

Treatment options for a large facial lentigo maligna



Juliet Smith, Lisa Byrom, Jim Muir

CASE

A man aged in his late 70s presented with an 11-year history of a pigmented lesion on his left cheek, which he reported has recently changed in size and colour. There is scarring and hypopigmentation in the area secondary to previous excision of a basal cell carcinoma (BCC) and use of imiquimod for solar keratosis many years earlier.

The patient is of Fitzpatrick skin type 2 with a lifelong history of recreational sun exposure, multiple facial and trunk BCCs and no history of melanoma. Medical history includes type 2 diabetes, benign prostate hyperplasia, polymyalgia rheumatica, hypertension, peripheral vascular disease and macular degeneration; the patient is an ex-heavy smoker and lives alone.

QUESTION 1

On examination there is an asymmetrical, variably pigmented patch with background actinic damage on the left cheek measuring 50 mm×45 mm (Figure 1). Dermoscopy reveals a non-uniform lesion coloured brown, grey, pink and white, with an irregular structure, annular granular pattern, rhomboid

structures, grey dots/concentric circles and structureless areas (Figure 2).

What is your favoured diagnosis? What differentials might be considered?

QUESTION 2

What would the initial management involve?

ANSWER 1

The most likely diagnosis is lentigo maligna (LM), an in situ melanoma confined to the epidermis, or lentigo maligna melanoma (LMM), where it has become invasive.

Typical features of differential diagnoses are provided in Table 1.

ANSWER 2

Clinical examination for lymphadenopathy and hepatosplenomegaly is crucial. LM can harbour significant subclinical extension and occult dermal invasion.

Recommended initial management for suspected melanoma is excisional biopsy with 2-mm margins.¹ Due to the size and location of this lesion, a central incisional biopsy of the lesion was performed (see below), alternatives being multiple punch/shave scouting biopsies (any partial biopsy carries risk of false negatives).^{1,2}

Determination of lesion extent is crucial to management planning. Techniques, including

dermoscopy, Woods light examination and mapping biopsies, can aid lesion delineation, and were subsequently performed (see below). Confocal microscopy might be used for this purpose but has limited availability.³

CASE CONTINUED

An incisional biopsy, 18 mm×10 mm×8 mm, showed Clark Level 1 (in situ melanoma) of the LM type. Melanin incontinence was present, with no evidence of adnexal extension or dermal invasion.

To determine lesion extent, four 2-mm diameter circumferential mapping punch



Figure 1. Pigmented facial lesion.

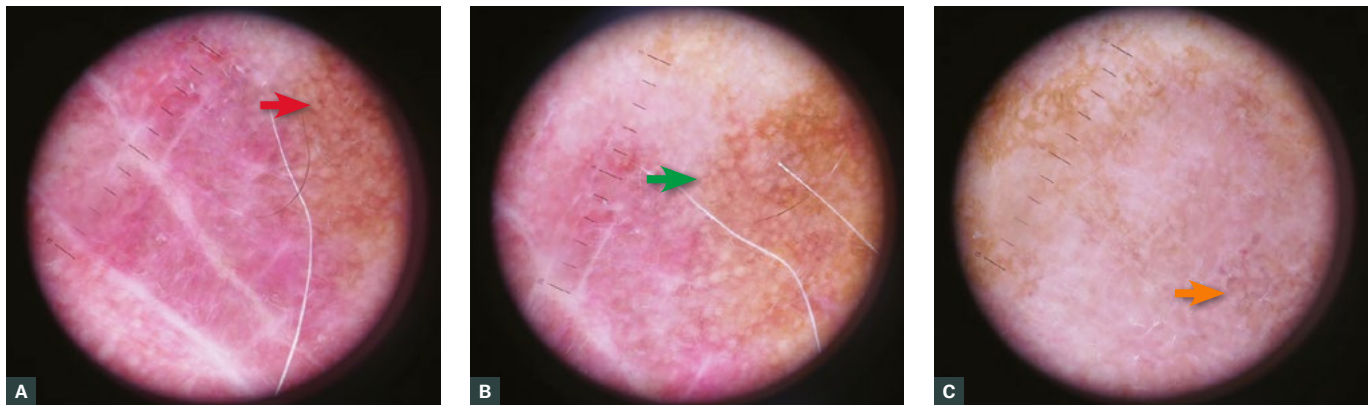


Figure 2. Dermoscopic features of the lesion. (A) Rhomboid structures (red arrow). (B) Structureless areas (green arrow). (C) Grey dots/grey concentric circles (orange arrow).

Table 1. Typical features of differential diagnoses

Condition	Macroscopic features	Dermoscopic features
Lentigo maligna/lentigo maligna melanoma	<ul style="list-style-type: none"> Often >6 mm, irregular shape, variable pigmentation, tan, light and dark brown, pink, red and white colours, with a smooth surface 	<ul style="list-style-type: none"> Asymmetrical pigmented follicular openings, rhomboid structures, grey pseudonetwork, annular granular pattern, grey dots and globules⁴
Pigmented intraepidermal carcinoma/Bowen's disease	<ul style="list-style-type: none"> Typically irregular scaly macular to raised lesions Might be several centimetres in diameter, orange-red or brown in colour, with areas of pigmentation and smooth, hyperkeratotic or ulcerated 	<ul style="list-style-type: none"> Dots, structureless areas, dots in linear arrangement, coiled vessels⁴
Pigmented actinic keratosis	<ul style="list-style-type: none"> Flat or thickened papule or plaque Pigmented or white, yellow or red Scaly, warty or horny surface 	<ul style="list-style-type: none"> Hyperpigmented follicular openings, brown structureless areas, annular-granular structures Occasionally angulated superficial brown lines Perifollicular inner grey halo, grey rhomboidal structures⁴
Solar lentigo	<ul style="list-style-type: none"> Flat macule or patch, multiple colours, including yellow, light or dark brown, regular or irregular border, dry surface, well circumscribed, moth-eaten outline 	<ul style="list-style-type: none"> Faint pigment network, fingerprint structures, uniform pigmentation, elongated rete ridges⁴
Lentiginous nevus	<ul style="list-style-type: none"> Irregular borders, multiple colours, including pink, red, tan, brown and black, and a diameter usually >5 mm 	<ul style="list-style-type: none"> Reticular, globular, asymmetrical pigmentation, regression structures, irregular vascular pattern, blue-grey areas⁴
Melasma	<ul style="list-style-type: none"> Bilateral, blotchy, brownish facial pigmentation, light-to-dark brown macules or patches with irregular borders 	<ul style="list-style-type: none"> Light to dark brown reticular network, brown dots, granules and globules, arcuate and annular structures, sparing of perifollicular region and openings of sweat glands, arcuate and annular pattern⁴
Seborrhoeic keratosis	<ul style="list-style-type: none"> Well demarcated, flat or raised, waxy papules and patches, central verrucous changes Skin coloured, yellow, grey, light brown, dark brown, black or mixed colours 	<ul style="list-style-type: none"> Comedo-like openings, milia-like cysts, hairpin vessels, fingerprinting, moth eaten and sharply demarcated, network-like structures, pseudonetwork⁴
Pigmented basal cell carcinoma	<ul style="list-style-type: none"> Areas of pigmentation, nodular or plaque, pearly appearance, telangiectasia, central ulceration 	<ul style="list-style-type: none"> Leaf-like structures, blue-grey ovoid nests, spoke-wheel areas, blue-grey globules, arborising telangiectasia and ulceration⁴

For those interested in a more comprehensive comparison, see Todorovic-Zivkovic et al.⁵

biopsies were taken 10 mm from the lesion. The left inferior specimen showed LM. The other biopsies showed actinic damage and basal layer hyperpigmentation associated with a focally lentiginous pattern.

QUESTION 3

What are the management options for this patient?

ANSWER 3

The clinical situation is poorly defined LM on the left cheek, where invasive disease has not been excluded, in an elderly patient with multiple medical comorbidities. First-line treatment is excision with 5- to 10-mm margins, although lesions larger than 10 mm or on the head and neck might require a wider margin.^{3,6} Recurrence rates for excised LM are approximately 6% at five years, but the risk of recurrence increases for lesions with a large diameter and/or those that are poorly defined.⁷

Excision options for this patient are as follows:

- excision with a repair that enables re-excision in the event of histological margin involvement³

- excision with margin control (eg Mohs surgery or staged excision with delayed definitive repair).^{4,7}

If surgery is refused or contraindicated, options include:

- radiation treatment^{8,9}
- imiquimod^{10,11}
- careful observation (only in selected cases with very elderly patients, or those with significant comorbidities and after review at multidisciplinary clinic with informed consent).¹²

Alternative treatment options (Table 2) might be discussed.

CASE CONTINUED

The patient refused surgery.

QUESTION 4

What would be your response in this situation?

ANSWER 4

When a patient declines optimal treatment, their reasons must be thoroughly and sensitively examined. Potential barriers include procedural fear, previous poor

surgical experience, concerns about scarring, concerns regarding general anaesthesia and cost. Factors to consider include allowing time in the consultation for all questions to be addressed, a support person to be present and organising patient follow-up. Comprehensive documentation of the discussion and patient consent are crucial.¹⁷

CASE CONTINUED

The patient was reviewed at a multidisciplinary clinic (plastic surgery, radiation oncology and dermatology) where excision was strongly recommended. Despite counselling, the patient declined, citing his age, frailty, comorbidities, lesion location, unwillingness to endure a substantial procedure and firm personal beliefs about illness and medicine. He was advised that surgical treatment had the highest cure rate and that the lesion could progress to invasive malignancy and metastatic disease. He understood these risks, and a support person attended appointments with him.

The patient opted to trial imiquimod treatment encompassing 10 mm outside the lesion margins. An adequate inflammatory response was achieved for

Table 2. Non-surgical treatment options

Treatment modality	Recurrence rate	Advantages	Disadvantages
Radiation therapy	<ul style="list-style-type: none"> • Mean recurrence estimate of 11.5%¹³ or recurrence estimate of 5% in 3 years with electron-specific treatment⁸ 	<ul style="list-style-type: none"> • Titration to adequate depth¹³ • Treat undetected invasion¹³ • Nil general anaesthetic or surgery 	<ul style="list-style-type: none"> • Multiple treatments • Short-term side effects: <ul style="list-style-type: none"> - Pain - Fatigue - Erythema - Desquamation - Mucosal irritation (not this particular patient) • Long-term side effects: <ul style="list-style-type: none"> - Radiation-induced optic neuropathy (not this particular patient) - Alopecia - Telangiectasia - Hyperpigmentation - Secondary malignancy
Imiquimod	<ul style="list-style-type: none"> • Mean recurrence estimate ranges between 0% and 40%^{14,15} 	<ul style="list-style-type: none"> • Non-invasive • Patient-controlled home treatment 	<ul style="list-style-type: none"> • Lack of long-term outcome data¹⁶ • Might not treat adnexal extension or occult invasion

12 weeks. Current guidelines suggest daily use for five days a week for 12 weeks.¹¹ Unfortunately, there was a poor clinical response, with dermoscopically evident extensive residual disease at review four weeks after treatment completion.

A six-week course of radiation treatment was prescribed. The patient experienced short-term mild erythema, dry desquamation and mild irritation to the treatment field managed with emollients. Mild hyperpigmentation was the only late toxicity noted.

There was complete clinical and dermoscopic clearance 12 weeks after treatment, and the patient remains free of any sign of local or systemic recurrence at the three-year follow-up (Figure 3).

Key points

- Large-diameter LM on sites of functional and cosmetic importance presents a difficult management issue.
- Surgical excision remains first-line treatment. When this is not possible due to lesion extent, comorbidities or patient preference, the patient should be counselled as to the available options and their relative risks and benefits.
- Radiation therapy, imiquimod or careful observation in selected patients are alternatives if surgery is not possible or declined.
- Strategies to support patients' autonomous and informed decision-making process include adequate consultation time, the facilitation of a support person and regular follow-up sessions.



Figure 3. Clinical and dermoscopic clearance at 12 weeks' post treatment.

Authors

Juliet Smith FRACGP, BMed, MPhil, General Practitioner, Institute of Dermatology, South Brisbane, Qld

Lisa Byrom MBBS, MPhil, BPhy, FACD, Consultant Dermatologist FACD, Mater Hospital, Brisbane, Qld

Jim Muir MBBS, FACD, Consultant Dermatologist FACD and Director, Mater Hospital, Brisbane, Qld

Competing interests: None.

Funding: None.

Provenance and peer review: Not commissioned, externally peer reviewed.

Correspondence to:

juliet.smith1@sydney.edu.au

References

- Sladden MJ, Nieweg OE, Howle J, Coventry BJ, Thompson JF. Updated evidence-based clinical practice guidelines for the diagnosis and management of melanoma: Definitive excision margins for primary cutaneous melanoma. *Med J Aust* 2018;208(3):137–42. doi: 10.5694/mja17.00278.
- Kasprzak JM, Xu YG. Diagnosis and management of lentigo maligna: A review. *Drugs Context* 2015;4:212281. doi: 10.7573/dic.212281.
- Guitera P, Moloney FJ, Menzies SW, et al. Improving management and patient care in lentigo maligna by mapping with in vivo confocal microscopy. *JAMA Dermatol* 2013;149(6):692–98. doi: 10.1001/jamadermatol.2013.2301.
- Soyer HP, Argenziano G, Hofmann-Wellenhof R, Zalaudek I. *Dermoscopy: The essentials*. 3rd edn. Elsevier, 2020.
- Todorovic-Zivkovic D, Argenziano G, Lallas A, et al. Age, gender, and topography influence the clinical and dermoscopic appearance of lentigo maligna. *J Am Acad Dermatol* 2015;72(5):801–08. doi: 10.1016/j.jaad.2015.01.030.
- Kunishige JH, Doan L, Brodland DG, Zitelli JA. Comparison of surgical margins for lentigo maligna versus melanoma in situ. *J Am Acad Dermatol* 2019;81(1):204–12. doi: 10.1016/j.jaad.2019.01.051.
- Hou JL, Reed KB, Knudson RM, et al. Five-year outcomes of wide excision and Mohs micrographic surgery for primary lentigo maligna in an academic practice cohort. *Dermatol Surg* 2015;41(2):211–18. doi: 10.1097/DSS.0000000000000248.
- Fogarty GB, Hong A, Scolyer RA, et al. Radiotherapy for lentigo maligna: A literature review and recommendations for treatment. *Br J Dermatol* 2014;170(1):52–58. doi: 10.1111/bjd.12611.
- Ballo MT, Ang KK. Radiation therapy for malignant melanoma. *Surg Clin North Am* 2003;83(2):323–42. doi: 10.1016/S0039-6109(02)00096-8.
- Tzellos T, Kyrgidis A, Mocellin S, Chan A, Pilati P, Apalla Z. Interventions for melanoma in situ, including lentigo maligna. *Cochrane Database Syst Rev* 2014;12:CD010308. doi: 10.1002/14651858.CD010308.pub2.
- Guitera P, Waddell A, Paton E, et al. A practical guide on the use of imiquimod cream to treat lentigo maligna. *Australas J Dermatol* 2021;62(4):478–85. doi: 10.1111/ajd.13720.
- Robinson M, Primiero C, Guitera P, et al. Evidence-based clinical practice guidelines for the management of patients with lentigo maligna. *Dermatology* 2020;236(2):111–16. doi: 10.1159/000502470.
- Read T, Noonan C, David M, et al. A systematic review of non-surgical treatments for lentigo maligna. *J Eur Acad Dermatol Venereol* 2016;30(5):748–53. doi: 10.1111/jdv.13252.
- Mahoney MH, Joseph MG, Temple C. Topical imiquimod therapy for lentigo maligna. *Ann Plast Surg* 2008;61(4):419–24. doi: 10.1097/SAP.0b013e31816714c8.
- Van Meurs T, Van Doorn R, Kirtschig G. Treatment of lentigo maligna with imiquimod cream: A long-term follow-up study of 10 patients. *Dermatol Surg* 2010;36:853–58. doi: 10.1111/j.1524-4725.2010.01560.x.
- Mora AN, Karia PS, Nguyen BM. A quantitative systematic review of the efficacy of imiquimod monotherapy for lentigo maligna and an analysis of factors that affect tumor clearance. *J Am Acad Dermatol* 2015;73(2):205–12. doi: 10.1016/j.jaad.2015.05.022.
- Australian Commission on Safety and Quality in Healthcare. Fact sheet for clinicians: Informed consent in health care. Australian Commission on Safety and Quality in Healthcare, 2020. Available at www.safetyandquality.gov.au/sites/default/files/2020-09/sq20-030_-_fact_sheet_-_informed_consent_-_nsqhs-8.9a.pdf [Accessed 2 October 2023].

correspondence ajgp@racgp.org.au