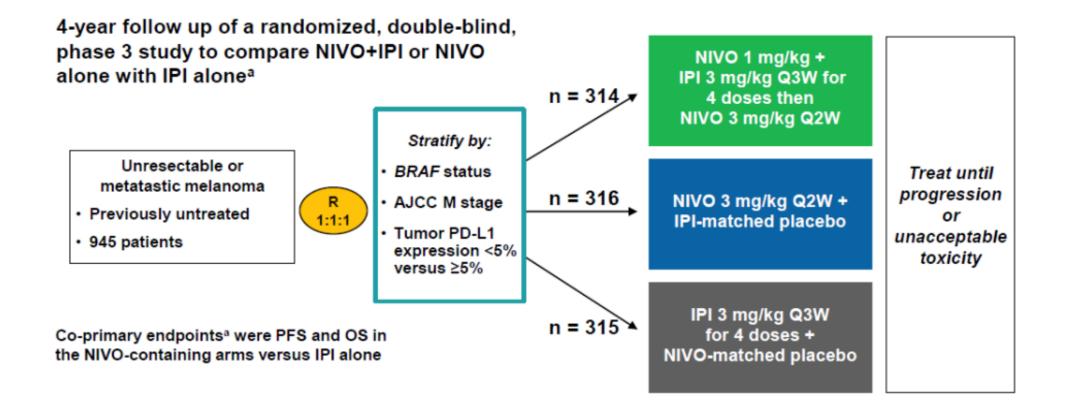
Immune checkpoint inhibitors





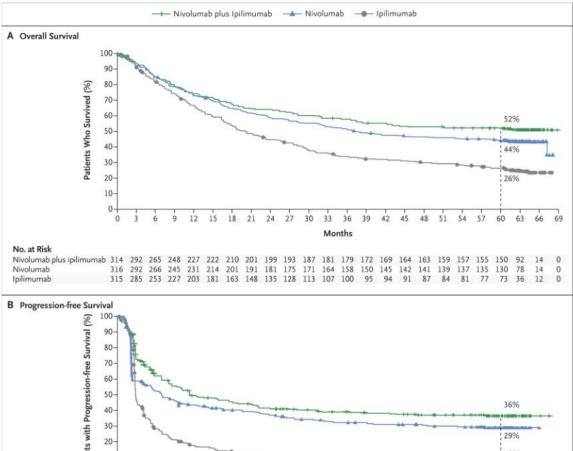


Met melanoma, BRAF WT Checkmate 067 trial (2017, 2019)



Checkmate 067 (2017, 2022) -Efficacy A Overall Survival

	lpi + Nivo	Nivo	lpi
mOS (mo)	72.1	36.9	19.9
PFS (mo)	11.5	6.9	2.9
5yr OS	52%	44%	26%
6.5yr MSS	NR	58.7	21.9
ORR	58%	45%	19%
CR	23%	19%	6%
PR	36%	26%	13%
SD	12%	9%	22%
PD	24%	38%	50%
mDOR (mo)	NR (61.9-NR)	NR (45.7 –NR)	19.2 (8.8 – 47.4)



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No. at Risk																									
Nivolumab plus ipilimumab	314	218	174	155	136	131	124	117	110	104	101	97	95	91	90	88	82	79	76	69	45	19	2	0	
Nivolumab	316	177	151	132	120	112	106	103	97	88	84	80	78	76	73	71	68	66	65	60	40	13	1	0	
Ipilimumab	315	136	78	58	46	42	34	32	31	29	28	26	21	19	18	18	17	15	15	15	11	8	1	0	
																									1

36 39

Months

33

42 45

48

12 15 18 21 24 27 30

29%

8%

63

66

51 54 57 60

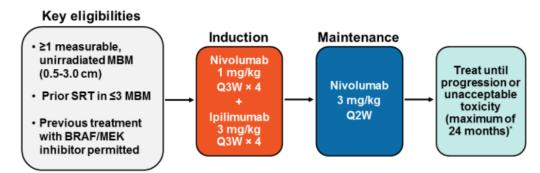
M1d Melanoma – Brain mets

- mOS of melanoma brain mets (MBM): 2.8-4 mo
- 2 trials: ABC (n=76) and CM 204 (n=94)
- ABC-X trial underway: Ipi/Nivo + SRT vs Ipi/Nivo

A: asymptomatic brain mets, no local Rx: Ipi/Nivo then NivoB: asymptomatic brain mets, no local Rx: NivoC: brain mets failed local Rx, neuro Sx, leptomeningeal disease: Nivo

	A (ipi+nivo)	B (nivo)	C (nivo)
All patients	n=35	n=25	n=16
ICR	51%	20%	6%
5-yr IC PFS	46%	15%	6%
5-yr OS	51%	34%	13%
Rx naïve	n=27	n=19	n=4
ICR (Rx naïve)	59%	21%	25%
5-yr IC PFS (Rx naïve)	52%	14%	
5-yr OS (Rx naïve)	55%	40%	25%
TRAE G3/4	63%	20%	13%

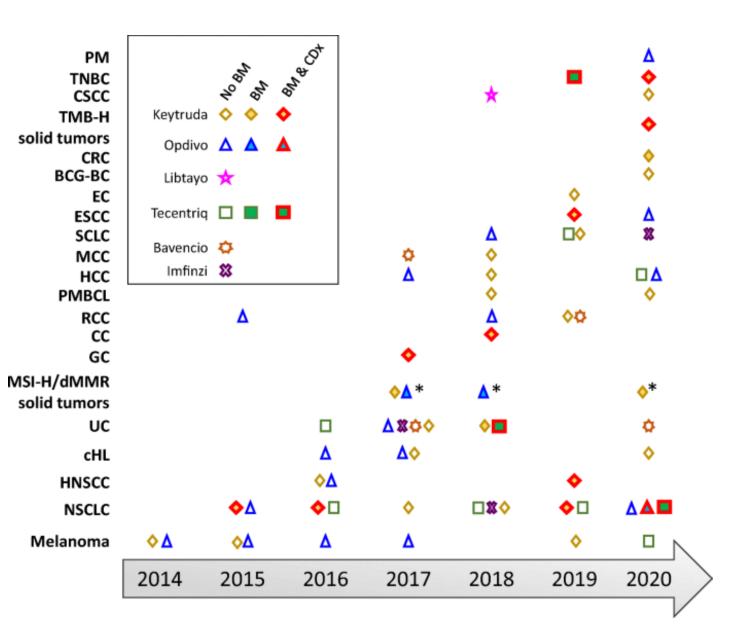
CM 204 schema



Variable	Intracranial (N=94)	Extracranial (N = 94)	Global (N=94)
Best overall response — no. (%)*			
Complete response	24 (26)	7 (7)	8 (9)
Partial response	28 (30)	40 (43)	40 (43)
Stable disease for ≥6 mo	2 (2)	6 (6)	5 (5)
Progressive disease	31 (33)	28 (30)	33 (35)
Could not be evaluated†	9 (10)	13 (14)	8 (9)
Objective response‡			
No. of patients	52	47	48
Percent of patients (95% CI)	55 (45–66)	50 (40–60)	51 (40-62)
Clinical benefit§			
No. of patients	54	53	53
Percent of patients (95% CI)	57 (47–68)	56 (46–67)	56 (46–67

ABC trial: https://www.thelancet.com/journals/lanonc/article/PIIS1470-2045(18)30139-6/abstract

Increasing indications



Stage 3 melanoma



Long-Term Follow-Up for Adjuvant Dabrafenib Plus Trametinib in Stage III BRAF-Mutated Melanoma: Final Results of the COMBI-AD Study

Axel Hauschild, Reinhard Dummer, Mario Santinami, Victoria Atkinson, Mario Mandala, Barbara Merelli, Vanna Chiarion-Sileni, Andrew Mark Haydon, Jacob Schachter, Dirk Schadendorf, Thierry Lesimple, Elizabeth Ruth Plummer, James Larkin, Monique Tan, Sachin Bajirao Adnaik, Paul Burgess, Tarveen Jandoo, <u>Georgina V. Long</u>





Presentation

https://bit.ly/Hauschild9500

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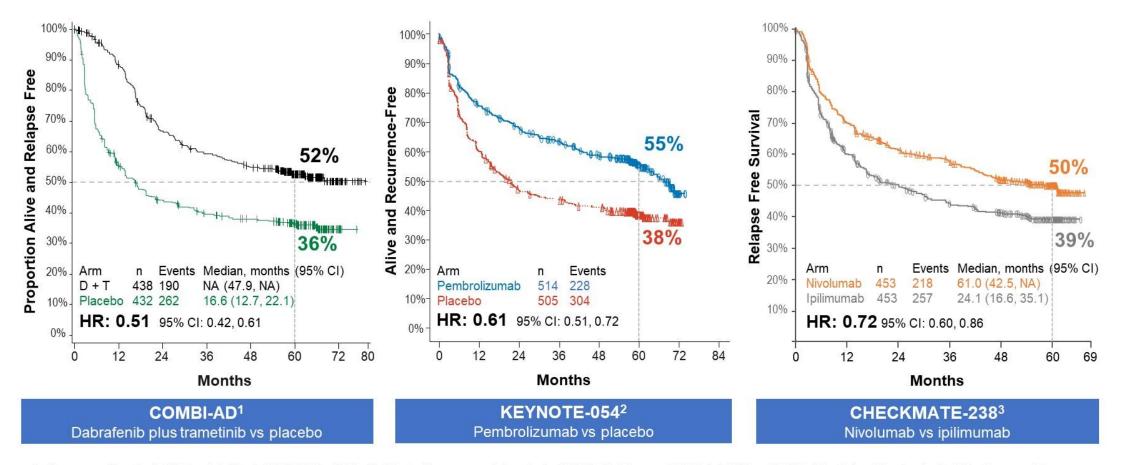
V. Long 🛛 🞯 @profglong

fglong 😏 @ProfGLongMIA

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Adjuvant Therapy in Resected Melanoma in 2024 RFS at 5 Years of Follow-Up



1. Dummer R, et al. N Engl J Med. 2020;383:1139-1148. 2. Eggermont A, et al. NEJM Evidence. 2022;1:EVIDoa2200214. 3. Larkin J, et al. Clin Cancer Res. 2023;29:3352-3361.

2024 ASCO ANNUAL MEETING

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Survival Endpoints at 5 Years of Follow-Up

Endpoint	COMBI-AD ¹	KEYNOTE-054 ²	CHECKMATE-238 ³		
Population	Dabrafenib + trametinib (n=438) vs placebo (n=432) <i>BRAF</i> V600E/K only	Nivolumab (n=453) vs ipilimumab (n=453)			
Melanoma	AJCC 7 th edition	AJCC 7 th edition	AJCC 7 th edition		
stage	Stage IIIA-C	Stage IIIA-C	Stage IIIB-C/IV		
RFS	52% vs 36%	55% vs 38%	50% vs 39%		
	HR: 0.51	HR: 0.61	HR: 0.72		
	95% CI: 0.42, 0.61	95% CI: 0.51, 0.72	95% CI: 0.60, 0.86		
DMFS	65% vs 54%	61% vs 44%	58% ^a vs 51% ^b		
	HR: 0.55	HR: 0.62	HR: 0.79		
	95% CI: 0.44, 0.70	95% CI: 0.52, 0.75	95% CI: 0.63, 0.99		
OS	Not analyzed ^c	Not analyzed	76% vs 72% HR: 0.86 95% CI: 0.66, 1.12		

^an=370; ^bn=366; ^cInadequate number of events to trigger the final analysis.

1. Dummer R, et al. *N Engl J Med.* 2020;383:1139-1148. 2. Eggermont A, et al. *NEJM Evidence*. 2022;1:EVIDoa2200214. 3. Larkin J, et al. *Clin Cancer Res.* 2023;29:3352-3361.



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Neoadjuvant therapy

ORIGINAL ARTICLE

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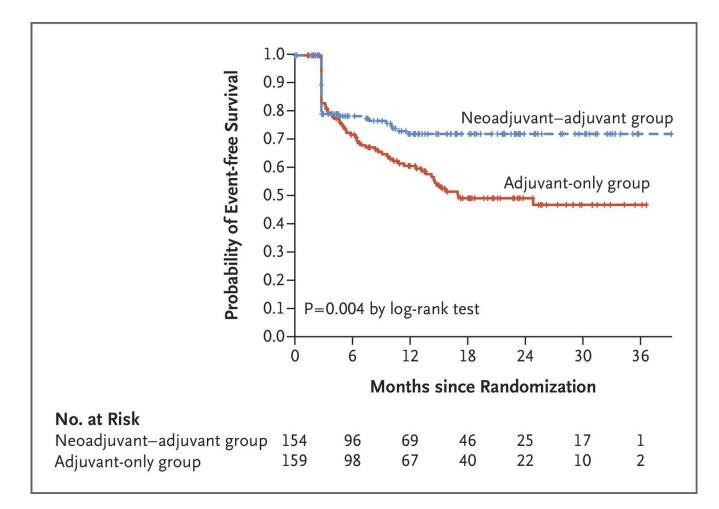
Neoadjuvant–Adjuvant or Adjuvant-Only Pembrolizumab in Advanced Melanoma

Authors: Sapna P. Patel, M.D. ^(D), Megan Othus, Ph.D., Yuanbin Chen, M.D., Ph.D., G. Paul Wright, Jr., M.D., Kathleen J. Yost, M.D., John R. Hyngstrom, M.D., Siwen Hu-Lieskovan, M.D., Ph.D., 45, and Antoni Ribas, M.D., Ph.D. Author Info & Affiliations

Published March 1, 2023 | N Engl J Med 2023;388:813-823 | DOI: 10.1056/NEJMoa2211437 | <u>VOL. 388 NO. 9</u> Copyright © 2023



Neoadjuvant therapy





Neoadjuvant Nivolumab Plus Ipilimumab Versus Adjuvant Nivolumab in Macroscopic, Resectable Stage III Melanoma: The Phase 3 NADINA Trial

Christian U. Blank, M.W. Lucas, R.A. Scolyer, B.A. van de Wiel, A.M. Menzies, M. Lopez-Yurda, A.C.J. van Akkooi, W.J. van Houdt, R.P.M. Saw, A. Torres-Acosta, S.N. Lo, G.A.P. Hospers, M.S. Carlino, J.W.B. de Groot, E. Kapiteijn, K.P.M. Suijkerbuijk, P. Rutkowski, S. Sandhu, A.A.M. van der Veldt, G.V. Long

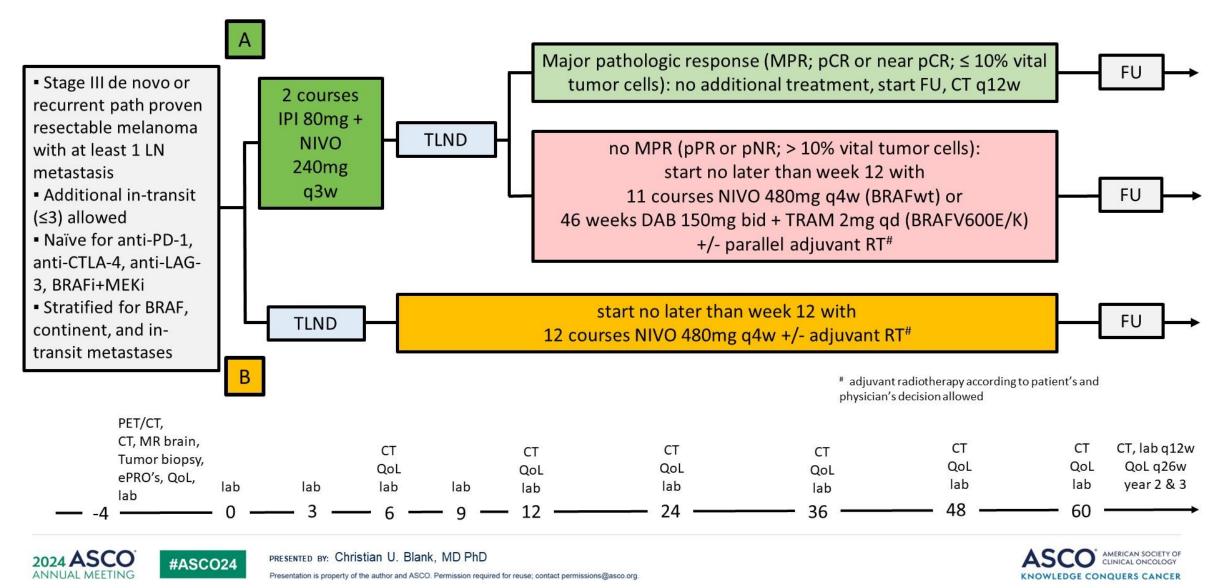


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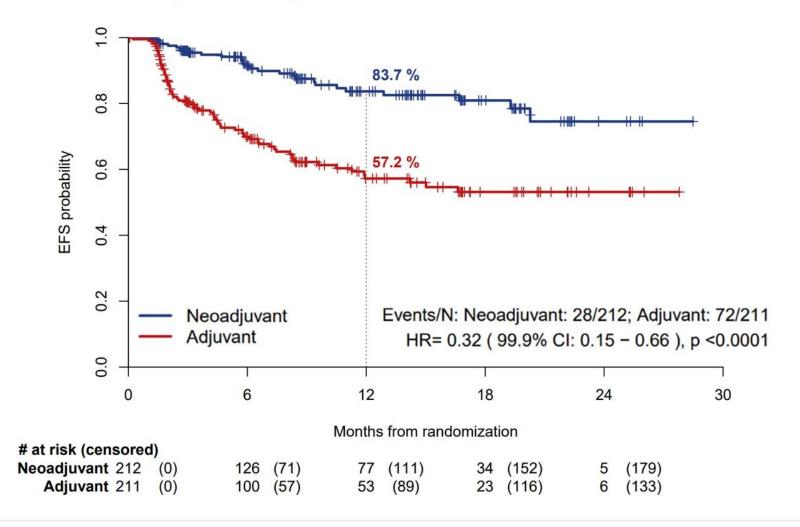
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NADINA - Trial Design



NADINA – Primary Endpoint: Event-Free Survival (EFS)



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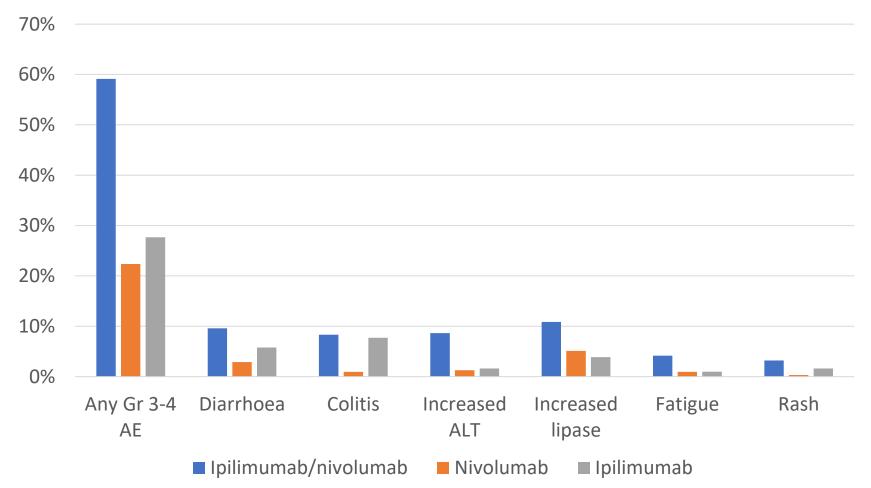
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Immune checkpoint inhibitors

Selected grade 3-4 toxicities from Checkmate-067 study in metastatic melanoma



Gastritis/duodenitis

- Much less common than lower GI toxicity; can occur together
- Incidence unclear, case reports/series; estimated <1%
- Symptoms include:
 - Nausea/vomiting (50-100%)
 - Abdominal pain (30-75%)
 - Dyspepsia (38%)
 - Bleeding (18% in one case series)
 - Concomitant lower GI symptoms eg. diarrhoea up to 50%
- Management guided by case series as for lower GI toxicity with steroids and anti-TNF-α agents in refractory cases

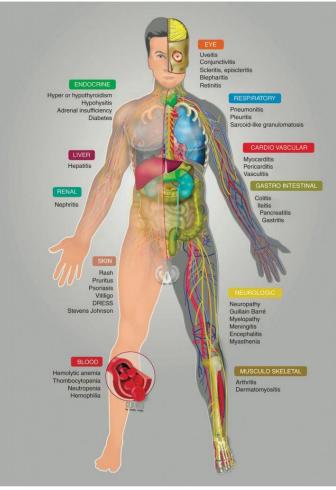
Endocrinopathies

- Clinically significant endocrinopathies in up to 10% patients
 - Most commonly hypothyroidism
 - Can include: hypoadrenalism, hypopituitarism / hypophysitis, diabetes
 - Incidence of ICI-associated diabetes <1%
- Irreversible
- Steroids rarely indicated
- Do not preclude further immunotherapy

Fever and systemic inflammatory syndromes

- Fever and systemic inflammatory complications of immunotherapy can occur including cytokine release syndrome (CRS) and haemophagocytic lymphohistiocytosis (HLH)
- Requires usual workup to exclude infectious causes
- Limited data to guide management but can be steroid responsive

Summary



Champiat et al, Ann Oncol, 2016

Metastasectomy

- For palliation of symptoms
- May be associated with improved long-term survival following complete curative resection
- Generally, best outcomes in pts with indolent disease, fewer metastatic sites and metasatatic disease that can be completely resected
- In pts with major response to systemic therapy, complete resection of residual mets may contribute to improved DFS and potential for cure
- Lung: 22% 5 yr survival, 16% 10 yr survival cf 0% after incomplete resection, SBRT is an option
- Liver: mOS 24.8mo vs 8mo (highly selected group, n= 58/1078)

Summary

- Metastatic Melanoma outcomes have vastly improved since IO + TT became available
- BRAF WT Melanoma:
 - Ipi / Nivo
 - Relatlimab / Nivo –available on PBS
 - Nivo
- BRAF MT Melanoma
 - IO then TT > TT then IO (Dream-Seq, SECOMBIT)
 - ? Ipi /Nivo over Relat/Nivo
 - D+T; Enco / Bini; Vemurafinib + Cobimetinib
- Brain mets
- Metastasectomy
- Questions:
 - Ipi / Nivo vs Rela / Nivo
 - Does Ipi / Nivo still have activity following PD on Relat/Nivo (currently not allowed on PBS)

Summary

 Systemic therapies are moving further forward in the patient journey

Thank you!