

Supplementary Information

Table S1. General characteristics of haematological parameters and inflammatory indices in patients with endometrial cancer (n = 528)

Characteristic	Whole population	Training cohort	Validation cohort	P value
HGB (g/L)				0.637
Mean ±(SD)	119.90 ± (16.54)	120.15 ± (16.42)	119.47 ± (16.76)	
Median (IQR)	123.00 (111.00-131.00)	123.00 (111.00-131.00)	123.00 (112.00-131.00)	
Albumin (g/L)				0.527
Mean ±(SD)	41.62 ± (5.10)	41.52 ± (4.93)	41.80 ± (5.37)	
Median (IQR)	42.00(39.00-45.00)	42.00 (39.00-45.00)	43.00 (39.00-45.00)	
Lymphocyte ($\times 10^9/\text{L}$)				0.808
Mean ±(SD)	1.68 ± (0.57)	1.69 ± (0.58)	1.65 ± (0.55)	
Median (IQR)	1.66 (1.25-2.04)	1.68 (1.25-2.09)	1.61 (1.25-1.99)	
Blood platelet ($\times 10^9/\text{L}$)				0.266
Mean ±(SD)	221.59 ± (75.12)	218.91 ± (75.32)	226.08 ± (74.73)	
Median (IQR)	211.50 (176.00-259.00)	206.00 (175.5-249.00)	214 (177.00-262.50)	
HALP score				0.344
Mean ±(SD)	42.67 ± (23.51)	43.38 ± (23.66)	41.47 ± (23.27)	
Median (IQR)	41.58 (24.30-53.33)	42.39 (25.86-53.51)	40.39 (23.70-52.58)	

Abbreviations: HGB, Hemoglobin; HALP, hemoglobin, albumin, lymphocyte, and platelet. IQR, interquartile range.

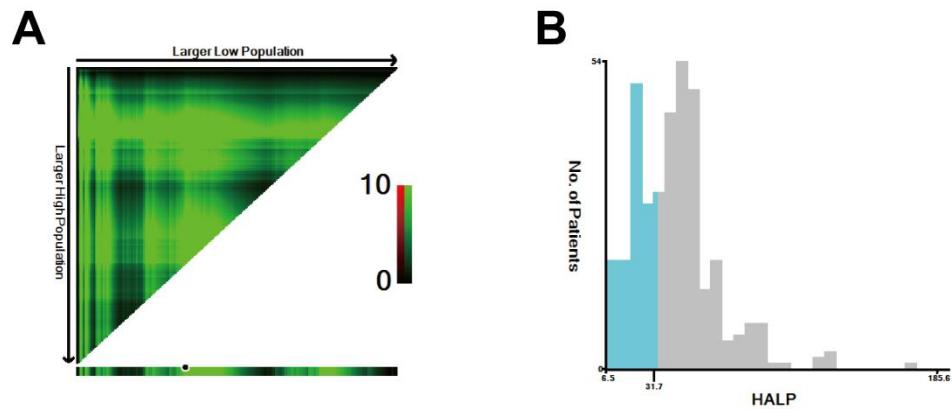


Figure S1. The optimal cutoff value of HALP score determined by X-tile (value = 31.70).

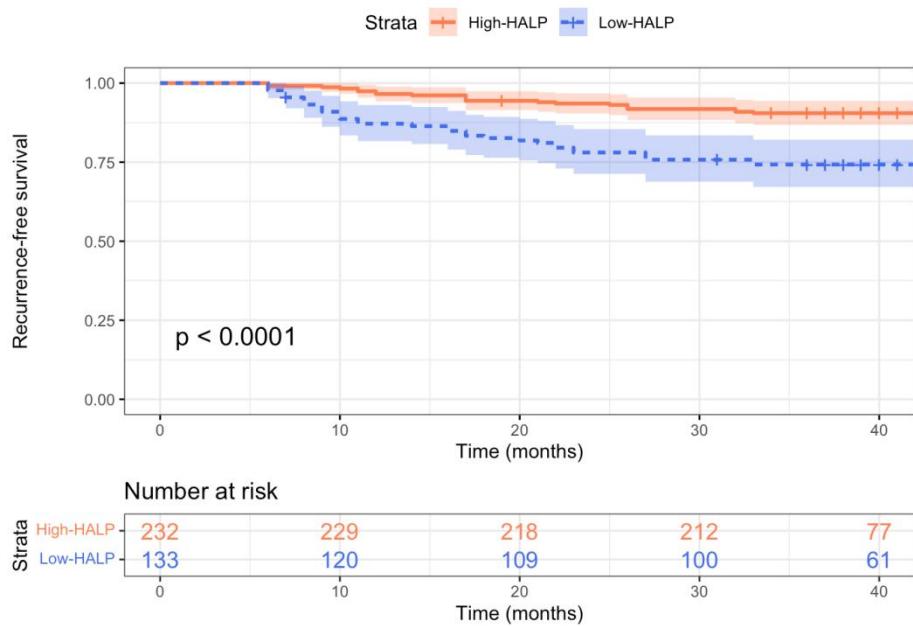


Figure S2. RFS of stage I-III endometrial cancer stratified by HALP score.

Table S2. Variable risk score and RFS probability prediction in patients with stage I-III endometrial cancer

Characteristics	Risk score	Total score	Probability of RFS
Age		Total Points for 1-year RFS	Probability of 1-year RFS
< 60 years	0	357	0.2
≥ 60 years	33	342	0.3
FIGO staging		328	0.4
I	0	314	0.5
II	15	299	0.6
III	42	281	0.7
Grade		258	0.8
G1-G2	0	220	0.9
G3	49	Total Points for 2-year RFS	Probability of 2-year RFS
LVSI		321	0.2
Negative	0	306	0.3
Positive	49	292	0.4
Ca125		278	0.5
≤ 35 U/ml	0	263	0.6
> 35 U/ml	36	245	0.7
ProMisE molecular subgroup		222	0.8
p53-wt	51	184	0.9
<i>POLE</i> EDM	0	Total Points for 3-year RFS	Probability of 3-year RFS
MMR-D	75	303	0.2
p53-abn	100	289	0.3
Adjuvant treatment		275	0.4
Yes	0	261	0.5
No	50	246	0.6
HALP score		228	0.7
High	0	204	0.8
Low	39	167	0.9

Abbreviations: FIGO, International Federation of Gynecology and Obstetrics; LVSI, lymphovascular space invasion; G1-G2, grade 1/2; G3, grade 3; *POLE* EDM, polymerase epsilon (*POLE*) exonuclease domain mutation; p53-wt, p53 wide-type; p53-abn, p53-abnormal; MMR-D, deficient mismatch repair; HALP, hemoglobin-albumin-lymphocyte-platelet.

Table S3. C-index for the nomogram and model variables in the training and external validation cohorts

Risk stratification system	Training cohort		External validation cohort	
	C-index	95% CI	C-index	95% CI
ProMisE molecular subgroup	0.678	(0.611-0.745)	0.691	(0.607-0.775)
FIGO stage^a	0.736	(0.673-0.799)	0.688	(0.604-0.772)
Nomogram Model	0.875	(0.836-0.914)	0.859	(0.806-0.912)

^a Adopt FIGO 2023 staging criteria for endometrial cancer.

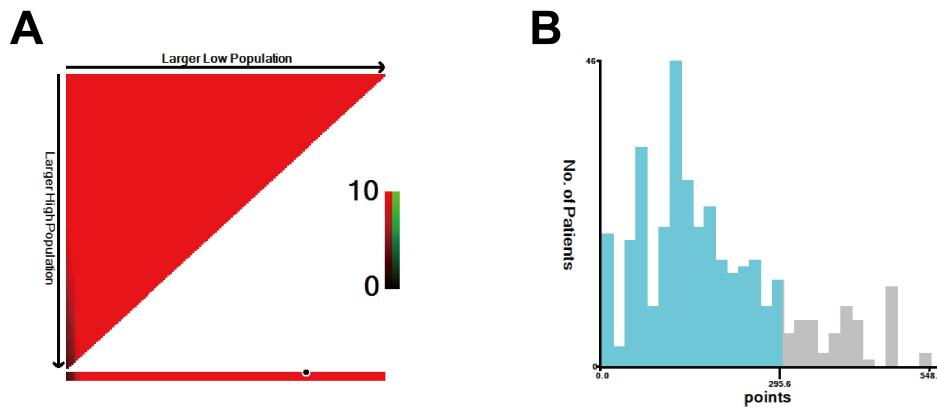


Figure S3. The optimal cutoff value for nomogram risk score determined by X-tile; and the RFS of endometrial cancer patients stratified by the scores (optimal cutoff value = 295.6 pts).

Supplementary materials

Description: Related Computerized Programs for Nomogram Development Using R

APPENDIX

R code for developing the RFS nomogram model and calculating the nomogram score for each patient

```
library(survival)
library(rms)
dd <- datadist(data)
options(datadist = "dd")
F <- cph(Surv(time, status) ~ Age + FIGO + Grade + LVSI + Ca125 +
ProMisE_molecular + HALP + Adjuvant_treatment, data = data, x = TRUE, y =
TRUE, surv = TRUE)
# Developing nomogram models
survival <- Survival(f)
survival1 <- function(x)survival(12,x)
survival2 <- function(x)survival(36,x)
survival3 <- function(x)survival(60,x)
nom <- nomogram(f, fun=list(survival1,survival2,survival3), lp = FALSE, fun.at =
c(0.1, seq(0.1,0.9,by=0.1), 0.9), funlabel = c("1-year RFS", "3-year RFS","5-year
RFS"))
plot(nom)
# Calculating the nomogram score for each patient
library(nomogramFormula)
data = na.omit(data)
dd<-datadist(data)
options(datadist="dd")
f <- cph(Surv(time, status) ~ Age + FIGO + Grade + LVSI + Ca125 +
ProMisE_molecular + HALP + Adjuvant_treatment, data=data, x=TRUE, y=TRUE,
surv = TRUE)
```

```

results <- formula_rd(nom)
results$formular
data$points <- points_cal(formula = results$formular, rd=data)

For calculating the C-index and 95% CI in both the training and validation cohorts

library(survival)
library(rms)
fit<-coxph(Surv(time, status) ~ Age + FIGO + Grade + LVSI + Ca125 +
ProMisE_molecular +HALP + Adjuvant_treatment, data = Ytraindata)
survConcordance(Surv(Ytraindata$time,Ytraindata$status)~predict(fit,Ytraindata))
survConcordance(Surv(Ytestdata$time,Ytestdata$status)~predict(fit,Ytestdata))

# Other coding methods
library(rms)
library(foreign)
library(survival)
ddist <- datadist(data)
options(datadist = 'ddist')
# C-index confidence interval: C-index-1.96*se/C-index+1.96*se
fmla1 <- as.formula(Surv(time, status) ~ Age + FIGO + Grade + LVSI + Ca125 +
ProMisE_molecular +HALP + Adjuvant_treatment)
cox2 <- coxph(fmla1, data=data)
summary(cox2)

```

R code for ROC curve for the nomogram model

```

cox_m <- coxph(Surv(time, status) ~ Age + FIGO + Grade + LVSI + Ca125 +
ProMisE_molecular +HALP + Adjuvant_treatment, data = data)
cox_m1<-step(cox_m, direction = "both")
risk_score<-predict(cox_m1, type="risk", newdata=data)
risk_level<-as.vector(ifelse(risk_score > median(risk_score), "High", "Low"))
#colnames(data) <- c("id", "survival_time", "status", "risk_score", "predict_time")

```

```

write.table(cbind(id=rownames(cbind(data$time, data$status, risk_score, risk_level)),
cbind(data$time,data$status,risk_score,risk_level)),"risk_score.txt",sep="\t",quote=F,r
ow.names=F)

# Plotting the ROC curves for 1-year RFS

library(survivalROC)

colnames(data) <- c("id", "survival_time", "status", "risk_score", "predict_time")

predict_time <- 12*1

myroc <- survivalROC(Stime = data$survival_time, status=data$status, marker =
data$risk_score, predict.time = predict_time, method="KM")

pdf("ROC_1-year_RFS_in_Training_set.pdf")
plot(mycoc$FP, myroc$TP, type="l", xlim=c(0,1), ylim=c(0,1), col="blue",
xlab="FP", ylab="TP", lwd=2.5, main=paste("1-year
RFS","AUC=",round(mycoc$AUC,3)))
abline(0,1)

dev.off()

# Plotting the ROC curves for 3-year RFS

library(survivalROC)

colnames(data) <- c("id", "survival_time", "status", "risk_score", "predict_time")

predict_time <- 12*3

myroc <- survivalROC(Stime = data$survival_time, status=data$status, marker =
data$risk_score, predict.time = predict_time, method="KM")

pdf("ROC_3-year_RFS_in_Training_set.pdf")
plot(mycoc$FP, myroc$TP, type="l", xlim=c(0,1), ylim=c(0,1), col="blue",
xlab="FP", ylab="TP", lwd=2.5, main=paste("3-year
RFS","AUC=",round(mycoc$AUC,3)))
abline(0,1)

dev.off()

# Plotting the ROC curves for 5-year RFS

predict_time<-12*5

```

```

myroc<-survivalROC(Stime = data$survival_time, status = data$status, marker =
data$risk_score, predict.time=predict_time,method="KM")
pdf("ROC_5-year RFS in Training set.pdf")
plot(mycoc$FP, myroc$TP, type="l", xlim=c(0,1), ylim=c(0,1), col="red",
xlab="FP", ylab="TP", lwd=2.5, main=paste("5-year RFS","AUC=", round(mycoc$AUC,3)))
abline(0,1)
dev.off()

```

R code for calibration curve of the nomogram model

```

library(survival)
library(rms)
Ytraindata <- data
f1 <- cph(Surv(Ytraindata$time, Ytraindata$status==1)~Age + FIGO + Grade + LVSI
+ Ca125 + ProMisE_molecular + HALP + Adjuvant_treatment, data = Ytraindata,
x=TRUE, y=TRUE, surv=TRUE, time.inc=3*12)
cal<-calibrate(f1, cmethod = 'KM', method = "boot", u=3*12, m=120, B=1000)
plot(cal, #Input the calibrate model
      lwd=1, #Line segment width
      lty=1, #Line segment type
      errbar.col=c(rgb(0,0,0,maxColorValue = 255)), #Specify the error line color
      xlim = c(0.7,1),ylim = c(0.7,1), #Set the axis range
      xlab ="Nomogram-Predicted Probability of 3-year RFS (proportion)", #Set the
      x-axis label
      ylab="Actual 3-year RFS (proportion)", #Set the y-axis label
      col=c(rgb(255,0,0,maxColorValue =255))) abline(0,1,lty = 3, lwd = 2, col =
      c(rgb(0,118,192,maxColorValue=255)))
lines(cal[,c('mean.predicted','KM')], type = 'b',lwd = 2, col =
      c(rgb(192,98,83,maxColorValue = 255)), pch = 16)
abline(0,1,lty = 3,lwd = 2,col = c(rgb(0,118,192,maxColorValue=255)))

```

```

lines(cal[,c('mean.predicted','KM')], type = 'b',lwd = 2, col =
c(rgb(192,98,83,maxColorValue = 255)),pch = 16)

#1_year

f1 <- cph(Surv(Ytraindata$time, Ytraindata$status==1)~Age + FIGO + Grade + LVSI
+ Ca125 + ProMisE_molecular +HALP + Adjuvant_treatment, data = Ytraindata,
x=TRUE, y=TRUE, surv=TRUE, time.inc=1*12)

cal<-calibrate(f1, cmethod = 'KM', method = "boot", u=1*12, m=120, B=1000)

plot(cal, lwd=1, lty=1, errbar.col=c(rgb(0,0,0,maxColorValue = 255)), xlim = c(0.8,1),
ylim = c(0.7,1), xlab ="Nomogram-Predicted Probability of 1-year RFS (proportion)",
ylab="Actual 1-year RFS (proportion)", col=c(rgb(255,0,0, maxColorValue =255)))

abline(0,1,lty = 3, lwd = 2, col = c(rgb(0,118,192,maxColorValue=255)))

lines(cal[,c('mean.predicted','KM')], type = 'b',lwd = 2, col =
c(rgb(192,98,83,maxColorValue = 255)),pch = 16)

#5_year

f1 <- cph(Surv(Ytraindata$time, Ytraindata$status==1)~Age + FIGO + Grade + LVSI
+ Ca125 + ProMisE_molecular +HALP + Adjuvant_treatment, data = Ytraindata,
x=TRUE, y=TRUE, surv=TRUE, time.inc=5*12)

cal<-calibrate(f1, cmethod = 'KM', method = "boot", u=5*12, m=120, B=1000)

plot(cal, lwd=1, lty=1, errbar.col=c(rgb(0,0,0,maxColorValue = 255)), xlim = c(0.8,1),
ylim = c(0.7,1), xlab ="Nomogram-Predicted Probability of 1-year RFS (proportion)",
ylab="Actual 1-year RFS (proportion)", col=c(rgb(255,0,0, maxColorValue =255)))

abline(0,1,lty = 3, lwd = 2, col = c(rgb(0,118,192,maxColorValue=255)))

lines(cal[,c('mean.predicted','KM')], type = 'b',lwd = 2, col =
c(rgb(192,98,83,maxColorValue = 255)),pch = 16)

```

For calculating the C-index and 95% CI of model variables for RFS

```

# ProMisE molecular subgroup

library(survival)

library(rms)

fit1 <- coxph(Surv(time,status)~ProMisE_molecular, data = Ytraindata)

```

```

survConcordance(Surv(Ytraindata$time, Ytraindata$status)~predict(fit1, Ytraindata))
survConcordance(Surv(Ytestdata$time, Ytestdata$status)~predict(fit1, Ytestdata))

# FIGO stage

library(survival)
library(rms)

fit1<- coxph(Surv(time,status)~FIGO_stage, data = Ytraindata)
survConcordance(Surv(Ytraindata$time, Ytraindata$status)~predict(fit1, Ytraindata))
survConcordance(Surv(Ytestdata$time, Ytestdata$status)~predict(fit1, Ytestdata))

```

**For calculating the C-index and 95% CI of different risk stratification systems
for RFS**

```

# Model A

library(survival)
library(rms)

fit1 <- coxph(Surv(time,status)~age + surgical_staging + Grade + LVSI + FIGO, data
= Ytraindata)
survConcordance(Surv(Ytraindata$time, Ytraindata$status)~predict(fit1, Ytraindata))
survConcordance(Surv(Ytestdata$time, Ytestdata$status)~predict(fit1, Ytestdata))

```

```

# Model B

library(survival)
library(rms)

fit1 <- coxph(Surv(time,status)~SII, data = Ytraindata)
survConcordance(Surv(Ytraindata$time, Ytraindata$status)~predict(fit1, Ytraindata))
survConcordance(Surv(Ytestdata$time, Ytestdata$status)~predict(fit1, Ytestdata))

```

```

# Model C

library(survival)
library(rms)

```

```

fit1 <- coxph(Surv(time,status)~Age + Ca125 + FIGO + LVSI + P53, data =
Ytraindata)

survConcordance(Surv(Ytraindata$time,Ytraindata$status)~predict(fit1, Ytraindata))

survConcordance(Surv(Ytestdata$time,Ytestdata$status)~predict(fit1, Ytestdata))

# Our model

library(survival)

library(rms)

fit<-coxph(Surv(time, status) ~ Age + FIGO + Grade + LVSI + Ca125 +
ProMisE_molecular +HALP + Adjuvant_treatment, data = Ytraindata)

survConcordance(Surv(Ytraindata$time,Ytraindata$status)~predict(fit,Ytraindata))

survConcordance(Surv(Ytestdata$time,Ytestdata$status)~predict(fit,Ytestdata))

```