

Clinical and Pathological Predictors of Unfavourable Outcomes in Thin and *In Situ* Melanomas: A Retrospective Cohort Study from Taiwan

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Thin melanoma, defined as tumour thickness ≤ 1.0 mm, and melanoma *in situ* are increasingly diagnosed. However, acral-predominant Asian populations appear to experience higher rates of adverse outcomes than Western cohorts. This study aimed to identify prognostic factors among Taiwanese patients with early-stage melanoma. We reviewed 178 melanoma *in situ*/thin melanoma cases diagnosed from 1995 to 2025 at a tertiary centre and analysed overall survival, melanoma-specific survival, recurrence-free survival, and distant metastasis-free survival using Cox and Fine-Gray competing-risk models. During a mean follow-up of 81 months, adverse events were not uncommon, including recurrence (8.8%), distant metastasis (4.7%) and melanoma-specific mortality (5.1%). Age was the most consistent predictor across endpoints in both Cox and competing-risk analyses. A Breslow thickness ≥ 0.8 mm independently increased the risk of melanoma-specific mortality, whereas ulceration and mitotic activity were not significant predictors. In conclusion, melanoma *in situ* and thin melanoma in this acral-predominant Asian cohort are not uniformly low risk. Older patients or those with tumours approaching or exceeding 0.8 mm in thickness warrant closer, risk-adapted surveillance.

Key words: age; cutaneous malignant melanoma; melanoma; prognosis; risk factors; survival analysis.

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The global incidence of melanoma continues to increase, largely due to increased detection of early-stage lesions – melanoma *in situ* (MIS) and thin melanoma (TM; Breslow thickness ≤ 1.0 mm) – through screening and awareness (1–3). Although melanoma has long been regarded as a malignancy of fair-skinned populations in Western countries, recent epidemiological reports from East Asia indicate a growing disease burden (4–6). Importantly, Asian

SIGNIFICANCE

Melanoma *in situ* and thin melanoma (1.0 mm or less) are often thought to be low risk. However, our study shows that some patients, especially those older than 60 years and those with Breslow thickness ≥ 0.8 mm lesions, can still face serious outcomes. By studying patients in Taiwan, where melanoma commonly appears on the feet and hands, we found that even early-stage disease is not always harmless. Our findings help doctors recognise who may need closer follow-up and raise public awareness supporting earlier care and potentially reducing preventable deaths.

patients often present with distinct clinicopathologic and molecular features, suggesting Western prognostic data may not apply to Asian populations (6–8).

In Western populations, early-stage melanoma is generally associated with excellent long-term outcomes, with 5 year melanoma-specific survival (MSS) rates approaching 97–99% for thin lesions (9–11). Nonetheless, a subset of patients with MIS or TM experience recurrence, distant metastasis or melanoma-specific death, and Breslow thickness ≥ 0.8 mm has emerged as a clinically relevant risk threshold (11–16).

In Asian populations, however, prognostic patterns may differ (1). Acral lentiginous melanoma (ALM) accounts for a larger proportion of cases and is frequently diagnosed at a later stage, contributing to poorer outcomes (6, 17, 18). In addition, ALM displays distinct molecular features compared with Western cohorts predominantly composed of superficial spreading melanoma (SSM), raising concerns regarding the direct applicability of Western-derived prognostic models to acral-predominant Asian populations (6, 19, 20).

Despite increasing reports, evidence specifically addressing MIS and TM in Asian patients remains limited. Prior reports from Taiwan and Korea have suggested higher recurrence and mortality than would be expected based on Western data and have highlighted older age, acral location, and male sex as adverse prognostic factors in localized melanoma (21–23). Because non-melanoma-related death is relatively common in ageing Asian populations and may preclude observation of recurrence or metastasis, competing-risk

methods are needed to obtain clinically realistic estimates of absolute risk (16, 24). Accordingly, the present study aimed to identify clinicopathological predictors of recurrence, distant metastasis and death in a Taiwanese cohort of early-stage melanoma. Using Cox proportional hazards models for overall survival (OS) and MSS, together with Fine–Gray competing-risk regression for recurrence and distant metastasis, we specifically examined whether age and Breslow thickness retain independent prognostic value in an acral-predominant population, and whether the commonly used 0.8 mm Breslow thickness threshold remains prognostically relevant in this setting.

MATERIALS AND METHODS

Study design and patient cohort

This retrospective study at National Taiwan University Hospital (NTUH) was approved by the institutional Research Ethics Committee (NTUH-REC No. 202503155RINA) adhering to the Declaration of Helsinki. We identified patients diagnosed with melanoma at NTUH between 1 January 1995 and 28 March 2025 using the electronic medical records. Of 638 records screened (Fig. S1), we excluded cases with Breslow thickness >1.0 mm or American Joint Committee on Cancer (AJCC) 8th edition stage >II ($n=458$), loss to follow-up ($n=1$) and unspecified histological subtype ($n=1$), leaving 178 early-stage (MIS and TM) patients.

We collected demographics, tumour features and follow-up information. Four outcomes were defined from the biopsy date: OS, time to death from any cause; MSS, time to melanoma-related death; distant metastasis-free survival (DMFS), time to first distant metastasis; and recurrence-free survival (RFS), time to first recurrence (local, regional nodal or distant). Causes of death and event dates were ascertained from electronic medical records and verified through linkage with the Taiwan National Death Registry.

Statistical analysis

Continuous variables were summarised as mean±standard deviation (SD), and categorical variables as frequencies and percentages; between-group comparisons used *t* test for continuous data and χ^2 or Fisher exact test for categorical data, as appropriate. Survival analyses were performed using the Kaplan–Meier method to estimate OS, MSS, DMFS and RFS. Group differences were assessed with the log-rank test.

Univariable Cox regression identified factors associated with survival outcomes. Age at diagnosis was modelled primarily as a continuous variable (per year); analyses stratified at ≤60 vs >60 years were provided for clinical interpretability. Variables

with $p<0.10$ in univariable analysis were subsequently entered into multivariable Cox models to determine independent predictors, and results were expressed as hazard ratios (HRs) with 95% confidence intervals (CIs). Because non-melanoma-related deaths were frequent, Fine–Gray competing-risk regression was used as the primary approach for MSS, DMFS and RFS (reporting subdistribution hazard ratios [SHRs]); Kaplan–Meier/Cox results for these endpoints are presented as complementary, descriptive analyses. All statistical analyses were conducted using SAS version 9.4 (SAS Institute, Cary, NC, USA) and R version 4.3. Statistical significance was defined as 2-tailed $p<0.05$.

RESULTS

A total of 178 patients with early-stage melanoma were included (Fig. S1). Of these, 89 (50.0%) had MIS and 89 (50.0%) had TM (**Table I**). Age and sex distribution were comparable between groups. The mean follow-up duration was longer in TM than MIS (89.99±95.74 vs 72.48±76.98 months); however, the difference was not statistically significant.

Anatomic distribution and histologic subtype showed marked differences between groups. Acral lesions were more frequent in MIS than in TM (87.6% vs 57.3%; $p<0.0001$), consistent with ALM being the predominant histologic subtype among MIS (87.6%). In TM, while the proportion of SSM (36.0%) was significantly higher than in MIS, ALM (57.3%) remained the most frequent histologic subtype. Ulceration was infrequent in TM (7/89, 7.9%) and a mitotic rate ≥ 1 /mm² was more common in TM than MIS (28.8% vs 13.2%, $p=0.045$). Tumour-infiltrating lymphocytes also differed significantly ($p<0.0001$): most MIS lesions had absent TILs (88.0%), whereas nonbrisk TILs predominated in TM (78.3%). Partial regression (19.6% vs 6.3%, $p=0.046$) and associated melanocytic lesions (21.4% vs 5.9%, $p=0.021$) were also more frequent in TM. Among TM cases, the mean Breslow thickness was 0.67±0.26 mm. Surgical wait time was shorter for MIS than TM (24.58±16.36 vs 28.45±22.76 days), but the difference was not statistically significant.

Among 171 patients with available recurrence data, 15 recurrences (8.77%) were identified, including 5 recurrences in MIS (5.68%) and 10 in TM (12.05%) (**Table II**). Among 172 patients with distant metastasis data, 8 distant metastases (4.65%) occurred: 1 (1.14%) in MIS and 7 (8.33%) in TM. Mortality data were available in 177 patients. Overall, 36 patients (20.34%) died, including 11 deaths in MIS (12.36%) and 25 in TM (28.41%). Among 175 patients with known causes of death, melanoma-specific deaths accounted for 9 cases (5.14%) – 2 in MIS (2.27%) and 7 in TM (8.05%) – whereas 27 deaths (15.43%) were unrelated to melanoma (10.23% in

Table I. Baseline clinicopathological characteristics of melanoma *in situ* and thin melanoma

Characteristic	Melanoma <i>in situ</i> (n=89)	Thin melanoma (n=89)	p-value
Sex, n (%)			0.548
Female	46 (51.7)	50 (56.2)	
Male	43 (48.3)	39 (43.8)	
Age at diagnosis, years, mean (SD)	60.12±14.46	59.47±15.57	0.773
Tumour location, n (%)			<0.0001*
Trunk	1 (1.1)	15 (16.9)	
Head and neck	2 (2.3)	3 (3.4)	
Extremities	8 (9.0)	20 (22.5)	
Acral	78 (87.6)	51 (57.3)	
Histological subtype, n (%)			<0.0001*
ALM	78 (87.6)	51 (57.3)	
SSM	7 (7.9)	32 (36.0)	
NM	1 (1.1)	3 (3.4)	
LMM	3 (3.4)	3 (3.4)	
Breslow thickness, mm, mean (SD)		0.67±0.26	
Ulceration, n (%)			
Absent		82 (92.1)	
Present		7 (7.9)	
Mitotic rate, n (%)			0.045*
<1/ mm ²	46 (86.8)	42 (71.2)	
≥1/ mm ²	7 (13.2)	17 (28.8)	
Tumour-infiltrating lymphocytes, n (%)			<0.0001*
Absent	44 (88.0)	7 (11.7)	
Nonbrisk	5 (10.0)	47 (78.3)	
Brisk	1 (2.0)	6 (10.0)	
Tumour regression, n (%)			0.046*
Absent	45 (93.8)	45 (80.4)	
Present	3 (6.3)	11 (19.6)	
Associated melanocytic lesion, n (%)			0.021*
No	48 (94.1)	44 (78.6)	
Yes	3 (5.9)	12 (21.4)	
Surgical wait time, days, mean (SD)	24.58±16.36	28.45±22.76	0.227
Follow-up time, months, mean (SD)	72.48±76.98	89.99±95.74	0.181

p-values were obtained using *t*-test (continuous) and χ^2 or Fisher exact test (categorical). Denominators reflect available data for each variable.

*Statistically significant.

ALM: acral lentiginous melanoma; LMM: lentigo maligna melanoma; NM: nodular melanoma; SSM: superficial spreading melanoma.

MIS vs 20.69% in TM). One patient was recorded as having melanoma-specific death according to the Taiwan National Death Registry despite no documented distant metastasis in the available clinical records, likely reflecting incomplete retrospective information regarding metastatic progression.

Overall survival showed a borderline difference between MIS and TM (log-rank $p=0.0719$), whereas MSS and RFS did not differ significantly ($p=0.1301$ and $p=0.1847$, respectively). In contrast, DMFS was significantly worse in TM (log-rank $p=0.0470$) (Fig.S2). By age group (≤ 60 vs >60 years), older patients had increased overall mortality risk ($p<0.0001$), increased melanoma-specific mortality risk ($p=0.0100$), and poorer DMFS ($p=0.0187$), but not RFS ($p=0.1778$) (Fig.S3).

Cause-specific Cox proportional hazards models were used to estimate melanoma-specific hazard ratios, whereas Fine-Gray subdistribution hazard models were applied to account for the competing risk of non-melanoma-related death. For OS (Table III, Fig. 1), age was a strong predictor both as a continuous variable (HR 1.11 per year; 95% CI, 1.07–1.15; $p<0.0001$)

Table II. Clinical outcomes in melanoma *in situ* and thin melanoma

Outcome	Melanoma <i>in situ</i>	Thin melanoma	Total
Recurrence			
Total patients, <i>N</i>	88	83	171
Event, <i>n</i> (%)	5 (5.68)	10 (12.05)	15 (8.77)
Distant metastasis			
Total patients, <i>N</i>	88	84	172
Event, <i>n</i> (%)	1 (1.14)	7 (8.33)	8 (4.65)
Deaths			
Total patients, <i>N</i>	89	88	177
Event, <i>n</i> (%)	11 (12.36)	25 (28.41)	36 (20.34)
Melanoma-specific deaths			
Total patients, <i>N</i>	88	87	175
Event, <i>n</i> (%)	2 (2.27)	7 (8.05)	9 (5.14)
Nonmelanoma deaths			
Total patients, <i>N</i>	88	87	175
Event, <i>n</i> (%)	9 (10.23)	18 (20.69)	27 (15.43)

For each endpoint, the total number of patients with available data (denominator) is indicated in the header. Events are reported as *n* (%) of that denominator.

and as a dichotomous variable (>60 vs ≤ 60 years: HR 12.60; 95% CI, 4.31–36.82; $p<0.0001$). TM showed a borderline association with increased overall mortality risk compared with MIS (HR 1.91; 95% CI, 0.93–3.93; $p=0.077$). Acral location (HR 2.11; 95% CI, 0.91–4.88; $p=0.081$) and ulceration (HR 2.63; 95% CI, 0.92–7.53; $p=0.072$) showed similar borderline associations with increased overall mortality. In multivariable analysis, age remained the only independent predictor of OS (adjusted HR 1.10; 95% CI, 1.07–1.14; $p<0.0001$), whereas TM retained a near-significant association (adjusted HR 2.13; 95% CI, 0.98–4.61; $p=0.056$).

For MSS, age remained significant in both univariable and multivariable models (continuous: HR 1.07; 95% CI, 1.01–1.13; $p=0.027$; adjusted HR 1.06; 95% CI, 1.01–1.12; $p=0.022$). In Cox analysis, Breslow thickness ≥ 0.8 mm showed a borderline association with increased melanoma-specific mortality but did not reach statistical significance after multivariable adjustment (adjusted HR 4.80; 95% CI, 0.93–24.80; $p=0.061$) (Table III, Fig. 1).

For RFS, age (continuous) was associated with an increased risk of recurrence (HR 1.05 per year; 95% CI, 1.01–1.10; $p=0.026$), whereas age >60 years and other clinicopathologic variables were not significant (Table SI). For DMFS, age >60 years was associated with a higher hazard of distant metastasis (HR 8.28; 95% CI, 1.01–67.65; $p=0.049$) with a similar trend for age as a continuous variable (HR 1.05; 95% CI, 0.99–1.11; $p=0.082$) (Table SI, Fig. 1).

Competing-risk analyses using Fine-Gray models are presented in Table SII. For MSS, age remained an independent predictor after accounting for competing deaths (SHR 1.05 per year; 95% CI, 1.02–1.08; $p=0.001$) and age >60 years was associated with markedly increased risk (SHR 8.25; 95% CI, 1.08–62.79; $p=0.042$). In the competing-risk framework, Breslow thickness ≥ 0.8 mm remained prognostically

Table III. Univariable and multivariable Cox regression analyses for overall mortality and melanoma-specific mortality

Variable	Overall mortality		Melanoma-specific mortality	
	Univariable HR (95% CI); <i>p</i>	Multivariable HR (95% CI); <i>p</i>	Univariable HR (95% CI); <i>p</i>	Multivariable HR (95% CI); <i>p</i>
Sex				
Female	Reference		Reference	
Male	1.24 (0.64–2.41); 0.520		0.85 (0.23–3.16); 0.808	
Age				
Continuous (per year)	1.11 (1.07–1.15); <0.0001*	1.10 (1.07–1.14); <0.0001*	1.07 (1.01–1.13); 0.027*	1.06 (1.01–1.12); 0.022*
Group >60 y (vs ≤60)	12.60 (4.31–36.82); <0.0001*		9.46 (1.18–75.96); 0.035*	
Tumour location				
Trunk	Reference			
Head and neck	1.07 (0.12–9.82); 0.954			
Extremities	0.26 (0.05–1.42); 0.120			
Acral	1.15 (0.39–3.39); 0.798			
Tumour site (Binary)				
Nonacral	Reference	Reference		
Acral	2.11 (0.91–4.88); 0.081	1.34 (0.54–3.32); 0.530		
Histological subtype				
ALM	Reference			
SSM	0.47 (0.18–1.21); 0.117			
NM	0.34 (0.04–2.65); 0.306			
LMM	0.82 (0.11–6.05); 0.845			
Melanoma type				
Melanoma <i>in situ</i>	Reference	Reference	Reference	
Thin melanoma	1.91 (0.93–3.93); 0.077	2.13 (0.98–4.61); 0.056	3.16 (0.66–15.23); 0.152	
Breslow thickness				
<i>In situ</i>	Reference		Reference	Reference
<0.8 mm	2.21 (0.99–4.90); 0.052		1.75 (0.25–12.43); 0.576	1.80 (0.25–12.81); 0.559
≥0.8 mm	1.63 (0.70–3.81); 0.259		4.68 (0.90–24.17); 0.066	4.80 (0.93–24.80); 0.061
Ulceration				
Absent	Reference	Reference	Reference	
Present	2.63 (0.92–7.53); 0.072	1.04 (0.35–3.13); 0.944	2.30 (0.29–18.45); 0.432	
Mitotic rate				
<1/mm ²	Reference		Reference	
≥1 /mm ²	1.50 (0.51–4.41); 0.459		4.35 (0.38–49.48); 0.236	
Tumour-infiltrating lymphocytes				
Absent	Reference			
Brisk	2.46 (0.52–11.59); 0.254			
Nonbrisk	1.93 (0.26–14.00); 0.518			
Tumour regression				
Absent	Reference			
Partial	0.32 (0.04–2.59); 0.287			
Associated melanocytic lesion				
Absent	Reference		Reference	
Present	0.57 (0.12–2.64); 0.468		1.47 (0.13–17.09); 0.757	
Surgical wait time (days)	1.00 (0.99–1.00); 0.581		0.95 (0.87–1.04); 0.241	

The multivariable models included variables with *P* < 0.10 in the univariable analysis.

*Statistically significant.

ALM:acral lentiginous melanoma; CI:confidence interval; HR:hazard ratio; LMM:lentigo maligna melanoma; NM:nodular melanoma; SSM:superficial spreading melanoma.

relevant and reached statistical significance (SHR 5.17; 95% CI, 1.02–26.20; *p*=0.048).

For DMFS (Table SII), age (continuous) remained significant (SHR 1.04; 95% CI, 1.01–1.06; *p*=0.005). TM, compared with MIS, showed a large effect size with wide CIs (SHR 6.13; 95% CI, 0.75–50.06; *p*=0.091). Breslow thickness ≥0.8 mm showed a borderline association with DMFS in univariable Fine–Gray analysis (SHR 8.02; 95% CI, 0.90–71.27; *p*=0.062) but did not reach statistical significance. Gray test demonstrated a borderline difference in DMFS between MIS and TM (*p*=0.0520) (Fig.S4).

For RFS (Table SII), age (continuous) was the only consistent predictor (SHR 1.03 per year; 95% CI, 1.01–1.06; *p*=0.013).

The 8 patients who developed distant metastasis are described in Table SIII. Primary tumours ranged from

MIS to Breslow thickness 1.0 mm; metastasis occurred 14–119 months after diagnosis; and metastatic sites included liver, bone, central nervous system, spleen, lung and lymph node. The occurrence of distant disease and melanoma-related death in patients with very thin lesions underscores the heterogeneity of risk even within MIS/TM.

DISCUSSION

In this cohort of patients with MIS and TM, adverse outcomes were not negligible despite restriction to early-stage disease. Across analytic approaches, older age and Breslow thickness ≥0.8 mm emerged as the most consistent prognostic determinants, underscoring clinically meaningful risk heterogeneity among lesions traditionally considered low risk.

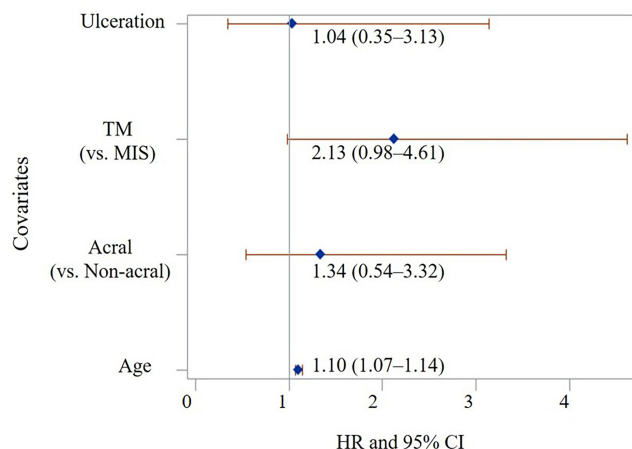
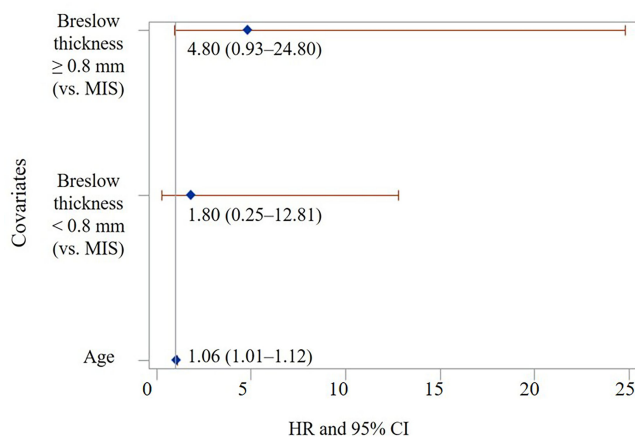
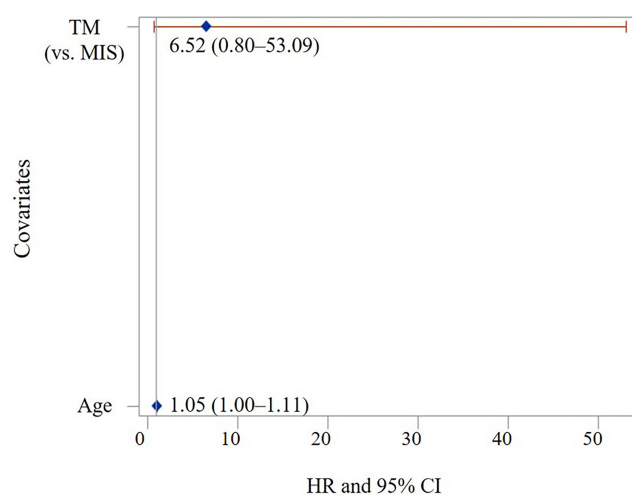
(A) OS**(B) MSS****(C) DMFS**

Fig. 1. Forest plots of multivariable Cox proportional hazards regression analyses for (A) overall survival, (B) melanoma-specific survival, (C) distant metastasis-free survival.

Our observed event rates exceeded those reported in Western reports. According to the AJCC 8th edition, 5-year MSS rates approach 99 % for T1a

and 97–99% for T1b tumours, reflecting the generally favourable outcomes of TMs in Western populations, where melanomas predominantly arise in intermittently sun-exposed sites such as the trunk (9, 25). In contrast, our acral-predominant cohort demonstrated a cumulative melanoma-specific mortality rate of 5.1 % over a mean follow-up of 81 months. This discrepancy suggests that prognostic patterns in acral-predominant Asian populations may differ from those reported in predominantly nonacral Western cohorts. However, neither ALM subtype nor acral location independently predicted increased mortality risk within our cohort, likely reflecting limited statistical power due to the small number of events, and these findings should therefore be interpreted cautiously (1, 6, 9, 15).

In our cohort, TM and MIS each accounted for half of the cases, with ALM representing the leading histologic subtype. Even patients with MIS experienced measurable adverse events, contrasting with the uniformly favourable prognosis often described in Western populations, where SSM predominates (1, 6, 9, 15). Prior studies from East Asia have similarly documented higher recurrence or mortality rates in localized acral melanoma, suggesting that anatomic site, delayed recognition, and distinct clinicopathologic features may exert greater influence on outcomes in this setting (6, 18, 23). The higher burden of acral disease, together with the older age distribution of our patients, likely contributes to the comparatively less favourable real-world outcomes observed in this Asian population (6, 26).

Breslow thickness ≥ 0.8 mm retained independent prognostic significance for melanoma-specific mortality after competing-risk adjustment, supporting its continued relevance as a clinically meaningful threshold in acral-predominant Asian populations (11, 12, 16). Our findings align with large Australian registry data demonstrating a substantially increased risk of death for tumours measuring 0.8–1.0 mm compared with those < 0.8 mm (16). These observations reinforce the global validity of the 0.8 mm cut-off adopted in the AJCC 8th edition (25) and extend its applicability to older Asian cohorts. Importantly, they highlight that submillimetre differences within the TM category remain clinically informative across diverse populations (16, 27).

The prognostic impact of age persisted across survival endpoints, including melanoma-specific survival and distant metastasis-free survival, even after accounting for competing mortality. This indicates that the association between older age and adverse outcomes cannot be explained solely by non-melanoma-related death. In our cohort, patients older than 60 years demonstrated a markedly increased risk of melanoma-specific death. While age is a recognised prognostic factor in Western populations, it often carries less relative weight than tumour thickness in large registry

analyses (27). In contrast, in our acral-predominant cohort, age emerged as the most consistent predictor across models, suggesting that host-related factors may exert a comparatively stronger influence in this clinical context.

Notably, several risk factors established in Western cohorts – including male sex, nonacral anatomic sites such as the head, neck, or posterior trunk, ulceration, and mitotic activity – did not demonstrate independent prognostic significance in our multivariable analysis (15, 16). This divergence likely reflects the distinct clinical composition of our cohort. The predominance of ALM and the limited number of nonacral cases reduced statistical power to detect associations for site-specific variables. These findings suggest that prognostic patterns derived from predominantly nonacral Western populations may not be directly transferable to acral-predominant Asian cohorts, particularly in early-stage disease (15, 16). In addition, differences in tumour thickness distribution may partly explain the less favourable outcomes observed in our cohort, as the mean Breslow thickness in our TM cohort (0.67 mm) was slightly higher than that reported in the Swedish population (0.6 mm) (9).

Taken together, these results challenge the assumption that MIS and TM uniformly confer minimal risk in Asian populations. Meaningful rates of distant metastasis and melanoma-specific mortality were observed even within the ≤ 1 mm category. Older age and Breslow thickness ≥ 0.8 mm consistently identified patients at higher risk, supporting consideration of risk-adapted surveillance strategies in this setting. Heightened vigilance in the evaluation of subtle acral lesions may also be warranted (10, 12). Future investigations integrating molecular and objective measures of tumour biology and host response may further refine individualized risk stratification, particularly given the occurrence of late distant events arising from very thin primaries (7, 28).

Limitations

This retrospective, single-centre design and modest sample size limit the generalisability and constrain subgroup precision. Molecular testing was incomplete, reflecting real-world practice and limiting assessment of genotype–prognosis relationships. Nonetheless, the prolonged observation window and uniform data capture provide clinically relevant, long-term outcomes for an underrepresented, acral-predominant Asian cohort.

Conclusion

In this acral-predominant Taiwanese cohort, MIS/TM were not uniformly low risk. Older age and Breslow thickness ≥ 0.8 mm independently identified patients at

increased risk, supporting risk-adapted surveillance in Asian populations.

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Data availability statement: The data that support the findings of this study are available from the corresponding author upon reasonable request.

Ethics declarations & trial registry information: This study was approved by the Research Ethics Committee of National Taiwan University Hospital (NTUH-REC No. 202503155RINA) and adhered to the Declaration of Helsinki. Informed consent was waived by the committee due to the retrospective nature of the study.

The authors have no conflicts of interest to declare.

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