

## I. Survival Analysis

### A. Core Functions & Properties

$S(t) = P(T > t) = \exp[-H(t)]; h(t) = \frac{f(t)}{S(t)} = -\frac{d \log S(t)}{dt};$

$H(t) = \int_0^t h(u)du = -\log S(t)$

**Properties:**  $S(0) = 1, S(\infty) = 0; h(t) \geq 0$  always;  $H(0) = 0, H(t)$  monotone non-decreasing

**Censoring:** Right-censoring (most common); observe  $(Y, \delta)$  where  $Y = \min(T, C), \delta = I(T \leq C)$

**Assumption:** Non-informative censoring (censoring independent of failure time)

### B. Kaplan-Meier Estimator

$\hat{S}(t) = \prod_{t_i \leq t} (1 - \frac{d_i}{n_i}) = \prod_{t_i \leq t} \frac{n_i - d_i}{n_i}$  where  $d_i$ =events,  $n_i$ =at risk just before  $t_i$

**KM Table (7 columns):**  $j, t_j, \text{interval}, n_j, d_j, p_j = 1 - d_j/n_j, \hat{S}(t_j)$

**Key:** Censored at  $t_j$  removed from risk set at  $t_j^+$ ; censoring  $\downarrow n_{j+1}$  but NOT  $\uparrow d_j$ ; if max time censored,  $\hat{S}(t)$  doesn't reach 0

**Greenwood Variance:**  $\widehat{\text{Var}}(\hat{S}) = \hat{S}^2 \sum_{t_i \leq t} \frac{d_i}{n_i(n_i - d_i)}$  (undefined when  $n_i = d_i$ )

**95% CI:** Log-log:  $\exp(-e^{C_u}), \exp(-e^{C_l})$  where  $\sigma^2 = \frac{1}{[\log \hat{S}]^2} \sum \frac{d_i}{n_i(n_i - d_i)}; C_u = \log[-\log \hat{S}] + 1.96\sigma$

**Nelson-Aalen:**  $\hat{H}(t) = \sum_{t_i \leq t} \frac{d_i}{n_i}; \tilde{S}(t) = \exp[-\hat{H}(t)]$  (asymptotically  $\equiv$  KM)

**Quantiles:**  $\hat{t}_p = \min\{t_j : \hat{S}(t_j) < 1 - p\}$ ; if  $\hat{S}(t_j) = 1 - p$  exactly,  $\hat{t}_p = (t_j + t_{j+1})/2$

Median:  $\hat{t}_{0.5} = \min\{t_j : \hat{S}(t_j) < 0.5\}$ ; Q1:  $\hat{t}_{0.25} = \min\{t_j : \hat{S}(t_j) < 0.75\}$

**Mean:**  $\hat{\mu} = \int_0^\infty \hat{S}(t)dt$  (underestimated if max censored)

**Discrete:**  $\hat{\mu} = \sum_{j=1}^{J-1} \hat{S}(t_{j-1})(t_j - t_{j-1})$  where  $t_0 = 0$

**Variance (HW2):**  $\text{Var}(\hat{\mu}) = \frac{n_d}{n_d - 1} \sum_{j=1}^{J-1} \frac{A_j^2 d_j}{n_j(n_j - d_j)}$  where  $A_j = \sum_{k=j+1}^J \hat{S}(t_{k-1})(t_k - t_{k-1}), n_d = \sum d_j$

**RMST:**  $\hat{\mu}(\tau) = \int_0^\tau \hat{S}(t)dt$  (more robust; choose  $\tau$ =max follow-up)

### C. Survival Curve Comparison

Both Log-rank and Wilcoxon can be extended to K groups

$H_0 : S_1(t) = S_0(t)$  for all  $t \leq \tau$

**Weighted Test:**  $Q = \frac{[\sum w_j(d_{1j} - e_{1j})]^2}{\sum w_j^2 \nu_{1j}} \sim \chi_1^2$

$e_{1j} = \frac{n_{1j} d_j}{n_j}; \nu_{1j} = \frac{n_{1j} n_{0j} d_j (n_j - d_j)}{n_j^2 (n_j - 1)}$

- **Log-rank:**  $w_j = 1$  (equal weights, optimal under PH)
- **Wilcoxon:**  $w_j = n_j$  (early-time sensitive, since  $n_j$  larger early)
- **Curves cross  $\Rightarrow$  both tests lose power;** use RMST/landmark
- K groups:  $\text{df}=K-1$

### D. Cox Proportional Hazards

$h(t, \mathbf{x}) = h_0(t) \exp(\boldsymbol{\beta}^T \mathbf{x})$  (no  $\beta_0$ !)

$\log \frac{h(t, \mathbf{x})}{h_0(t)} = \boldsymbol{\beta}^T \mathbf{x}; S(t, \mathbf{x}) = [S_0(t)]^{\exp(\boldsymbol{\beta}^T \mathbf{x})}$

**HR Interpretation:**

- Binary ( $x=0$  vs 1):  $\text{HR} = e^\beta$
- Continuous (k-unit  $\uparrow$ ):  $\text{HR} = e^{k\beta}$
- Multi-class:  $\text{HR}_j = e^{\beta_j}$  (vs ref);  $\text{HR}(j : k) = e^{\beta_j - \beta_k}$

- **HR = instantaneous relative risk  $\neq$  cumulative risk ratio**

**95% CI:**  $\exp(\hat{\beta} \pm 1.96 \cdot SE)$

**Partial Likelihood:**  $L_p(\boldsymbol{\beta}) = \prod_{j=1}^m \frac{\exp(\mathbf{x}_j^T \boldsymbol{\beta})}{\sum_{k \in R(t_j)} \exp(\mathbf{x}_k^T \boldsymbol{\beta})}$

$m$ =distinct failure times;  $R(t_j)$ =risk set;  $\log L_p = \sum [\mathbf{x}_j^T \boldsymbol{\beta} - \log \sum_{k \in R} \exp(\mathbf{x}_k^T \boldsymbol{\beta})]$

**Ties:** EFRON (recommended) approximates exact; BRESLOW faster but less accurate

**PH Assumption Checks:**

1. Log-log plot:  $\log[-\log \hat{S}(t)]$  vs  $\log t$  (parallel  $\Rightarrow$  PH)
2. Schoenfeld residuals: **assess ph / resample;**
3. Time interaction: add  $X \times \log(t)$ ; if sig  $\Rightarrow$  violates PH

**Non-PH Solutions:**

- Stratify: **strata Z;** (cannot estimate HR for Z)
- Time-varying coef:  $X \times t$  or  $X \times \log(t)$
- TDC: model **(tstart,tstop)\*status(0)=...**

**Residuals:**

- **Martingale**  $\in (-\infty, 1]$ :  $M_i = \delta_i - \hat{H}_i(t_i)$ ; check functional form via plot vs  $X$ +lowess; **horizontal lowess  $\Rightarrow$  correct form**, non-horizontal  $\Rightarrow$  need transformation
- **Deviance:**  $D_i = \text{sign}(M_i) \sqrt{-2[M_i + \delta_i \log(\delta_i - M_i)]}$ ;  $|D_i| > 2$  noteworthy,  $> 3$  outlier
- **Dfbeta:**  $dfbeta_i = \hat{\beta} - \hat{\beta}_{(-i)}$ ;  $|dfbeta| > 2/\sqrt{n}$  influential

**Confounding:**  $\Delta \hat{\beta}\% = 100 \times \frac{|\hat{\theta} - \hat{\beta}|}{|\hat{\beta}|}$ ;  $> 20\% \Rightarrow$  confounder

**Interaction:**  $h(t) = h_0(t) \exp(\beta_1 X_1 + \beta_2 X_2 + \beta_3 X_1 X_2)$ ;  $\text{HR}_{X_1}(X_2 = a) = e^{\beta_1 + \beta_3 a}$

**Variance (group comparison):**  $\text{Var}(a \times \hat{\beta}_j - b \times \hat{\beta}_k) = a^2 \times \text{Var}(\hat{\beta}_j) + b^2 \times \text{Var}(\hat{\beta}_k) - 2ab \text{Cov}(\hat{\beta}_j, \hat{\beta}_k)$

**Hypothesis Tests:**

- **Wald:**  $(\hat{\beta}/SE)^2 \sim \chi_1^2$  (single param); Type 3 (multiple)
- **LRT:**  $G = -2[\log L_{\text{reduced}} - \log L_{\text{full}}] \sim \chi_{df}^2$  (preferred for nested models)
- **Score:** Based on  $U(\beta)$ ; used in PH test

### E. Key Points

**Semi-parametric:** Cox has nonparametric  $h_0(t)$  + parametric  $\boldsymbol{\beta}$   
**Model comparison:** LRT (Likelihood Ratio Test, preferred when nested), Wald, Score test;  $\text{df}=\#$  parameters difference

**Stratified analysis:** Control confounders; assumes common treatment effect across strata

**Baseline hazard:** Cannot estimate  $h_0(t)$  directly, but can get  $S_0(t)$  via baseline

## II. Categorical Data

### A. GLM Framework

1. Distribution:  $Y \sim \text{Normal/Binomial/Poisson/NB}$
2. Linear predictor:  $\eta = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p$
3. Link:  $g\{E(Y|X)\} = \eta$

Model	Dist	Link	$g(\mu)$
Linear	Normal	Identity	$\mu$
Logistic	Binomial	Logit	$\log \frac{\mu}{1-\mu}$
Poisson	Poisson	Log	$\log \mu$
NB	NB	Log	$\log \mu$

Variance: Normal= $\sigma^2$ ; Binomial= $\mu(1-\mu)$ ; Poisson= $\mu$ ; NB= $\mu + \alpha\mu^2$

**Canonical link:** Logit for Binomial, Log for Poisson

**Fitted values:** Logistic:  $\hat{p} = \frac{e^{\hat{\eta}}}{1+e^{\hat{\eta}}}$  where  $\hat{\eta} = \hat{\beta}_0 + \hat{\beta}_1 X_1 + \dots$

**Interpretation:**

- Linear: " $\beta_j$ " = unit change in mean
- Logistic: " $e^{\beta_j}$ " = odds ratio (OR)
- Poisson/NB: " $e^{\beta_j}$ " = rate ratio (RR)

### B. Multinomial Logit (unordered)

$\log \frac{P(Y=j)}{P(Y=K)} = \alpha_j + \beta_j^T \mathbf{X}, j = 1, \dots, K-1$

**Expanded:**  $\log \frac{P(Y=j)}{P(Y=3)} = \alpha_j + \beta_{j1} X_1 + \beta_{j2} X_2 + \dots + \beta_{jp} X_p$  (each class  $j$  has own  $\beta_j$ )

**Example (Y=3 levels):**  $\log \frac{P(Y=1)}{P(Y=3)} = \alpha_1 + \beta_{11} \text{Age} + \beta_{12} \text{Sex}$ ;

$\log \frac{P(Y=2)}{P(Y=3)} = \alpha_2 + \beta_{21} \text{Age} + \beta_{22} \text{Sex}$

**df:** Continuous X:  $K-1$ ; q-level categorical X:  $(K-1)(q-1)$

**Interpretation:**

- $\beta_{jk}$ : change in log-odds of class  $j$  vs ref  $K$  per unit increase in  $X_k$
- $e^{\beta_{jk}}$ =RRR (relative risk ratio): multiplicative change in odds( $Y=j$ )/odds( $Y=K$ )
- Example:  $\beta_{11} = 0.5 \Rightarrow$  1-yr age increase multiplies odds( $Y=1$  vs  $Y=3$ ) by  $e^{0.5} = 1.65$
- Compare classes:  $e^{\beta_{1k} - \beta_{2k}}$ =RRR for class 1 vs class 2

**Probability:**  $P(Y = j) = \frac{\exp(\alpha_j + \beta_j^T \mathbf{X})}{1 + \sum_{k=1}^{K-1} \exp(\alpha_k + \beta_k^T \mathbf{X})}$ ;  $P(Y = K) = \frac{1}{1 + \sum_{k=1}^{K-1} \exp(\alpha_k + \beta_k^T \mathbf{X})}$

### C. Ordinal (Cumulative)Logit (ordered)

$\log[P(Y \leq j)] = \log \left[ \frac{P(Y \leq j)}{P(Y > j)} \right] = \alpha_j + \boldsymbol{\beta}^T \mathbf{X}, j = 1, \dots, K-1$

**Expanded:**  $\log[P(Y \leq j)] = \alpha_j + \beta_1 X_1 + \beta_2 X_2 + \dots + \beta_p X_p$  (same  $\boldsymbol{\beta}$  for all  $j$ )

**Example (Y=4 levels):**

- $\log \frac{P(Y \leq 1)}{P(Y > 1)} = \alpha_1 + \beta_1 \text{Age} + \beta_2 \text{Sex}$
- $\log \frac{P(Y \leq 2)}{P(Y > 2)} = \alpha_2 + \beta_1 \text{Age} + \beta_2 \text{Sex}$  (note: same  $\beta_1, \beta_2$ !)
- $\log \frac{P(Y \leq 3)}{P(Y > 3)} = \alpha_3 + \beta_1 \text{Age} + \beta_2 \text{Sex}$

**Proportional Odds:** same  $\boldsymbol{\beta}$  for all cutpoints; only  $\alpha_j$  differs

**df:** Continuous X: 1; q-level X:  $q-1$  (only ONE  $\boldsymbol{\beta}$  across all equations!)

**Proportional Odds Assumption Test (HW6):**

1. **Hypotheses:**  $H_0$ : slopes  $(\beta_1, \beta_2, \dots)$  equal across logit equations;  $H_a$ : different slopes needed
2. **Test:** Score chi-square test;  $\text{df}=(K-2)(q-1)$  where  $K$ =response levels,  $q$ =predictor levels
3. Example:  $Y=4$  levels,  $X$  4-group  $\Rightarrow \text{df}=(4-2) \times (4-1) = 6$
4. **P-value:**  $P(\chi_{df}^2 \geq S)$  where  $S$ =test statistic
5. **Decision:**  $p > 0.05 \Rightarrow$  fail to reject  $H_0 \Rightarrow$  use Ordinal;  $p < 0.05 \Rightarrow$  reject  $H_0 \Rightarrow$  use Multinomial

**Interpretation:**

- $\beta_k$ : change in log-cumulative-odds per unit increase in  $X_k$  (**same effect for all cutpoints**)
- $e^{\beta_k}$ =cumulative OR: multiplicative change in odds( $Y \leq j$ ) for all  $j$

- Default  $P(Y \leq j)$ :  $\beta > 0 \Rightarrow$  higher odds of  $Y \leq j \Rightarrow$  tend to lower levels
- Example:  $\beta_1 = 0.3 \Rightarrow$  1-yr age increase multiplies odds( $Y \leq$  any cutpoint) by  $e^{0.3} = 1.35$
- Key: **ONE**  $\beta$  per predictor applies to **ALL**  $(K - 1)$  equations; only intercepts  $\alpha_j$  vary

**Hypothesis Test for  $\beta_k$ :**

- $H_0 : \beta_k = 0$  vs  $H_a : \beta_k \neq 0$  (no effect of  $X_k$  on cumulative odds)
- Wald test:  $z = \frac{\hat{\beta}_k}{SE(\hat{\beta}_k)} \sim N(0, 1)$ ;  $p = 2P(|Z| > |z|)$
- $p < 0.05 \Rightarrow$  reject  $H_0 \Rightarrow X_k$  significantly affects ordered outcome

**Probability Calculation (HW6):**

**Ordinal:**  $P(Y \leq j|X) = \frac{e^{\alpha_j + \beta X}}{1 + e^{\alpha_j + \beta X}}$ ; then  $P(Y = j) = P(Y \leq j) - P(Y \leq j - 1)$ ;  $P(Y = K) = 1 - P(Y \leq K - 1)$

**Multinomial:**  $P(Y = j|X) = \frac{e^{\beta_{0j} + \beta_j X}}{1 + \sum_{k=1}^{K-1} e^{\beta_{0k} + \beta_k X}}$ ;  $P(Y = K) = \frac{1}{1 + \sum_{k=1}^{K-1} e^{\beta_{0k} + \beta_k X}}$

**D. Poisson Regression**

$\log\{E(Y|X)\} = \beta_0 + \beta^T X$

**Rate model:**  $\log\{E(Y|X)\} = \log(n) + \beta_0 + \beta^T X$

$\log(n)$ =offset (known, no estimate);  $n$ =exposure (person-years, months, etc.)

**Rate Ratio Derivation (HW7):** Binary  $X$ :  $RR = \frac{E(Y|X=1)}{E(Y|X=0)} = \frac{\exp(\beta_0 + \beta_1 \cdot 1 + \dots)}{\exp(\beta_0 + \beta_1 \cdot 0 + \dots)} = e^{\beta_1}$

**Interpretation:** Binary:  $RR = e^{\beta}$ ; Continuous k-unit:  $RR = e^{k\beta}$

**Goodness Of Fit:** Deviance= $-2(\log L_{current} - \log L_{saturated})$ ;

Pearson  $\chi^2 = \sum \frac{(y_i - \hat{\mu}_i)^2}{\hat{\mu}_i}$

**Overdispersion Check:** Scale=Deviance/df or Pearson  $\chi^2$ /df;  $\approx 1 \Rightarrow$  ok;  $> 1 \Rightarrow$  mild;  $> 1.5 \Rightarrow$  serious

**Pearson residual:**  $r_i = \frac{y_i - \hat{y}_i}{\sqrt{\hat{y}_i}}$ ;  $|r_i| > 2 \Rightarrow$  poor fit

**Overdispersion Solutions:**

- **Quasi-Poisson:** scale=pearson or scale=deviance;  $\hat{\beta}$  unchanged,  $SE_{new} = SE_{old} \times \sqrt{\phi}$  where  $\phi$ =Scale;  $CI_{new}$  wider
- **NB:** dist=negbin;  $Var(Y) = \mu + \alpha\mu^2$ ; when  $\alpha \rightarrow 0 \Rightarrow$  Poisson
- **ZIP/ZINB:** Excess zeros; two-part: Logistic (0 vs  $> 0$ ) + Poisson/NB (counts)

**NB vs Poisson Test (HW7):**

1. **Hypotheses:**  $H_0 : \alpha = 0$  vs  $H_a : \alpha \neq 0$  (no overdispersion vs overdispersion exists)
2. **Test:** t-test for  $\alpha$  parameter (in SAS output: "Dispersion" parameter)
3. **From SAS:** Find "Analysis Of Maximum Likelihood Parameter Estimates" table; look for "Dispersion" row
4. Value: estimate of  $\alpha$  (e.g.,  $\hat{\alpha} = 10.10$ ); Std Error; Wald 95% CI
5. **P-value:** from "Pr > ChiSq" column (e.g.,  $p < 0.0001$ )
6. **Decision:**  $p < 0.05 \Rightarrow$  reject  $H_0 \Rightarrow$  NB preferred;  $p > 0.05 \Rightarrow$  Poisson adequate
7. Example:  $\hat{\alpha} = 10.10$ ,  $p < 0.0001 \Rightarrow$  reject  $H_0$  at 95% level  $\Rightarrow$  NB more appropriate

**Rate Ratio Estimation (HW7):**

- **Point estimate:**  $\widehat{RR} = \exp(\hat{\beta})$  (same formula for both Poisson and NB)
- **95% CI:**  $\exp(\hat{\beta} \pm 1.96 \cdot SE)$  where SE from respective model
- **Compare models:** NB typically gives wider CI (larger SE) than Poisson due to accounting for overdispersion
- Example: Male vs Female absence rate ratio
  - Poisson:  $\widehat{RR} = e^{0.321} = 1.38$ , 95% CI: (1.25, 1.52)
  - NB:  $\widehat{RR} = e^{0.321} = 1.38$ , 95% CI: (1.15, 1.65) (wider!)

- **Conclusion change?** If Poisson CI excludes 1 but NB CI includes 1  $\Rightarrow$  significance changes (Poisson overconfident)

**E. Key Concepts & Formulas**

**Multinomial vs Ordinal:**

- Multinomial: unordered,  $(K - 1)$  equations, each with own  $\beta_j$
- Ordinal: ordered, PO  $\Rightarrow$  same  $\beta$  for all cutpoints
- PO test  $p < 0.05 \Rightarrow$  reject PO  $\Rightarrow$  use Multinomial

**Parameter & df count:**

- Multinomial: cont X:  $K - 1$  param; q-level X:  $(K - 1)(q - 1)$  param
- Ordinal: cont X: 1 param; q-level X:  $q - 1$  param
- PO test df:  $(K - 2)(q - 1)$

**CI for linear combination:**  $\hat{\beta}_1 - \hat{\beta}_2$ :  $SE = \sqrt{Var(\hat{\beta}_1) + Var(\hat{\beta}_2) - 2Cov(\hat{\beta}_1, \hat{\beta}_2)}$

**Model selection:** LRT for nested models (df=# param diff); AIC/BIC for non-nested; QIC for GEE

## III. Longitudinal Data

**A. GEE** (population-averaged)

$g\{E[Y_{ij}|X_{ij}]\} = \mathbf{X}_{ij}^T \beta$

**Working Correlation:**

Type	Matrix	Param
Independence	$\begin{pmatrix} 1 & 0 \\ 0 & 1 \end{pmatrix}$	0
Compound Symmetry	$\begin{pmatrix} 1 & \rho \\ \rho & 1 \end{pmatrix}$	1
AR(1)	$\begin{pmatrix} 1 & \rho \\ \rho & 1 \end{pmatrix} \rho^{ j-k }$	1
Unstructured	$\begin{pmatrix} 1 & \rho_{12} \\ \rho_{21} & 1 \end{pmatrix}$	$k(k - 1)/2$

**SE:** Robust/Sandwich (default, preferred); Model-based (if corr correct)

**Selection:** QIC (smaller better); cannot use AIC/BIC

**Interpretation: Marginal** — "at population level, treatment increases odds by  $e^{\beta}$ "

**Key:** Even if working correlation misspecified,  $\hat{\beta}$  consistent with robust SE; efficiency loss only

**B. Model Selection & Strategy**

**Correlation selection:** Try multiple structures (IND, EXCH, AR(1), UN); compare using QIC (smaller is better)

- **EXCH:** All correlations equal; good for repeated measures with no time order
- **AR(1):** Correlation decays with time distance; good for equally spaced time series
- **UN:** Most flexible but many parameters; may not converge with many time points
- **IND:** No correlation; baseline comparison

**Convergence:** UN may fail with many time points; use simpler structures (EXCH, AR)

**Time as Categorical (HW8):**  $E[Y_{ij}] = \beta_0 + \beta_1 X_{t_2} + \beta_2 X_{t_4} + \beta_3 X_{t_8} + \beta_4 trt + \beta_5 (X_{t_2} \times trt) + \beta_6 (X_{t_4} \times trt) + \beta_7 (X_{t_8} \times trt)$  where  $X_{t_j}$ =dummy for time  $j$

**Interaction Test (HW8):**  $H_0 : \beta_5 = \beta_6 = \beta_7 = 0$  vs  $H_a$ : not  $H_0$ ; Type 3 Wald test; df=# interaction terms

**Interaction in GEE (time continuous):**  $E[Y_{ij}] = \beta_0 + \beta_1 time + \beta_2 trt + \beta_3 (time \times trt)$

- $\beta_3$ : difference in time trend between treatment groups
- Treatment A change:  $\beta_1$ ; Treatment B change:  $\beta_1 + \beta_3$ ; Difference:  $\beta_3$

**Interpretation guidelines:**

- Marginal interpretation: "at population level, treatment increases odds by  $e^{\beta}$ "
- Always report with 95% CI
- Robust SE preferred (default in GEE)

**C. Step-by-Step Calculation Guide**

**1. KM Table Construction (HW1):**

1. List all unique event times (exclude censored-only times)
2. For each  $t_j$ : count  $n_j$  (at risk before  $t_j$ ),  $d_j$  (events at  $t_j$ )
3. Censored at  $t$ : include in  $n_j$  for  $t_j \leq t$ , exclude from  $n_{j+1}$  if  $t_j = t$
4. Compute  $p_j = 1 - d_j/n_j$ , then  $\hat{S}(t_j) = \prod_{i \leq j} p_i$

**2. Interaction HR with CI (HW5 - Complete Steps):**

1. Model:  $h = h_0 \exp(\beta_1 X_1 + \beta_2 X_2 + \beta_3 X_1 X_2)$ ; Compare:  $X_1 = 0$  vs  $X_1 = 1$  at  $X_2 = a$
2. **Point estimate:**  $HR = \frac{h(X_1=0, X_2=a)}{h(X_1=1, X_2=a)} = \exp(-\beta_1 - \beta_3 a)$ ; plug in  $\hat{\beta}$ 's
3. **Variance:**  $Var(-\hat{\beta}_1 - \hat{\beta}_3 a) = Var(\hat{\beta}_1) + a^2 Var(\hat{\beta}_3) + 2a \cdot Cov(\hat{\beta}_1, \hat{\beta}_3)$
4. Get  $Var(\hat{\beta}_1)$ ,  $Var(\hat{\beta}_3)$ ,  $Cov(\hat{\beta}_1, \hat{\beta}_3)$  from CovB matrix; compute variance
5.  $SE = \sqrt{Var}$
6. 95% CI:  $\exp[(-\hat{\beta}_1 - \hat{\beta}_3 a) \pm 1.96 \cdot SE]$

**Example (HW5):**  $HR = \exp(-1.09033 - 0.40491) = 0.224$ ;  $Var = 0.2302 + 0.4539 + 2(-0.227) = 0.2301$ ;  $SE = 0.4797$ ; CI=[0.088, 0.574]

**3. Ordinal Probability (HW6):**

1. Get  $\hat{\alpha}_j$  and  $\hat{\beta}$  from output
2. Compute  $P(Y \leq j) = \frac{\exp(\hat{\alpha}_j + \hat{\beta} X)}{1 + \exp(\hat{\alpha}_j + \hat{\beta} X)}$  for  $j = 1, \dots, K - 1$
3.  $P(Y = 1) = P(Y \leq 1)$ ;  $P(Y = j) = P(Y \leq j) - P(Y \leq j - 1)$  for  $j \geq 2$
4.  $P(Y = K) = 1 - P(Y \leq K - 1)$

**4. Quasi-Poisson CI Adjustment (HW7):**

1. Get  $\hat{\beta}$  and  $SE_{old}$  from Poisson output
2. Get Scale= $\phi$  (Deviance/df or Pearson  $\chi^2$ /df)
3.  $SE_{new} = SE_{old} \times \sqrt{\phi}$
4. New 95% CI:  $\exp(\hat{\beta} \pm 1.96 \cdot SE_{new})$

**5. Mean Variance Calculation (HW2):**

1. Compute  $\hat{\mu}$  using discrete formula
2. For each  $j$ , calculate  $A_j = \sum_{k=j+1}^J \hat{S}(t_{k-1})(t_k - t_{k-1})$  (remaining area)

3. Compute  $Var(\hat{\mu}) = \frac{n_d}{n_d - 1} \sum_{j=1}^{J-1} \frac{A_j^2 d_j}{n_j(n_j - d_j)}$  where  $n_d = \sum d_j$

4.  $SE = \sqrt{Var}$ ; 95% CI:  $\hat{\mu} \pm 1.96 \cdot SE$

**6. GEE Mean Change (HW8 - Categorical Time):**

1. Model:  $E[Y] = \beta_0 + \beta_1 X_{t_2} + \beta_2 X_{t_4} + \beta_3 X_{t_8} + \beta_4 trt + \beta_5 (X_{t_2} \times trt) + \beta_6 (X_{t_4} \times trt) + \beta_7 (X_{t_8} \times trt)$
2. **Trt A:** baseline= $\beta_0$ ; at time 2= $\beta_0 + \beta_1$ ; change= $\beta_1$
3. **Trt B:** baseline= $\beta_0 + \beta_4$ ; at time 2= $\beta_0 + \beta_1 + \beta_4 + \beta_5$ ; change= $\beta_1 + \beta_5$
4. **Difference in change:**  $(\beta_1 + \beta_5) - \beta_1 = \beta_5$  (this is the interaction coefficient!)
5. Interpretation:  $\beta_5$ =mean difference of temperature change from baseline to 2hrs between two treatments

**7. Log-log CI for S(t) (HW1):**

1. Compute  $Var[\log\{-\log(\hat{S})\}] = \frac{1}{[\log(\hat{S})]^2} \sum_{t_j \leq t} \frac{d_j}{n_j(n_j - d_j)}$
2.  $SE = \sqrt{Var}$ ; compute limits:  $L = \log[-\log(\hat{S})] - 1.96 \cdot SE$ ,  $U = \log[-\log(\hat{S})] + 1.96 \cdot SE$
3. 95% CI for  $S(t)$ :  $[\exp(-e^U), \exp(-e^L)]$

# Typical Exam Question Types (Based on HW1-8)

## I. Survival Analysis

- **KM table:** Construct 7-column table; handle censoring ( $\downarrow n_{j+1}$ , NOT  $\uparrow d_j$ )
- **Median & mean:** Find  $\min\{t_j : \hat{S}(t_j) < 0.5\}$ ; mean area under curve
- **Log-rank vs Wilcoxon:** Compare test stats; explain why differ (early weight)
- **Interaction HR:**  $\log(HR) = \beta_1 + \beta_3 x_2$ ;  $SE^2 = Var(\beta_1) + x_2^2 Var(\beta_3) + 2x_2 Cov(\beta_1, \beta_3)$
- **Survival prediction:**  $S(t|X) = [S_0(t)]^{\exp(\beta^T X)}$  using baseline
- **Group comparison:**  $Var(\beta_j - \beta_k) = Var(\beta_j) + Var(\beta_k) - 2Cov(\beta_j, \beta_k)$

## II. Categorical Data

- **PO test:** Check  $p$ -value;  $p < 0.05 \Rightarrow$  use Multinomial;  $p > 0.05 \Rightarrow$  use Ordinal
- **Probability calc:** Ordinal: cumulative then subtract; Multinomial: softmax formula
- **OR interpretation:** Ordinal: cumulative OR (same for all cut-points); Multinomial: RRR vs ref
- **df calculation:** Ordinal cont X:  $df=1$ ; Ordinal q-level X:  $df=q-1$ ; Multinomial:  $df=(K-1)(q-1)$
- **Overdispersion check:** Look at Scale (Dev/df or  $\chi^2/df$ ); if  $> 1.5 \Rightarrow$  serious problem
- **Quasi-Poisson:**  $\hat{\beta}$  stays same,  $SE_{new} = SE_{old} \times \sqrt{\phi}$ ; recalculate CI
- **NB vs Poisson:** Test  $H_0 : \alpha = 0$ ; compare point estimates and CI width
- **R/R interpretation:**  $e^{\beta}$  with 95% CI; state "adjusting for other

variables"

- **Conclusion change:** NB gives wider CI  $\Rightarrow$  may lose significance

## III. Longitudinal Data

- **Correlation structure:** Try EXCH, AR(1), UN; compare QIC (smaller better)
- **Time as categorical:** Multiple dummy variables; different  $\beta$  for each time
- **Interaction**  $time \times trt$ :  $\beta_3$ =difference in time trend between groups
- **Mean change calc:** Trt A:  $\beta_1$  (time main); Trt B:  $\beta_1 + \beta_3$ ; Diff:  $\beta_3$
- **Hypothesis test:** Test  $H_0 : \beta_3 = 0$  (or all interaction terms=0); use Type 3 Wald;  $df=\#$  interaction terms
- **Robust SE:** Always use (default); model-based SE assumes correct correlation