

Development and validation of survival prediction nomograms for patients with early-stage rectal cancer: a population-based study

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Background: The incidence of colorectal cancer (CRC) has been rising in recent years, with a concurrent increase in early-stage rectal cancer (ESRC) cases. This study aimed to investigate risk factors and developed nomograms to predict overall survival (OS) and cancer-specific survival (CSS) in ESRC patients in order to improve clinical outcomes across diverse patient subgroups.

Methods: Risk factors were investigated in ESRC patients by analyzing data from the Surveillance, Epidemiology, and End Results (SEER) database. We developed and validated nomograms to predict OS and CSS after dividing patients into two risk groups. Then we assessed the potential benefits of various therapies across subgroups after propensity score-matching (PSM).

Results: T stage, tumor grade, age, carcinoembryonic antigen (CEA) levels, tumor size, and surgical options emerged as independent risk factors through univariate and multivariate Cox regression analyses, contributing to the OS nomogram; while for CSS, the identified risk factors were tumor grade, age, elevated CEA levels and surgical options. The Concordance-index of the nomogram surpassed that of the American Joint Committee on Cancer (AJCC) 7th staging system, with values of 0.69 (C-index, 0.64–0.74) in the training set and 0.65 (C-index, 0.62–0.68) in the testing set. The receiver operating characteristic (ROC) analysis revealed area under the curve (AUC) values of 0.70, 0.70, and 0.67 for 1-, 3-, and 5-year OS in the development cohort, with comparable results in the validation cohort. Calibration plots demonstrated strong alignment between predicted and observed outcomes. Decision curve analysis (DCA) confirmed the nomogram's superior clinical utility relative to the AJCC 7th staging system, with similar findings for CSS. Kaplan-Meier curves illustrated significant differences in OS and CSS between low- and high-risk groups. Notably, radiation and chemotherapy conferred no benefit, while low-risk patients, especially younger individuals, may benefit from local resection.

Conclusions: This study presents a comprehensive prognostic analysis of patients with ESRC and developed predictive nomograms for OS and CSS. Subgroup analyses highlight the potential benefits of local resection in younger patients with low risk.

Keywords: Early-stage rectal cancer (ESRC); radiation; chemotherapy; local resection; prognosis

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Introduction

The colorectal cancer (CRC) is the second leading cause of cancer-related mortality and the third most prevalent cancer worldwide. In 2022, it accounted for over 1.8 million new diagnoses and 881,899 deaths (1). Advances in imaging technology have led to a marked rise in the detection of early-stage rectal cancer (ESRC) in recent decades, including tumors classified as T1-2N0M0 (2). Although 5-year survival rates for ESRC have shown improvement, there has been a marked increase in ESRC incidence, particularly in individuals younger than 50 years (3). For patients with ESRC, radical resection is recognized as the gold standard for treatment option, significantly reducing the risk of local recurrence and metastasis (4). However, advancements in endoscopic techniques have led to widespread recognition of local resection via endoscopy (5). Therefore, radical resection may not be the optimal treatment for patients with ESRC. Radical resection may lead to loss of organ function, surgical complications, and short-term low anterior resection syndrome (LARS) in the rectum. Furthermore, 75% of patients reported different degrees of gastrointestinal and urinary dysfunction after the procedure (6). One study indicated that adjuvant chemoradiotherapy in conjunction with local excision effectively eliminated occult metastatic lymph nodes in the rectal mesentery and residual cancer cells in the rectal wall, thereby significantly reducing the risk of tumor recurrence in T1 stage rectal cancer (7). A meta-analysis revealed that

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Key findings

 The developed nomograms are capable to predict overall survival (OS) and cancer-specific survival (CSS) of early-stage rectal cancer (ESRC) patients. Radiation and chemotherapy showed no benefit, while low-risk patients, especially younger individuals, may benefit from local resection.

What is known and what is new?

- The incidence of ESRC is increasing with few relative prognostic studies.
- Based on a large population-level data, and we developed well-performing nomograms to predict OS and CSS after dividing patients into two risk groups. Then we assessed the potential benefits of various therapies across subgroups after propensity score-matching.

What is the implication, and what should change now?

 Identifying high-risk patients with ESRC and make targeted medical management may improve outcomes of this population. neoadjuvant chemotherapy offered several advantages for locally advanced rectal cancer (LARC) patients, including the eradication of occult micrometastases, reduced ostomy restoration time, improved treatment adherence, a significant increase in the pathological complete response (pCR) rate and enhanced R0 resection and tumor downstaging rates (8). Local resection alone may serve as an effective treatment option for patients with ESRC who lack poor prognostic pathology (9). However, it is crucial to recognize that the risk of lymph node metastasis following simple local excision can range from 12% to 28% in patients with intermediate-stage disease (10). Trans-abdominal resection is generally recommended for T2N0M0 rectal cancer, as local excision alone has been associated with local recurrence rates ranging from 11% to 45% in patients with T2 lesions (11). Chemoradiotherapy is regarded as the standard treatment for LARC, having demonstrated a reduction in disease recurrence and an improvement in overall survival (OS) outcomes (12). In addition, a study underscored the significant disease burden of ESRC, which exhibited distinct clinicopathological and molecular features compared to late-onset cases, including differences in tumor differentiation, tumorinfiltrating lymphocytes, and molecular phenotypes (13). The treatment strategies and outcomes also differ between ESRC and the late-onset rectal cancers (14). Nonetheless, the prognostic implications and effects of additional chemoradiotherapy in patients with ESRC continue to be debated (11,15). Moreover, population-based studies on prognosis and treatment options for patients with ESCC are limited, highlighting the significance to evaluate the function of chemoradiotherapy in this cohort.

This study utilized data from the Surveillance, Epidemiology, and End Results (SEER) database to identify prognostic factors and develop nomograms for predicting OS and cancer-specific survival (CSS) in patients with ESRC. Furthermore, it aimed to evaluate treatment strategies across subgroups to identify the most effective therapeutic approaches for this population. We present this article in accordance with the TRIPOD reporting checklist (available at https://tcr.amegroups.com/article/view/10.21037/tcr-24-1888/rc).

Methods

Database source and patients selection

In the retrospective cohort study, clinicopathological data from ESRC patients in the SEER database was analyzed

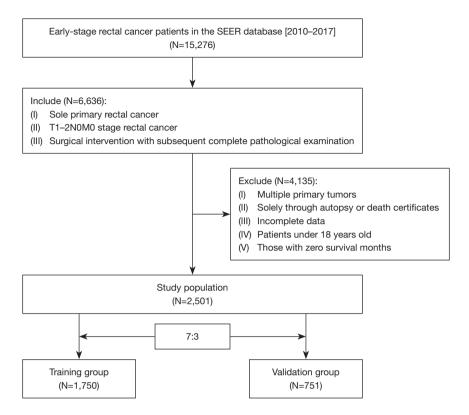


Figure 1 The selection process of patients in the research. SEER, Surveillance, Epidemiology, and End Results.

from January 1, 2010 to December 31, 2017 by use of SEER*Stat 8.4.0.1 software. Data information collected included age, race, tumor grade, sex, histological grade, the American Joint Committee on Cancer (AJCC) 7th stages (T, N, M), tumor size, regional nodes examined, treatment sequence (surgery before, after, or without chemoradiotherapy), preoperative carcinoembryonic antigen (CEA) levels, perineural invasion, and radiotherapy and chemotherapy information. The inclusion criteria were defined: (I) individuals with rectal cancer as their sole primary malignancy; (II) those diagnosed with T1-2N0M0 stage rectal cancer; and (III) patients who underwent surgical intervention with subsequent complete pathological examination. Exclusion criteria were as follows: (I) patients presenting with multiple primary tumors; (II) cases identified solely through autopsy or death certificate reports; (III) individuals with incomplete data on any of the inclusion parameters; (IV) patients aged under 18 years; and (V) those with zero survival months. The final cohort comprised 2,501 patients, stratified into a training set of 1,750 and a test set of 751 for validation of predictive nomograms (Figure 1). The primary outcomes

of interest were OS and CSS, with OS defined as the interval from diagnosis to the date of death from any cause or last follow-up, and CSS as the period from diagnosis to death attributed to cancer or last follow-up. This study utilized publicly available data from the SEER database, with all patient data de-identified prior to collection, thus exempting it from ethics committee approval or informed consent. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013).

Propensity score-matching (PSM)

To reduce bias and confounding factors in subgroups among ESRC patients, we employed PSM. The method accounted for age, sex, race, T stage, surgical options, tumor grade, tumor size, CEA level, perineural invasion, and chemoradiotherapy. We used the logistic regression to get propensity scores. Subgroups of ESRC patients were matched in a 1:1 ratio after PSM. We compared clinicopathological variables before and after PSM by use of the χ^2 test to evaluate the effectiveness of the matching method.

Statistics analysis

This study employed the Mann-Whitney U test to assess disparities in continuous variables, with categorical variables subjected to chi-squared analysis. Univariate and multivariate Cox regression analyses were performed to calculate odds ratios (ORs) and 95% confidence intervals (CIs). Prognostic variables identified as significant (P<0.05) from the multivariate Cox regression were integrated into a nomogram, facilitating a graphical representation for the estimation of 1-, 3-, and 5-year survival probabilities.

Hazard ratios (HRs) along with their 95% CIs were reported for all significant findings. The optimal threshold for tumor size was determined utilizing X-tile software (version 3.6.1, Yale University). Model performance was evaluated with the concordance index (C-index) and receiver operating characteristic (ROC) curves, with the area under the curve (AUC) being calculated. Calibration plots were constructed to compare predicted versus actual 1-, 3-, and 5-year survival outcomes. The clinical applicability of the predictive model was appraised through decision curve analysis (DCA). The study population was stratified into high- and low-risk categories on the basis of their total nomogram scores. Survival disparities, as measured by OS and CSS, were compared across subgroups using the Kaplan-Meier method complemented by the log-rank test. Statistical analysis was performed using the Statistical Package for the Social Sciences (SPSS) 22.0 (International Business Machines Corporation, Armonk, NY, USA) and R version 4.2.0.

Results

Basic characteristics of the patients

Our study comprised a total of 2,501 patients diagnosed with ESRC, with 1,750 patients designated to the training set and 751 to the validation set. The demographic and clinical characteristics of patients with ESRC in both the training and validation cohorts for OS were detailed in *Table 1*. The median age of the patient cohort was 63 years, with a median survival duration of 59 months. Notably, nearly half of the patients (43.14%) were younger than 60 years, and approximately one-fifth (21.99%) underwent local resection. Chemotherapy was administered to 575 patients (22.99%), while 576 patients (23.03%) received radiotherapy. Additionally, 393 patients (17.24%) underwent neoadjuvant therapy prior to surgery, and 151 patients (6.63%) received adjuvant therapy post-surgery. Statistical analyses, including the Mann-Whitney *U* and Chi-squared tests, revealed no

significant differences in feature distributions between the training and validation sets. The demographic and clinical characteristics for CSS in both cohorts yielded comparable results (see Table S1).

Prognostic factors for OS and CSS

To identify prognostic indicators for OS, we performed univariate and multivariate Cox regression analyses on the training set (*Table 2*). These analyses identified several risk factors associated with OS, including age at diagnosis, T stage, tumor grade, CEA level, and surgical options. Notably, T2 stage (HR: 1.23; 95% CI: 0.94–1.63), poor tumor grade (HR: 1.81; 95% CI: 1.25–2.62), older age (HR: 2.55; 95% CI: 1.93–3.36), elevated CEA levels (HR: 1.80; 95% CI: 1.40–2.32), and larger tumor size (HR: 1.41; 95% CI: 1.08–1.84) were all linked to worse OS outcomes. In contrast, radical resection (HR: 0.64; 95% CI: 0.48–0.85) was associated with improved OS prognosis.

Additionally, the risk factors for CSS mirrored those for OS (Table S2). Poor tumor grade (HR: 1.90; 95% CI: 1.13–3.18), older age (HR: 1.77; 95% CI: 1.20–2.61), and positive CEA levels (HR: 1.59; 95% CI: 1.07–2.36) were similarly correlated with inferior CSS outcomes, while radical resection (HR: 0.61; 95% CI: 0.39–0.97) was linked to improved CSS prognosis.

Development and validation of prognostic nomograms

Utilizing the multivariate analysis results from the training cohort, we constructed nomograms to predict OS and CSS in ESRC patients (Figure 2). Each variable was assigned a score ranging from 0 to 100, reflecting its contribution to model accuracy. By aggregating these scores for each patient, we calculated a total point value to estimate the probabilities of 1-, 3-, and 5-year OS and CSS. Importantly, higher scores were associated with poorer prognoses. In the training set, the nomogram for OS exhibited a C-index of 0.69 (95% CI: 0.64–0.74), surpassing the 7th edition of the AJCC staging system, which recorded a C-index of 0.60 (95% CI: 0.54-0.65). And validation in an independent testing set yielded a C-index of 0.65 (95% CI: 0.62-0.68). For CSS, the C-index was 0.68 (95% CI: 0.61-0.76) in the training set and 0.64 (95% CI: 0.59-0.68) in the validation set, both surpassing the 7th AJCC staging system.

The ROC analysis revealed AUC values of 0.70, 0.70, and 0.67 in the training set for 1-, 3-, and 5-year OS, respectively. In the validation set, the AUC values for

Table 1 Baseline characteristics of patients for overall survival in training and test sets

Characteristics	Total (N=2,501)	Training set (N=1,750)	Test set (N=751)	P value
Age				0.08
<60 years	1,079 (43.14)	735 (42.00)	344 (45.81)	
≥60 years	1,422 (56.86)	1,015 (58.00)	407 (54.19)	
Sex				0.10
Female	1,064 (42.54)	726 (41.49)	338 (45.01)	
Male	1,437 (57.46)	1,024 (58.51)	413 (54.99)	
Race				0.81
White	2,041 (81.61)	1,426 (81.49)	615 (81.89)	
Others	460 (18.39)	324 (18.51)	136 (18.11)	
Grade				0.67
I/II	2,300 (91.96)	1,612 (92.11)	688 (91.61)	
III/IV	201 (8.04)	138 (7.89)	63 (8.39)	
Histology				0.29
Adenocarcinoma	2,442 (97.64)	1,705 (97.43)	737 (98.14)	
Others	59 (2.36)	45 (2.57)	14 (1.86)	
T stage				0.37
T1	1,171 (46.82)	809 (46.23)	362 (48.20)	
T2	1,330 (53.18)	941 (53.77)	389 (51.80)	
Surgery options				0.14
Local resection	550 (21.99)	399 (22.80)	151 (20.11)	
Radical resection	1,951 (78.01)	1,351 (77.20)	600 (79.89)	
Radiation				0.92
No	1,925 (76.97)	1,346 (76.91)	579 (77.10)	
Yes	576 (23.03)	404 (23.09)	172 (22.90)	
Chemotherapy				0.97
No	1,926 (77.01)	1,348 (77.03)	578 (76.96)	
Yes	575 (22.99)	402 (22.97)	173 (23.04)	
Treatment sequence				0.88
Only surgery	1,925 (76.97)	1,347 (76.97)	578 (76.96)	
Adjuvant therapy	165 (6.60)	118 (6.74)	47 (6.26)	
Neoadjuvant therapy	411 (16.43)	285 (16.29)	126 (16.78)	
CEA				0.89
Negative	1,985 (79.37)	1,387 (79.26)	598 (79.63)	
Positive	516 (20.63)	363 (20.74)	153 (20.37)	
Perineural invasion				0.68
No	2,443 (97.68)	1,708 (97.60)	735 (97.87)	
Yes	58 (2.32)	42 (2.40)	16 (2.13)	
Tumor size				0.85
<18 mm	889 (35.55)	620 (35.43)	269 (35.82)	
≥18 mm	1,612 (64.45)	1,130 (64.57)	482 (64.18)	

Data are presented as n (%). CEA, carcinoembryonic antigen.

Table 2 Univariate and multivariate Cox analysis for overall survival of patients in the training set

Characteristics -	Univariate analysis		Multivariate analysis	
	Hazard ratio (95% CI)	P value	Hazard ratio (95% CI)	P value
Age (years)		<0.001*		<0.001*
<60	Ref		Ref	
≥60	2.72 (2.07–3.57)		2.55 (1.93–3.36)	
Sex		0.82		
Female	Ref			
Male	0.97 (0.77–1.23)			
Race		0.07		
Others	Ref			
White	1.37 (0.98–1.90)			
Grade		0.002*		<0.001*
1/11	Ref		Ref	
III/IV	1.77 (1.23–2.54)		1.81 (1.25–2.62)	
Histology		0.01*		0.21
Others	Ref		Ref	
Adenocarcinoma	0.50 (0.28–0.86)		0.70 (0.39–1.23)	
T stage		0.02*		0.13
T1	Ref		Ref	
T2	1.34 (1.05–1.70)		1.23 (0.94–1.63)	
Surgery options		0.005*		0.003*
Local resection	Ref		Ref	
Radical resection	0.69 (0.53–0.90)		0.64 (0.48–0.85)	
Radiation		0.14		
No	Ref		Ref	0.17
Yes	1.22 (0.94–1.58)		1.57 (0.82–2.99)	
Chemotherapy		0.37		
No	Ref		Ref	
Yes	1.13 (0.87–1.48)		0.66 (0.34–1.27)	
CEA		<0.001*		<0.001*
Negative	Ref		Ref	
Positive	1.86 (1.45–2.39)		1.80 (1.40–2.32)	
Perineural invasion		0.69		
No	Ref		Ref	
Yes	1.17 (0.55–2.47)		1.11 (0.52–2.36)	
Tumor size (mm)		0.03*		0.01*
<18	Ref		Ref	
≥18	1.32 (1.02–1.71)		1.41 (1.08–1.84)	

 $^{^\}star,$ statistically significant. CEA, carcinoembryonic antigen; CI, confidence interval.

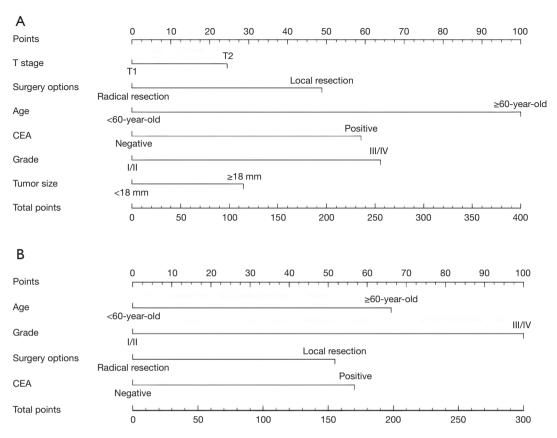


Figure 2 Nomograms to predict OS (A) and CSS (B) in ESRC patients. CEA, carcinoembryonic antigen; CSS, cancer-specific survival; ESRC, early-stage rectal cancer; OS, overall survival.

1-, 3-, and 5-year were 0.67, 0.64, and 0.62, respectively (Figure S1A-S1C). For CSS, ROC analysis also yielded promising results, with AUCs of 0.72, 0.71, and 0.63 for 1-, 3-, and 5-year survival in the training set, and 0.63, 0.69, and 0.62 in the validation set (Figure S1D-S1F). In the training set and validation set for both OS and CSS, calibration plots demonstrated strong concordance between predicted and observed outcomes (Figure S2). Furthermore, both in the training sets and validation sets, the DCA curves of OS and CSS illustrated enhanced clinical utility relative to the 7th AJCC staging system (Figure S3).

The effects of different treatment therapy in various subgroups

Patients were divided into two subgroups according to their prediction model scores: the high-risk group (≥128 points) and the low-risk group (<128 points) in the OS nomogram, as well as the low-risk group (<66 points) and the high-risk

group (≥66 points) in the CSS nomogram. Kaplan-Meier survival curves revealed significant differences in both OS and CSS between these groups (Figure 3). Following PSM, we assessed various treatment options across subgroups. Notably, radiation and chemotherapy provided no survival benefits for ESRC patients in any subgroup and were associated with worse OS and CSS in the low-risk group (Figure 4, Figure S4). Additionally, local resection did not yield significant survival differences compared to radical resection in the low-risk group, while poorer OS and CSS were observed in the high-risk group (Figure 5, Figure S5). In subgroups undergoing local resection, neither radiotherapy nor chemotherapy enhanced OS or CSS outcomes (Figure 6, Figure S6). Importantly, the subset of patients potentially benefiting from local resection appeared limited to younger individuals (Figure S7). Furthermore, the age subgroup analysis was influenced by biases inherent in the retrospective study, as all elderly patients were classified into the high-risk group.

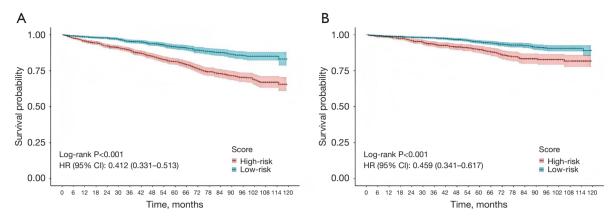


Figure 3 The Kaplan-Meier survival curves of OS (A) and CSS (B) between the high-risk group and the low-risk group. CI, confidence interval; CSS, cancer-specific survival; HR, hazard ratio; OS, overall survival.

Discussion

Radical resection remains the cornerstone treatment for ESRC and is widely endorsed in clinical practice (11,14). However, this conventional approach carries considerable risks, including stoma formation, anastomotic leaks, genitourinary dysfunction, fecal incontinence, and sexual dysfunction (16). Such complications significantly impair patients' quality of life post-resection. With the advent of advanced endoscopic techniques, local resection has emerged as a promising alternative for patients with ESRC (17). Furthermore, studies on prognostic outcomes in ESRC are limited, and comprehensive analyses of treatment options across various subgroups remain scarce (18,19).

In this study, we examined prognostic factors and devised predictive nomograms for OS and CSS in ESRC patients. Our analysis of 2,501 ESRC patients from the SEER database corroborated previous findings, identifying T stage, poor tumor grade, age, tumor size, elevated CEA levels, and surgical options as independent prognostic factors for OS and CSS. In rectal cancer, advanced T stage and lower tumor grade were well-established independent risk factors that significantly impact tumor survival, as evidenced by numerous studies (20,21). Likewise, elevated CEA level has been identified as critical to survival prognosis in patients with rectal cancer (22). Additionally, a recent study suggested that tumor size was associated with early metastasis and poor prognosis (23). What's more, we identified local resection as an independent risk factor for OS and CSS in ESRC patients, consistent with findings from previous studies (24,25). However, a study has found that local resection was safe and beneficial for selected

patients (26). One possible reason is that neoadjuvant chemoradiotherapy significantly improves survival outcomes in LARC but not in ESRC (27).

The predictive nomogram effectively stratified patients into low-risk and high-risk groups, with Kaplan-Meier analysis revealing significant disparities in OS and CSS between these cohorts. Following PSM in the training group, neither chemotherapy nor radiotherapy conferred survival benefits for ESRC patients, particularly in the low-risk group, where both treatments correlated with poorer OS and CSS. While radiotherapy is integral to the management of CRC, its efficacy is often limited by low tumor radiosensitivity and toxicity to adjacent normal tissues (28). In line with previous studies, our results showed that radiotherapy provided no benefit for ESRC patients across all subgroups. However, a study suggested that ESRC patients who underwent local resection may benefit from (neo) adjuvant therapy due to the risk of lymph node metastasis (29). Contrarily, our results indicated that radiotherapy did not enhance OS or CSS in those receiving local resection, possibly due to the low incidence of perineural invasion and the favorable tumor grades in the cohort. Chemoradiotherapy is the standard of care for LARC in conjunction with radical resection (11). Recently, the benefits of chemotherapy have been observed in ESRC (28,30). Yet, our analysis revealed no significant improvement in OS or CSS for any subgroup receiving chemotherapy. This aligns with recent studies indicating that adjuvant chemotherapy does not confer substantial advantages in early-stage rectal adenocarcinoma (31). A possible reason is that ESRC has low rates of recurrence and

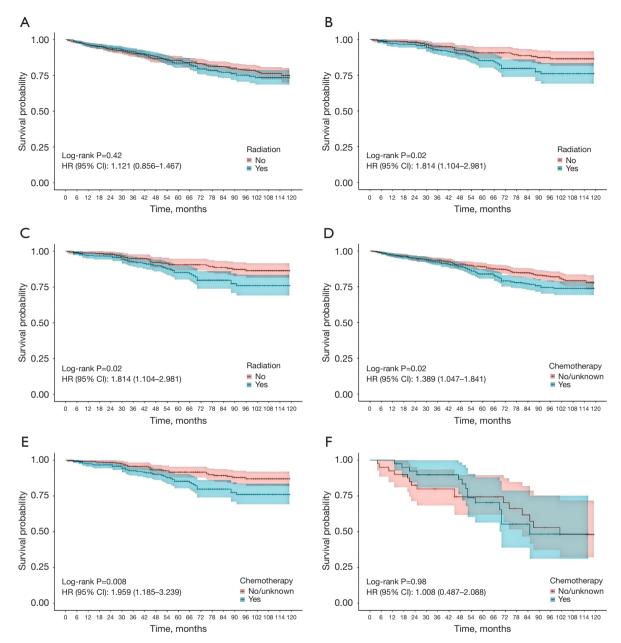


Figure 4 The Kaplan-Meier survival curves of OS under radiation therapy in all (A), low-risk group (B), high-risk group (C) patients and under chemotherapy in all (D), low-risk group (E), high-risk group (F) patients. CI, confidence interval; HR, hazard ratio; OS, overall survival.

distant metastasis (19). Previous research has indicated that radical resection yields better survival outcomes than local resection (29). Our study found no significant differences in OS or CSS within the low-risk group, particularly among younger patients. This may be attributed to the reduced rates of recurrence and metastasis in our selected low-risk cohort. Moreover, the limited sample size of patients

undergoing local resection and neoadjuvant therapy underscores the critical need for prospective clinical trials with large sample to validate these findings.

Otherwise, there are a growing number of studies showing the positive effect of neoadjuvant radiotherapy and chemotherapy in rectal cancer, especially in locally advanced disease. Spatola *et al.* conducted a study of 784 patients

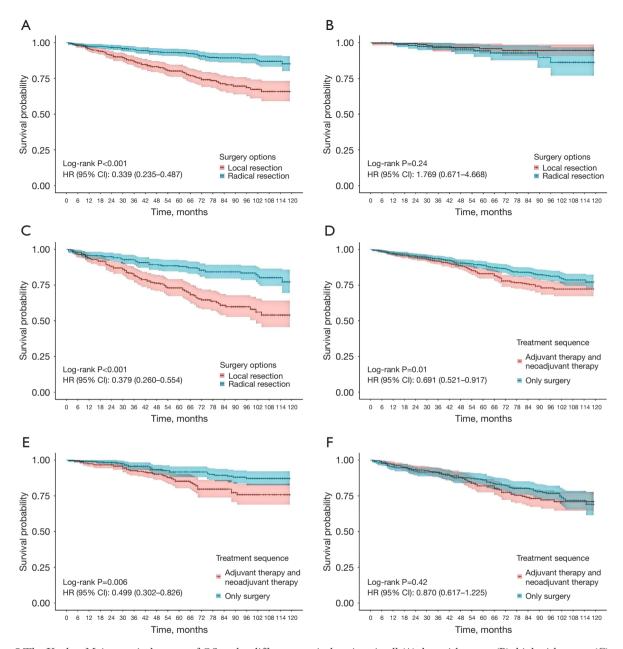


Figure 5 The Kaplan-Meier survival curves of OS under different surgical options in all (A), low-risk group (B), high-risk group (C) patients and under different treatment options in all (D), low-risk group (E), high-risk group (F) patients. CI, confidence interval; HR, hazard ratio; OS, overall survival.

with rectal cancer at 13 research centers, and found that neoadjuvant therapy was feasible for most patients in the comprehensive treatment of rectal cancer, which can increase the probability of undergoing sphincter preservation surgery and reduce the toxicity during treatment significantly (32). In addition, Lo *et al.* found that neoadjuvant therapy based on integrated intensified

treatment was proved to be feasible and well-tolerated in patients with LARC, showing high pCR rates (22.2%) and disease-free survival at a median follow-up of 30 months (33). In contrast, studies related to ESRC have been few and mixed. It is undeniable that our results have been elaborated from a perspective of the predictive model. Therefore, more large-scale prospective studies are needed

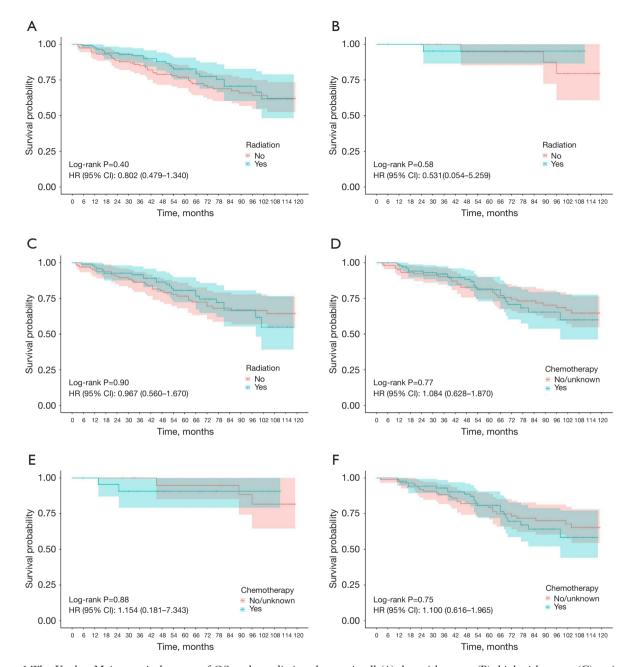


Figure 6 The Kaplan-Meier survival curves of OS under radiation therapy in all (A), low-risk group (B), high-risk group (C) patients and under chemotherapy in all (D), low-risk group (E), high-risk group (F) patients after local resection. CI, confidence interval; HR, hazard ratio; OS, overall survival.

to confirm our conclusions in the environment using multidisciplinary assessment.

A comprehensive understanding of OS and CSS is crucial for alleviating anxiety and enhancing the quality of life in patients, particularly those with initially poor prognoses. The development of OS and CSS nomograms enables clinicians to evaluate mortality risk and design suitable follow-up and monitoring strategies. This methodology provides critical insights into the evolving nature of postoperative survival, empowering both patients and clinicians to make informed decisions regarding therapeutic options.

However, this study had some limitations. First, the SEER database is deficient in key biomarker data, such as microsatellite instability and deficient mismatch repair status, both of which are essential prognostic indicators. Additionally, it offered only basic therapeutic records, lacking detailed information on surgical techniques, chemotherapy regimens, radiation doses, patient health status, and socio-economic factors that may influence survival outcomes. These gaps constrained the depth of our analysis, and future research should aim to incorporate these variables to better elucidate their impact. Furthermore, the retrospective design introduced potential selection bias; thus, prospective cohort studies or randomized controlled trials are necessary to validate our findings and mitigate bias.

Conclusions

Our study provided a comprehensive analysis of prognostic factors influencing OS and CSS in patients with ESRC, utilizing data from the SEER database. We have developed and validated prediction nomograms for both OS and CSS, evaluating the impact of various treatment options across distinct subgroups. While our model exhibited promising performance in forecasting survival outcomes for ESRC patients, further validation through multi-center studies is essential to confirm its applicability in clinical situation.

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Footnote

Reporting Checklist: The authors have completed the TRIPOD reporting checklist. Available at https://tcr.amegroups.com/article/view/10.21037/tcr-24-1888/rc

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have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013).

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