

RESEARCH ARTICLE

Early platelet elevation after complete remission as a prognostic marker of favourable outcomes in favourable- and intermediate-risk acute myeloid leukaemia: A retrospective study

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Abstract

Objectives: Platelet (PLT) recovery after chemotherapy is associated with the prognosis of patients with acute myeloid leukaemia (AML). This study aimed to explore the prognostic significance of early high PLT values in patients with *de novo* non-M3 AML who achieved first complete remission (CR).

Methods: A total of 206 patients with *de novo* non-M3 AML were analysed in this retrospective study. A receiver operating characteristic (ROC) curve was used to determine the optimal PLT cut-off. The overall survival (OS) and relapse-free survival (RFS) were assessed using Kaplan-Meier and Cox regression analyses.

Results: $312 \times 10^9/L$ was confined as the cut-off of the PLT count. The estimated 3-year OS of patients with high PLT was higher than that of their counterparts (72.3% vs. 34.6%, $p = 0.001$). In subgroup analysis, patients with high PLT had better OS in the favourable- and intermediate-risk (non-adverse-risk) AML ($p = 0.001$). The estimated 3-year RFS for the high and low PLT groups was 75.1% and 45.7% respectively ($p = 0.078$). Multivariate analyses revealed that high PLT count was an independent favourable variable for OS (HR = 0.264, $p < 0.001$) and RFS (HR = 0.375, $p = 0.011$) in the non-adverse-risk group.

Conclusion: Our results showed that early high PLT count recovery at first CR in non-adverse-risk AML patients is a positive prognostic marker for survival outcomes.

KEYWORDS

acute myeloid leukaemia, outcome, overall survival, platelet, relapse-free survival

1 | INTRODUCTION

Acute myeloid leukaemia (AML) is a malignant bone marrow failure disease characterised by the clonal proliferation of bone marrow haematopoietic stem progenitor cells. In the USA, 4.3/100,000

individuals develop the disease every year, and the 5-year survival rate is only 24%.¹ Currently, chemotherapy is the primary AML treatment modality, and the risk stratification of patients underlies chemotherapy and further treatment regimens.²⁻⁵ With the application and development of next-generation sequencing technology, the

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risk stratification of AML mainly relies on cytogenetic and molecular genetic characteristics.⁵ However, the easily available clinical indicators (such as age, white blood cell count [WBC] and platelet [PLT]) are also associated with the prognosis of patients with AML.⁶⁻¹³

Complete remission (CR) occurs when bone marrow blasts are <5%, circulating blasts and the blasts with Auer rods are absent, no extramedullary disease is detected, neutrophil (ANC) count is $\geq 1.0 \times 10^9/L$, and platelet count is $\geq 100 \times 10^9/L$.¹⁴ When all conditions of CR are fulfilled but the PLT count is $< 100 \times 10^9/L$, the patient is considered to be in CR with incomplete platelet recovery (CRp). Since the concept of CRp was first proposed by the International Working Group in 2003,¹⁵ several studies have evaluated the correlation between PLT count and AML prognosis. In 2005, Larson et al. conducted a study on the recurrence rate of patients experiencing CRp, which was significantly higher than that of patients who experienced CR.¹⁶ CR with an ANC count $< 1.0 \times 10^9/L$ and PLT count $< 100 \times 10^9/L$ was defined as CR with incomplete haematologic recovery (CRI).¹⁴ In 2017, a retrospective study revealed that CRp or CRI decreased the overall survival (OS) and increased the risk of recurrence compared to CR in AML.⁸ At present, it has been recognised that the prognosis of CR is better than CRI in AML.

However, no consensus has been reached regarding the correlation between patients with complete PLT recovery and prognosis. In this retrospective study, we collected newly diagnosed non-M3 AML patients to explore the importance of complete PLT recovery after induction chemotherapy on prognosis.

2 | MATERIALS AND METHODS

2.1 | Data collection

This retrospective single-centre study included *de novo* non-M3 AML patients treated at the Second Hospital of Shanxi Medical University from January 2014 to December 2020. The clinical data of the patients were collected from hospital medical records. All procedures involving the clinical characteristics, diagnosis and treatment of the patients followed the Declaration of Helsinki.

2.2 | Patients

This study included patients who were (1) *de novo* AML patients treated at the Second Hospital of Shanxi Medical University from January 2014 to December 2020, (2) achieved CR after chemotherapy and (3) administered at least one course of consolidation chemotherapy.

The exclusion criteria were as follows: (1) acute promyelocytic leukaemia (M3), (2) patients who had previously received chemotherapy and (3) patients who were refractory.

All patients with AML were diagnosed and classified based on morphology, immunology, cytogenetics and molecular biology according to the 2016 World Health Organization (WHO) criteria.¹⁷

Cytogenetic and molecular biology subgroups were classified according to the National Comprehensive Cancer Network (NCCN) guidelines version 1, 2017. Patients younger than 60 years old received the traditional '7+3' chemotherapy regimen, and patients older than 60 years old received priming chemotherapy, all patients with or without demethylation therapy. Of the 26 patients who did not achieve CR after the first induction chemotherapy, 5 received the second '7+3' induction chemotherapy, and 21 patients received second-line chemotherapy regimens. Finally, in 206 patients who achieved CR after induction or reinduction chemotherapy, the choice of consolidation chemotherapy regimens was based on the NCCN guidelines.⁵

Relapse-free survival (RFS) was defined as the time from the date of CR to the date of the first event (relapse or last follow-up) occurrence or death. OS was defined as the time from the date of diagnosis to the date of death or the last follow-up. Relapse was considered to have occurred when the bone marrow contained $\geq 5\%$ blasts, reappearance of blasts in the blood or the emergence of extramedullary leukaemia.

The highest PLT count 28–42 days after chemotherapy was documented and defined as an early PLT count (D28 PLT). Follow-up was conducted through medical record review or telephone, and the follow-up time was up to 1 March 2021.

2.3 | Statistical analysis

Receiver operating characteristic (ROC) curve analysis was used to determine the optimal platelet cut-off value. The patients' baseline characteristics were compared using the Mann-Whitney *U* test for continuous parameters and the chi-square test or Fisher's exact test for categorical parameters. OS and RFS were analysed using the Kaplan-Meier method, and differences between groups were assessed using the log-rank test. Univariate Cox regression analysis ($p < 0.10$) was included in the multivariate Cox regression analysis, in which $p < 0.05$ was considered statistically significant. All statistical analyses were performed using SPSS software (version 20.0; IBM, Armonk, NY, USA).

3 | RESULTS

3.1 | Patients' characteristics

A total of 206 patients were enrolled in the current study (Figure 1), and the median follow-up time was 13 months (range, 3–87 months). The median age of the patients was 46 (range: 12–78) years at the time of diagnosis, and the cohort comprised 113 (54.85%) males and 93 (45.15%) females. The 2016 WHO subtype classification for AML patients was as follows: 114 (55.34%) patients were in the recurrent genetic abnormalities group, nine (4.37%) patients were in the myelodysplasia-related changes group, 76 (36.89%) patients were in the not-otherwise specified groups, and seven (3.40%)

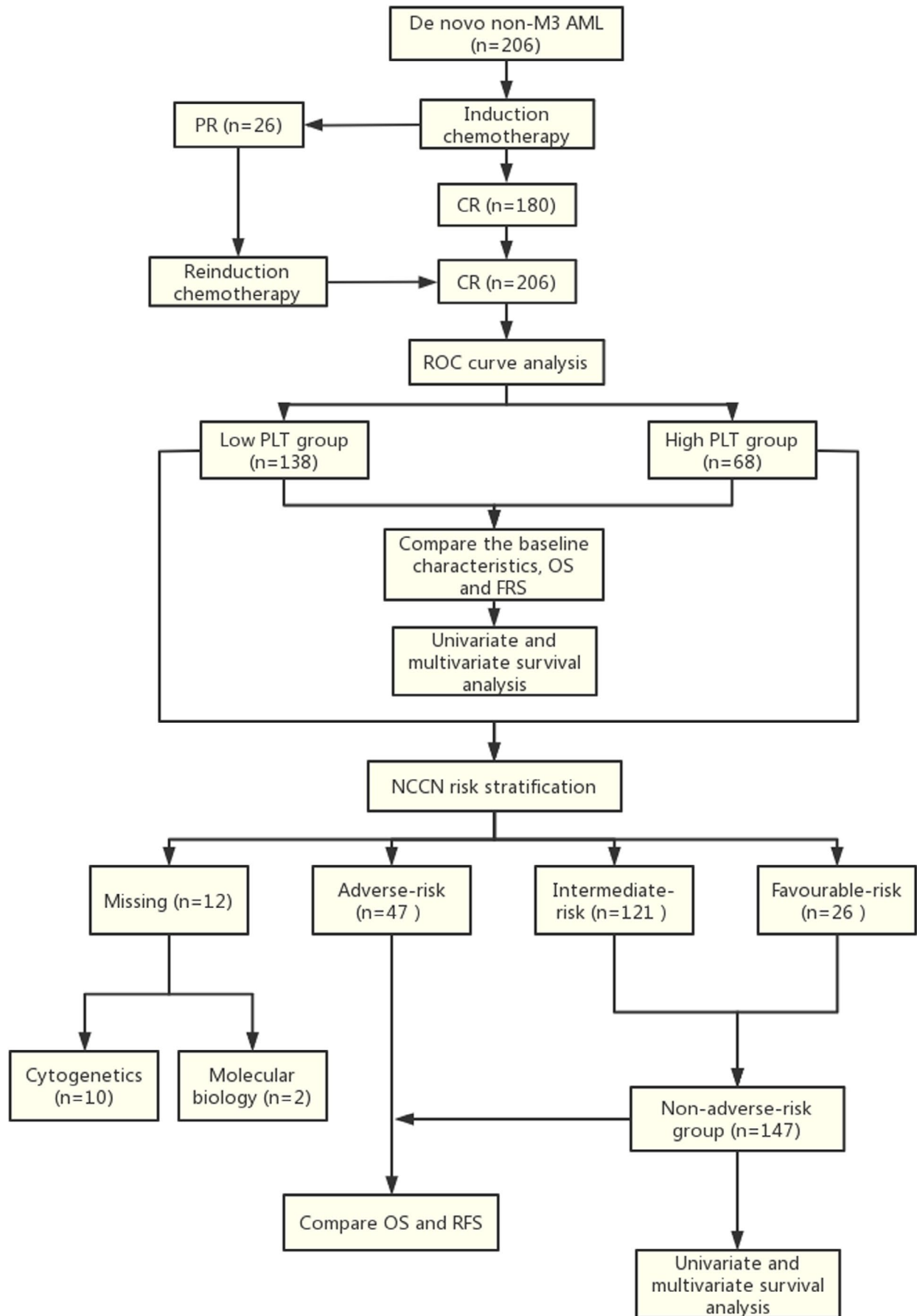


FIGURE 1 Study flow. M3, acute promyelocytic leukaemia; AML, acute myeloid leukaemia; PR, partial remission; CR, complete remission; ROC, receiver operating characteristic; PLT, platelet; OS, overall survival; RFS, relapse-free survival; NCCN, National Comprehensive Cancer Network

TABLE 1 Baseline characteristics of 206 patients with AML

Characteristics	All patients (n = 206)	Low PLT group	High PLT group	p
		100 ≤ PLT < 312 × 10 ⁹ /L (n = 138)	PLT ≥ 312 × 10 ⁹ /L (n = 68)	
Characteristics				
Median age (range), years	46 (12–78)	48(12–78)	44.5(12–68)	0.094
Gender				
Male, No. (%)	113 (54.85)	73 (52.90)	40 (58.82)	0.422
Female, No. (%)	93 (45.15)	65 (47.10)	28 (41.18)	
2016 WHO subtypes, No. (%)				
AML with recurrent genetic abnormalities	114 (55.34)	67 (48.55)	47 (69.12)	0.020
AML with myelodysplasia-related changes	9 (4.37)	7 (5.07)	2 (2.94)	
AML, not-otherwise specified	76 (36.89)	60 (43.48)	16 (23.53)	
Others	7 (3.40)	4 (2.90)	3 (4.41)	
Risk group, No. (%)				
Favourable	26 (12.62)	14 (10.15)	12 (17.65)	0.084
Intermediate	121 (58.74)	86 (62.32)	35 (51.47)	
Adverse	47 (22.81)	33 (23.91)	14 (20.59)	
Missing	12 (5.83)	5 (3.62)	7 (10.29)	
Initial infection, No. (%)				
Yes	133 (64.56)	87 (63.04)	46 (67.65)	0.516
NO	73 (35.44)	51 (36.96)	22 (32.35)	
Induction therapy with demethylation, No. (%)				
Yes	102 (49.51)	74 (53.62)	28 (41.18)	0.093
NO	104 (50.49)	64 (46.38)	40 (58.82)	
MRD < 5%, No. (%)				
Yes	185 (89.81)	124 (89.86)	61 (89.71)	0.973
NO	21 (10.19)	14 (10.14)	7 (10.29)	
Allogenic SCT, No. (%)				
Yes	24 (11.65)	15 (10.87)	9 (13.23)	0.619
NO	182 (88.35)	123 (89.13)	59 (86.76)	
Initial CBC				
Median WBC (range), 10 ⁹ /L	11.49 (0.51–274.58)	15.53 (0.80–274.58)	10.30 (0.51–185.30)	0.856
Median Hb (range), g/L	81.50 (27.00–157.20)	80.00 (27.00–157.20)	84.00 (40.00–148.00)	0.547
Median PLT (range), 10 ⁹ /L	33.00 (1.00–437.00)	32.50 (4.00–437.00)	34.00 (1.00–226.00)	0.761
Initial median MCV (range), fl	98.40 (30.90–128.90)	98.30 (30.90–121.90)	99.70 (77.10–128.90)	0.893
Initial median BMI (range), kg/m ²	23.47 (16.41–42.86)	23.84 (16.41–42.86)	22.95 (16.80–28.25)	0.715
D28 CBC after chemotherapy				
ANC, 10 ⁹ /L	2.59 (0.15–24.70)	2.55 (0.15–8.74)	2.85 (0.66–24.70)	0.007
Hb, g/L	111.00 (11.00–159.10)	114.00(11.00–159.10)	108.50 (68.00–141.00)	0.018
PLT, 10 ⁹ /L	255.00(100–695)	204.95 (100–311)	420.70 (313–695)	< 0.001

Abbreviations: 95%CI, 95% confidence interval; AML, acute myeloid leukaemia; ANC, neutrophil; BM, bone marrow; BMI, body mass index; CBC, cell blood count; Hb, haemoglobin; HR, hazard ratio; MCV, mean red blood cell volume; MRD, minimal residual disease; OS, overall survival; PLT, platelet; RFS, relapse-free survival; SCT, stem cell transplantation; WBC, white blood cell.

patients were in the other group. Based on cytogenetic and molecular genetic risk stratification, the favourable-risk group consisted of 26 (12.62%) patients; the intermediate-risk and adverse-risk

groups consisted of 121 (58.74%) and 47 (22.81%) patients, respectively, and the remaining 12 (5.83%) patients failed to be classified from records. A total of 133 (64.56%) patients were infected at the

time of diagnosis, 102 (49.51%) patients received demethylated drug induction chemotherapy, and 185 (89.81%) had minimal residual disease (MRD) <5% after induction chemotherapy. The initial median WBC count, haemoglobin (Hb) count, PLT, and mean red blood cell volume (MCV) were 11.49 (range: 0.51 – 274.58) $\times 10^9/L$, 81.50 (range: 27.00 – 157.20) g/L, 33 (range: 1.00 – 437.00) $\times 10^9/L$ and 98.40 (range: 30.90 – 128.90) fl respectively. The median body mass index (BMI) was 23.47 (range: 16.41 – 42.86) kg/m^2 . Blood cell analysis was performed after chemotherapy, in which the median ANC count was 2.59 (range: 0.15 – 24.70) $\times 10^9/L$, Hb count was 111.00 (range: 11.00 – 159.10) g/L, and PLT count was 255 (range: 100 – 695) $\times 10^9/L$. There were 24 (11.65%) patients who received allogeneic stem cell transplantation (SCT), the adverse-risk group, intermediate-risk group and favourable-risk group accounted for five, 16 and one respectively, while two patients were missing. This patient received allogeneic SCT in the favourable-risk group because of recurrence. A total of 83 deaths were recorded until 1 March 2021, and the estimated 3-year OS and RFS were 48.4% (95% confidence interval [CI]: 39.4–57.4%) and 59.6% (95% CI: 49.4–69.8%) respectively.

According to the ROC curve, $312 \times 10^9/L$ was confined as the cut-off value for PLT; hence, we divided the patients into high ($\geq 312 \times 10^9/L$) and low ($100 \leq PLT < 312 \times 10^9/L$). Subsequently, 138 patients had high D28 PLT counts, and 68 patients showed low D28 PLT counts. There were significant differences in the 2016 WHO subtypes and blood cell counts after chemotherapy between the two groups ($p < 0.05$). The clinical characteristics of the two groups of patients are summarised in Table 1.

3.2 | Prognostic value of D28 PLT

Patients with a high PLT count had a significantly superior RFS ($p = 0.020$, Figure 2A) and OS ($p < 0.001$, Figure 2B) compared with those with low PLT count, as assessed by the Kaplan-Meier test. The estimated 3-year OS of patients with high PLT was higher than that of their counterparts (72.3% vs. 34.6%, $p = 0.001$). The estimated 3-year RFS for the high and low PLT groups was 75.1% and 45.7% respectively ($p = 0.078$).

Next, we performed a univariate analysis for survival outcomes of 206 patients with AML with respect to age, sex, initial blood count, infection, BMI, induction chemotherapy with demethylation, risk group, MRD, allogeneic SCT and D28 blood count. The results are summarised in Table 2. We found that D28 PLT was significantly associated with RFS ($p = 0.017$) and OS ($p = 0.001$), and the MRD and allogeneic SCT were also correlated with RFS and OS. In the multivariate analysis, the results showed that D28 high PLT count was an independent predictor of better RFS (HR = 0.436, $p = 0.009$) and OS (HR = 0.386, $p < 0.001$), MRD < 5%, and allogeneic SCT was a favourable predictor of RFS and OS ($p < 0.05$), and D28 Hb ≥ 100 g/L was associated with superior RFS ($p < 0.05$).

To further analyse the prognostic significance of D28 PLT count in various cytogenetic and molecular genetic risk subgroups, we

divided the patients into adverse-risk and non-adverse-risk group (favourable- and intermediate-risk groups). In subgroup analysis, patients with high PLT had better OS in the non-adverse-risk group ($p = 0.001$), but not in the adverse-risk group ($p = 0.695$) (Figure 3B). Patients with high PLT also showed a better RFS trend ($p = 0.076$); however, this trend was not observed in patients in the adverse-risk group ($p = 0.450$) (Figure 3A). We also conducted univariate and multivariate survival analyses for non-adverse-risk patients, and the results are summarised in Table 3. Univariate analysis showed that age, induction chemotherapy with demethylation, MRD, D28 Hb and D28 PLT were significantly associated with RFS and OS, D28 ANC and allogeneic SCT were correlated with OS. Furthermore, D28 PLT $\geq 312 \times 10^9/L$ was a positive predictive marker of improved RFS (HR = 0.375, $p = 0.011$) and OS (HR = 0.264, $p < 0.001$) in multivariate analysis. Age, induction chemotherapy with demethylation,

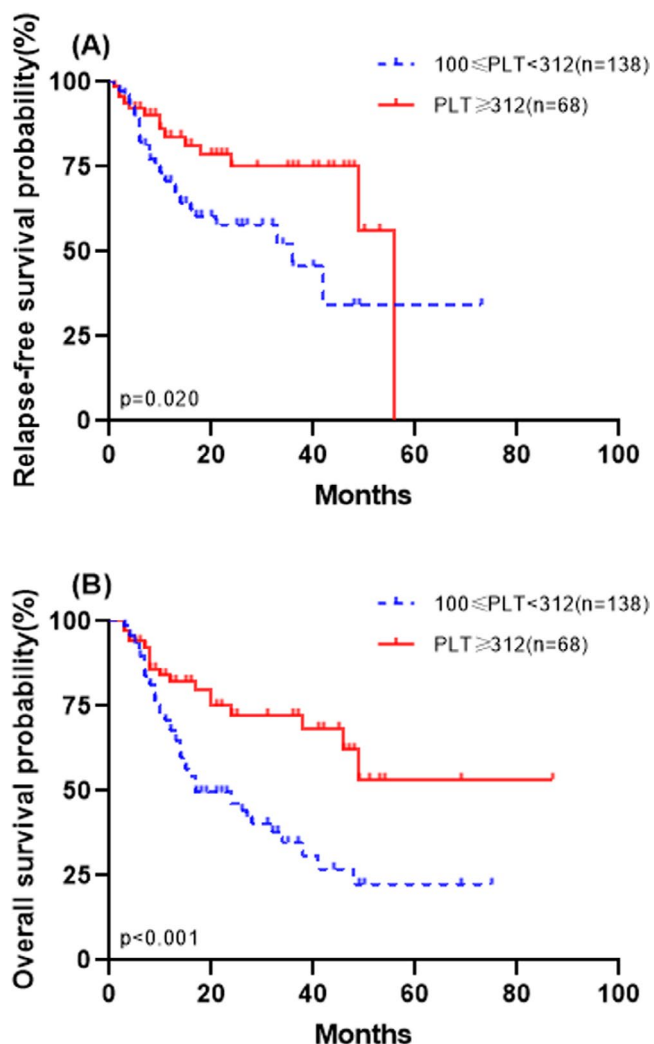


FIGURE 2 Survival outcomes analysis of patients with de novo non-M3 AML after CR according to PLT count. (A) Relapse-free survival was compared between low PLT group and high PLT group. (B) Overall survival was compared between low PLT group and high PLT group

TABLE 2 Univariate and multivariate analysis of 206 patients with AML for RFS and OS

Characteristics	RFS			OS		
	Univariate		Multivariate	Univariate		Multivariate
	HR (95%CI)	p	HR (95%CI)	HR (95%CI)	p	HR (95%CI)
Age(years, ≥ 60 vs. < 60)	1.436 (0.726–2.841)	0.298	-	1.320 (0.745–2.342)	0.342	-
Gender (male vs. female)	0.757 (0.419–1.366)	0.355	-	1.072 (0.656–1.754)	0.781	-
Initial WBC ($10^9/L$, ≥ 100 vs. < 100)	1.002 (0.349–2.875)	0.997	-	1.199 (0.554–2.593)	0.645	-
Initial Hb (g/L, ≥ 100 vs. < 100)	0.957 (0.499–1.838)	0.896	-	0.728 (0.409–1.297)	0.281	-
Initial PLT ($10^9/L$, 100–300 vs. < 100 or > 300)	1.181 (0.508–2.745)	0.698	-	1.046 (0.491–2.227)	0.907	-
Initial MCV (fl, 80–100 vs. > 100 or < 80)	1.414 (0.786–2.541)	0.247	-	1.094 (0.688–1.740)	0.704	-
Initial BMI (kg/m^2 , 18.5–23.9 vs. < 18.5 or > 23.9)	0.625 (0.346–1.132)	0.121	-	1.046 (0.659–1.661)	0.849	-
Initial infection (Yes vs. No)	0.907 (0.471–1.746)	0.770	-	1.441 (0.804–2.581)	0.220	-
Induction chemotherapy with demethylation (Yes vs. No)	0.883 (0.469–1.661)	0.699	-	0.898 (0.545–1.481)	0.675	-
Risk group (adverse vs. others)	1.146 (0.570–2.303)	0.702	-	1.312 (0.772–2.230)	0.315	-
MRD $< 5\%$ (Yes vs. No)	0.245 (0.114–0.523)	< 0.001	0.281 (0.137–0.574)	0.486 (0.251–0.943)	0.033	0.456 (0.243–0.855)
Allogenic SCT (Yes vs. No)	0.231 (0.054–0.978)	0.047	0.225 (0.054–0.936)	0.380 (0.149–0.972)	0.043	0.378 (0.151–0.946)
D28 ANC ($10^9/L$, 2–7 vs. < 2 or > 7)	1.162 (0.639–2.113)	0.623	-	1.273 (0.787–2.060)	0.326	-
D28 Hb (g/L, ≥ 100 vs. < 100)	0.568 (0.307–1.052)	0.072	0.540 (0.304–0.960)	0.652 (0.392–1.083)	0.098	0.643 (0.402–1.027)
D28 PLT ($10^9/L$, ≥ 312 vs. $100 \leq$ PLT < 312)	0.452 (0.235–0.870)	0.017	0.436 (0.234–0.811)	0.390 (0.225–0.675)	0.001	0.386 (0.227–0.657)

Abbreviations: 95%CI, 95% confidence interval; AML, acute myeloid leukaemia; ANC, neutrophil; BMI, body mass index; Hb, haemoglobin; HR, hazard ratio; MCV, mean red blood cell volume; MRD, minimal residual disease; OS, overall survival; PLT, platelet; RFS, relapse-free survival; SCT, stem cell transplantation; WBC, white blood cell.

MRD < 5% and D28 Hb ≥ 100 g/L were deemed predictive factors for FRS and OS ($p < 0.05$).

4 | DISCUSSION

Our study aimed to explore the prognostic significance of early PLT recovery in patients with *de novo* non-M3 AML who achieved CR. The results showed that the estimated 3-year OS of patients with high PLT was higher than that of their counterparts (72.3% vs. 34.6%, $p = 0.001$). The estimated 3-year RFS for the high and low PLT groups was 75.1% and 45.7% respectively ($p = 0.078$). Incorporating multiple clinical features for survival regression analysis showed that early high PLT count recovery $\geq 312 \times 10^9/L$ significantly prolonged OS and reduced the risk of recurrence in all patients, which is consistent with the research of Yamazaki et al. The study evaluated the blood count on day 28 and found that patients with $PLT \geq 500 \times 10^9/L$ and $Hb \geq 9$ g/dL were strong predictors with better RFS ($p = 0.009$, $p = 0.012$), while the PLT count $\geq 350 \times 10^9/L$ was an independent predictive factor for improved RFS ($p = 0.020$).¹⁰ Yanada et al. reported that $PLT \geq 320 \times 10^9/L$ were significantly associated with improved RFS at CR in AML patients.¹⁸ A retrospective analysis showed that patients with relatively early PLT recovery (from the beginning of induction chemotherapy to the first 25 days PLT count $\geq 20 \times 10^9/L$ for ≥ 3 days) have a longer 5-year OS (62% vs. 23%, $p < 0.001$) and disease-free survival (DFS) (57% vs. 15%, $p < 0.001$) rates than patients with late PLT recovery.⁶ Herein, we defined D28 PLT recovery as early, which further demonstrated that early bone marrow recovery was associated with prolonged OS and RFS. The above study showed that early recovery of high PLT count is strongly correlated with favourable prognosis in AML patients.

However, none of the previous studies ruled out the influence of cytogenetic and molecular biologic characteristics on survival prognosis. After risk stratification, our study included multiple clinical indicators to analyse the survival influence of platelets in patients with AML. The results showed that D28 $PLT \geq 312 \times 10^9/L$ was also a positive prognostic marker of RFS (HR = 0.375, $p = 0.011$) and OS (HR = 0.264, $p < 0.001$) in non-adverse-risk patients. Moreover, we also found that $Hb \geq 100$ g/L was a positive prognostic factor of better RFS and OS in non-adverse-risk AML patients. The findings also emphasised that if PLT recovery was early, the count was high, the bone marrow haematopoietic function was strong, and the patients' long-term survival was improved.

Some studies have shown that infection affects the secretion of thrombopoietin (TPO) and interleukin-6 (IL-6) in liver cells, which in turn alters the PLT count.^{19,20} However, in this study, no correlation was established between infection at the time of diagnosis and PLT count. In the survival analysis, infection did not show statistical significance in predicting survival prognosis. Mangaonkar et al.²¹ reached similar conclusions with respect to the association of infection during induction chemotherapy and PLT count to prognosis. Reportedly, decitabine improves PLT recovery by downregulating the IL-8 level in AML.²² However, no statistically significant difference

was detected in PLT irrespective of the demethylation drugs administered to patients. This phenomenon can be attributed to inconsistent chemotherapy regimens or small patient samples. Age is one of the most important prognostic factors for AML.⁵ Typically, elderly AML patients have a low long-term survival rate, many comorbidities and poor efficacy.^{23,24} The present study is consistent with this in non-adverse-risk patients; however, it has not yet been confirmed as an influencing factor of OS and RFS in all patients, which might be related to the fact that most of the patients with adverse risk treated in this study were individuals aged < 60 years old (40/47).

The current study has the following limitations: (1) It is a single-centre retrospective study; (2) the reasons for the correlation between PLT count and the prognosis of AML lack basic research, thereby requiring additional prospective multicentre clinical trials.

In conclusion, the current study demonstrated that early high PLT count in non-adverse-risk AML patients showed strong bone marrow haematopoietic recovery, which is a positive prognostic marker for survival outcomes. PLT count is a simple and easily available indicator of peripheral blood cells in AML patients and thus can serve as a convenient reference index for prognostic evaluation

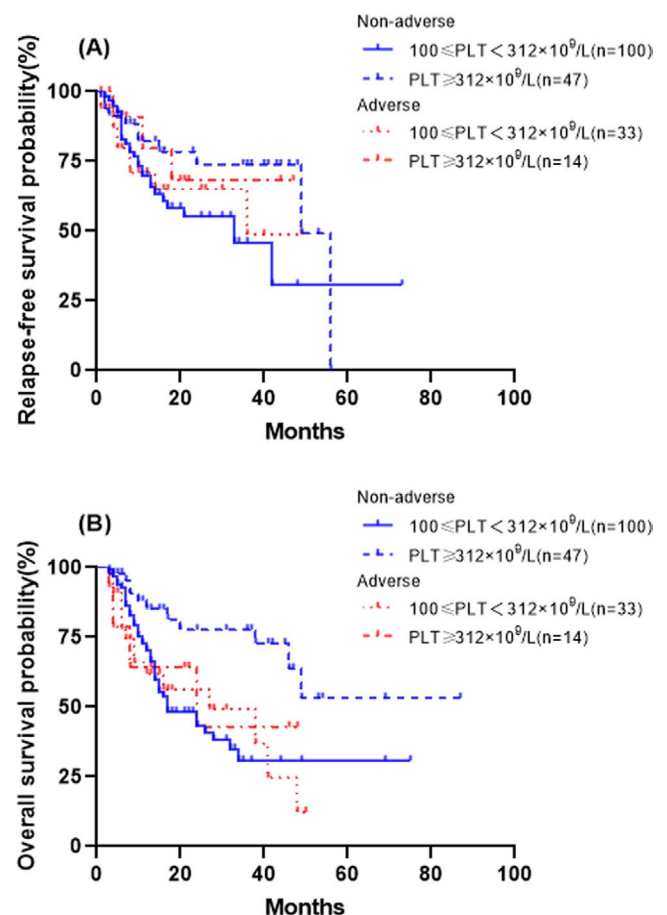


FIGURE 3 Survival outcomes analysis of patients with *de novo* non-M3 AML after CR according to PLT count. (A) Relapse-free survival was compared between non-adverse-risk group and adverse-risk group. (B) Overall survival was compared between non-adverse-risk group and adverse-risk group

TABLE 3 Univariate and multivariate analysis of non-adverse-risk patients with AML for RFS and OS

Characteristics	RFS			OS		
	Univariate		Multivariate	Univariate		Multivariate
	HR (95%CI)	p	HR (95%CI)	HR (95%CI)	p	p
Age(years, ≥60 vs. <60)	2.237 (1.003–4.989)	0.049	2.289 (1.066–4.915)	2.244 (1.128–4.465)	0.021	2.188 (1.137–4.209)
Gender (male vs. female)	0.792 (0.395–1.587)	0.510	-	0.937 (0.520–1.685)	0.827	-
Initial WBC (10 ⁹ /L, ≥100 vs. <100)	1.213 (0.266–5.531)	0.803	-	1.810 (0.582–5.630)	0.305	-
Initial Hb (g/L, ≥100 vs. <100)	1.167 (0.544–2.501)	0.692	-	0.738 (0.349–1.560)	0.426	-
Initial PLT (10 ⁹ /L, 100–300 vs. <100 or >300)	0.836 (0.258–2.714)	0.766	-	0.909 (0.320–2.579)	0.858	-
Initial MCV (fl, 80–100 vs. >100 or <80)	1.607 (0.814–3.175)	0.172	-	1.228 (0.694–2.173)	0.481	-
Initial BMI (kg/m ² , 18.5–23.9 vs. <18.5 or >23.9)	0.484 (0.225–1.044)	0.064	0.524 (0.253–1.083)	1.001 (0.554–1.809)	0.998	-
Initial infection (Yes vs. No)	0.856 (0.407–1.804)	0.683	-	1.233 (0.639–2.379)	0.532	-
Induction chemotherapy with demethylation (Yes vs. No)	0.402 (0.181–0.894)	0.025	0.447 (0.209–0.956)	0.485 (0.254–0.925)	0.028	0.497 (0.268–0.919)
MRD<5% (Yes vs. No)	0.233 (0.097–0.562)	0.001	0.246 (0.106–0.574)	0.336 (0.156–0.727)	0.006	0.313 (0.152–0.646)
Allogenic SCT (Yes vs. No)	0.272 (0.061–1.202)	0.086	0.310 (0.072–1.338)	0.266 (0.075–0.950)	0.041	0.326 (0.098–1.081)
D28 ANC (10 ⁹ /L, 2–7 vs. <2 or >7)	1.336 (0.667–2.677)	0.413	-	1.944 (1.054–3.583)	0.033	1.854 (1.023–3.361)
D28 Hb (g/L, ≥100 vs. <100)	0.326 (0.161–0.657)	0.002	0.361 (0.187–0.700)	0.447 (0.249–0.801)	0.007	0.444 (0.252–0.783)
D28 PLT (10 ⁹ /L, ≥312 vs. 100 ≤ PLT <312)	0.346 (0.159–0.754)	0.008	0.375 (0.176–0.798)	0.263 (0.130–0.533)	<0.001	0.264 (0.133–0.526)

Abbreviations: 95%CI, 95% confidence interval; AML, acute myeloid leukaemia; ANC, neutrophil; BMI, body mass index; Hb, haemoglobin; HR, hazard ratio; MCV, mean red blood cell volume; MRD, minimal residual disease; OS, overall survival; PLT, platelet; RFS, relapse-free survival; SCT, stem cell transplantation; WBC, white blood cell.

and treatment options. Nonetheless, additional prospective, large-sample studies are also needed to deeply explore the correlation between PLT count and prognosis of AML patients, and the reason for the elevated PLT.

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RJZ designed the study and reviewed the article. XLW designed the study, collected and analysed the data, and wrote the article. RQL, JHZ, SXY, YZW, ZFX and YHT collected the data. XLZ and LHY reviewed the article.

CONFLICT OF INTEREST

The authors declare that there are no competing interests.

DATA AVAILABILITY STATEMENT

The data used to support the findings of this study are available from the corresponding author upon request.

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