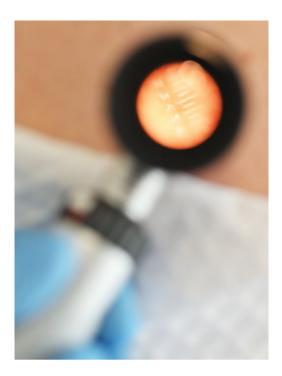
Skin cancer management:

Making the most of your pathologist



Simon P Clark, Olivia MM Clark, Cliff O Rosendahl

Background

Australia has the highest incidence of skin cancer globally. Early detection and treatment of skin cancer is critical for positive patient outcomes. General practitioners (GPs) play a central role in skin cancer management in Australia.

Objective

Collaboration between GPs and pathologists can improve the accuracy of skin cancer diagnosis. However, for improvement to occur, clear communication and high-quality specimens are essential.

Discussion

Inadequate clinical information and suboptimal biopsy specimens can hinder diagnosis. Improved communication, targeted training and selecting appropriate biopsy techniques are essential. A collaborative approach, guided by recommended techniques and clear guidelines, can minimise errors and improve patient outcomes in Australia's GP-led skin cancer management system. AUSTRALIA has the highest incidence of skin cancer globally. It has been estimated that in 2024, the healthcare costs for the three most common types of skin cancer basal cell carcinoma (BCC), squamous cell carcinoma (SCC) and melanoma - will exceed \$1 billion.¹ Each year, Australians experience an incidence of skin cancer far exceeding that in other populations, with two in three Australians diagnosed with skin cancer by the age of 70 years. For the most common forms - BCC, SCC and melanoma - vary widely in their presentation and prognosis and the complexity of treatment required. Among these, melanoma is particularly notable for its potential for fatal outcomes if not detected and treated early.

The high incidence and substantial burden of skin cancer in Australia not only reflect demographic and environmental factors, such as high levels of solar radiation, but also highlight the critical role of primary healthcare providers in managing this group of malignancies.² In contrast to other countries where dermatologists or surgeons manage most skin cancers, in Australia, general practitioners (GPs) manage the majority, playing a pivotal role in the detection, diagnosis and definitive treatment of skin cancers.³

This primary care-led approach is supported by Australia's healthcare

infrastructure, which facilitates the GP's role in skin cancer management through initiatives like skin cancer clinics and dedicated training programs in dermatoscopy and surgical skills. However, this model also results in a unique set of responsibilities and challenges for GPs, who must possess a high degree of skill in diagnostic and treatment procedures, as well as in navigating the complexities of when to refer patients to specialist care.

The histopathological evaluation of skin cancer is largely subjective. In most cases, the essential elements of a pathology report, diagnosis, evaluation of prognostic features and assessment of margin status (where relevant) are reliable. That is, the observations of one pathologist are likely to be repeated by a second pathologist and by the first pathologist on a second occasion. In certain situations, however, reliability is suboptimal, especially in the grading of dysplasia of naevi,4 the subtyping of BCCs,5 and the distinction between solar keratoses and invasive SCC.5 Furthermore, the evaluation of prognostic features6 and margin status7 might present problems of poor concordance.

In this setting, the collaboration between GPs and pathologists becomes critical. It is common knowledge that accurate histopathological diagnosis by pathologists informs clinical management decisions, affecting patient outcomes significantly. However, the nuances of interpreting pathology reports and the inherent limitations of histopathology might pose challenges, but these challenges can be mitigated by optimising communication between healthcare providers and the quality of skin pathology specimens.

Enhancing clinical communication and collaboration

The provision of comprehensive clinical data is deficient in many pathology specimen requests.8 Ferrara et al showed improved diagnostic reliability (and pathologists' confidence) with increments in provided clinical information, including clinical and dermatoscopic images.9 To facilitate this clinicopathological correlation, many Australian GPs send anonymised clinical and dermatoscopic images to the laboratory by email. An Australian study emphasised the risks of poor outcomes in cases where pathology request forms lacked adequate clinical information.10 It follows that improving the quality of clinical information conveyed by GPs to pathologists could significantly elevate the diagnostic accuracy and outcomes in skin cancer management.

Similarly, clear communication of histopathological evaluation of specimens requires unambiguous reports that are aligned to recommended treatments and clearly articulate any diagnostic uncertainty. Inconsistent terminology and inadequate contextual details frequently impede interpretation by clinicians. Targeted educational training co-delivered by GPs and pathologists can provide a foundation for better comprehension of the information requirements of each group.

Improving the quality of specimens

The quality of skin cancer specimens, in terms of their nature and extent, has a profound effect on the pathologist's ability to make accurate assessments. In Australia, the *Skin cancer clinical guidelines* highlight the desirability of excisional biopsy specimens while recognising that such specimens are not always appropriate.¹¹ The selection of biopsy technique should take into account the location of the lesion within the various skin layers and also the need, in many instances, for the architecture of the lesion to be assessed.

Sampling techniques, other than excisional biopsy, should be used judiciously. Partial punch biopsies, particularly small diameter punch biopsies, are associated with a significant rate of misdiagnosis, while often not demonstrating a lesion. In one study, 7% of cases sampled by punch biopsy and initially diagnosed as actinic keratosis were later found to harbour BCC or SCC.¹² Although it is likely that routine sampling of excisional specimens might also overlook foci of invasive cancer, the excisional technique ensures that the lesion is extirpated.

Great caution is required in the biopsy of suspected melanoma. The *Clinical practice* guidelines for the diagnosis and management of melanoma advise that 'the optimal biopsy approach for a suspicious pigmented lesion is complete excision with 2 mm clinical margin and upper subcutis'.¹³ Frequently a melanoma has its base transected in a shave biopsy specimen, limiting assessment of the Breslow thickness.¹⁴

More significantly, Kok et al, in a study based in Melbourne, showed that punch biopsy was associated with a false negative rate of diagnosis of more than 23% and shave biopsy with 4.5%, compared with less than 2% for excisional biopsy specimens.¹⁰ These under-called cases were associated with adverse outcomes, persistence or progression in 11.6% of the erroneously diagnosed punch biopsy specimens and in 1.7% of the shaved specimens but in only 0.7% of the excisional specimens. It is noteworthy that most (78%) of the misdiagnoses made on small punch biopsy specimens were due to misinterpretation by pathologists, while the remainder were due to sampling issues.

Very superficial shave biopsy specimens and small punch biopsy specimens often present the laboratory with technical difficulties in the provision of satisfactory microscopy slides. Shave biopsy specimens that do not include the reticular dermis (ie less than around 1-mm thick) tend to curl during fixation, resulting in microscopic sections that might not demonstrate the lesion or can result in sections that are tangential to the surface. Likewise, small punch biopsy specimens often shrink differentially, resulting in a specimen that is a truncated cone rather than a cylinder. The microscopy slides produced are, consequently, often obliquely sectioned. These technical problems might account for some of the difficulties in the interpretation of skin cancer pathology.

Incisional biopsies (ie partial biopsies through the diameter of a lesion) are less frequently used than punch or shave biopsies but are an excellent technique as the resulting slides closely resemble those of the excisional biopsy.¹⁵ An incisional biopsy allows the pathologist to evaluate the architecture of a lesion, and because the slides produced are identical, save for extent, to those of an excisional specimen, are likely to be associated with fewer diagnostic errors.

Ultimately, the goal of the biopsy is to achieve a balance between a minimally invasive procedure and the acquisition of pathologically valuable specimens. The choice of biopsy technique should be guided by the lesion's characteristics, the clinician's expertise and the available resources, aiming to maximise diagnostic accuracy while minimising patient discomfort, complications and the risk of incomplete excision, especially for those lesions where diagnostic discordance is relatively high. In these situations, complete excision ensures the lesion is destroyed, providing an added level of safety for the patient. Discussions with the pathologist can inform clinical decisions regarding the most appropriate biopsy technique.

Closing the loop

We have drawn attention to potential pitfalls in the pathological diagnosis of skin cancers. The harms that might result can be minimised with careful interpretation of the pathology report. Where there is discordance between the clinical and the pathological aspects, it is necessary for the clinician to ensure that the patient is managed correctly. Saving clinical and dermatoscopic images is important in this regard, as it provides a record that might be referred to once the pathology report has been received. It is customary to request that pathologists review the diagnoses in these situations, and because slides and tissue blocks are retained for many years, additional testing can be performed when re-evaluation or additional information is required. This additional testing might include examining deeper levels or the performance of

immunohistochemical stains. Opinions from

additional expert pathologists might also be

clinicopathological correlation in diagnosis

is the need for a good working relationship

between the clinician and the pathologist to

Aside from the immediate feedback

provided to the clinician by the histology

report, the pathology laboratory can also

clinical diagnosis and matching pathological

diagnosis can assist the clinicians to compare

In summary, we have highlighted the two

assessment of skin cancer cases, namely

but might be less well known to GPs.

skin cancer and improved practice.

skin cancer globally.

· Australia has the highest incidence of

pathology requests can lead to diagnostic

· Inadequate clinical information in

Quality of skin cancer specimens

pathological assessments.

diagnosis and treatment.

skin cancer management.

· Collaboration between GPs and

pathologists is crucial for accurate

· Awareness of issues in pathological

assessment can lead to more effective

significantly impacts the accuracy of

most important challenges in the pathological

adequacy of information and the skin cancer

Increasing awareness of these issues provides

an avenue for more effective management of

assist with audit activity. Collation of

facilitate communication, allowing resolution

Implicit in this goal of improved

of any diagnostic challenges.

performance with their peers.

Conclusion

Key points

challenges.

Authors

readily obtained.

University of Medical Sciences, Tehran, Iran Olivia MM Clark, BComm, Phlebotomist, Douglass Hanly Moir Pathology - Macquarie Park, Sydney, NSW

Simon P Clark MBChB, FRCPA, Dermatopathologist,

Douglass Hanly Moir Pathology - Macquarie Park, Sydney, NSW; Senior Lecturer, Faculty of Medicine, Cliff O Rosendahl MBBS, PhD, Professor, Faculty of Medicine, The University of Queensland, Brisbane, Qld; Visiting Professor, Department of Dermatology, Tehran University of Medical Sciences, Tehran, Iran Competing interests: None.

Fundina: None

Provenance and peer review: Commissioned, externally peer reviewed.

Correspondence to:

sclark@dhm.com.au

References

- Gordon LG, Leung W, Johns R, et al. Estimated healthcare costs of melanoma and keratinocyte skin cancers in Australia and Aotearoa New Zealand in 2021. Int J Environ Res Public Health 2022:19(6):3178. doi: 10.3390/ijerph19063178.
- 2 Thompson BS, Pandeya N, Olsen CM, et al. Keratinocyte cancer excisions in Australia: Who performs them and associated costs. Australas J Dermatol 2019;60(4):294-300. doi: 10.1111/ajd.13056.
- 3 Pandeya N, Olsen CM, Shalit MM, Dusingize JC, Neale RE, Whiteman DC. The diagnosis and initial management of melanoma in Australia: Findings from the prospective, population-based QSkin study. Med J Aust 2023;218(9):402-07. doi: 10.5694/mia2.51919.
- Elmore JG, Barnhill RL, Elder DE, et al. 4. Pathologists' diagnosis of invasive melanoma and melanocytic proliferations: Observer accuracy and reproducibility study. BMJ 2017;357:j2813. doi: 10.1136/bmj.j2813.
- Bingham SF; Department of Veteran Affairs Topical Tretinoin Chemoprevention Trial Group. Reliability of the histopathologic diagnosis of keratinocyte carcinomas. J Am Acad Dermatol 2007;57(2):279-84. doi: 10.1016/j.jaad.2007.03.021.
- 6 primary melanoma. Interobserver and intraobserver reliability, analyzed using H&E sections and immunohistochemistry. JDDG. J Dtsch Dermatol Ges 2016;14(9):910-15. doi: 10.1111/ddg.12797.
- Histopathologic recognition of involved margins of lentigo maligna excised by staged excision: An interobserver comparison study. Arch Dermatol 2003;139(5):595-604. doi: 10.1001/ archderm.139.5.595.
- 8. for skin lesions-Are we giving the pathologist sufficient clinical information? N Z Med J 2010;123(1325):53-58.
- 9. Ferrara G, Argenyi Z, Argenziano G, et al. The influence of clinical information in the histopathologic diagnosis of melanocytic skin neoplasms. PLoS One 2009;4(4):e5375. doi: 10.1371/journal.pone.0005375.
- 10. Kok Y, Scott K, Pham A, et al. The impact of incomplete clinical information and initial biopsy technique on the histopathological diagnosis of cutaneous melanoma. Australas J Dermatol 2021;62(4):e524-31. doi: 10.1111/ajd.13697.
- 11. Cancer Council Australia Keratinocyte Cancers Guidelines Working Party, Clinical practice guidelines for keratinocyte cancer. Sydney, NSW: Cancer Council Australia, 2024. Available at https://app.magicapp.org/#/guideline/n3QxOj [Accessed 16 April 2024].
- 12. Carag HR, Prieto VG, Yballe LS, Shea CR. Utility of step sections: Demonstration of additional pathological findings in biopsy

samples initially diagnosed as actinic keratosis. Arch Dermatol 2000;136(4):471-75. doi: 10.1001/ archderm.136.4.471

- 13. Cancer Council Australia Melanoma Guidelines Working Party. Clinical practice guidelines for the diagnosis and management of melanoma. Sydney, NSW: Cancer Council Australia, 2024, Available at https://app.magicapp.org/#/guideline/Lkk3pL [Accessed 16 April 2024].
- 14. de Menezes SL, Kelly JW, Wolfe R, Farrugia H, Mar VJ. The increasing use of shave biopsy for diagnosing invasive melanoma in Australia. Med J Aust 2019;211(5):213-18. doi: 10.5694/mja2.50289.
- 15. Pardasani AG, Leshin B, Hallman JR, White WL Fusiform incisional biopsy for pigmented skin lesions. Dermatol Surg 2000;26(7):622-24. doi: 10.1046/j.1524-4725.2000.00037.x.

correspondence ajgp@racgp.org.au

specimen. These issues have been canvassed in the dermatological and surgical literature

- 5. Jagdeo J, Weinstock MA, Piepkorn M,
- Garbe C, Eigentler TK, Bauer J, et al. Mitotic rate in
- 7. Florell SR, Boucher KM, Leachman SA, et al.
- Rademaker M, Thorburn M. Pathology referrals