HANDY REFERENCE SHEET – HRP/STATS 261, Discrete Data

2x2 Contingency Tables

	Disease (D)	No Disease (~D)
Exposed (E)	a	b
Unexposed (~E)	с	d

 $\frac{\text{Measures of Association}}{\text{Risk Ratio}} = \frac{a/(a+b)}{c/(c+d)}$

95% CI: RR * exp $\left[\frac{-1.96\sqrt{\frac{1-a/(a+b)}{a}+\frac{1-c/(c+d)}{c}}\right]}{c}$, RR * exp $\left[\frac{+1.96\sqrt{\frac{1-a/(a+b)}{a}+\frac{1-c/(c+d)}{c}}\right]$

Odds Ratio = $\frac{ad}{bc}$

95% CI: OR * exp
$$\left[\frac{-1.96\sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}}}{1.96\sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}}} \right]$$
, OR * exp $\left[\frac{+1.96\sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}}}{1.96\sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}}} \right]$

Difference in Proportions:

H₀: $p_{D/E} - p_{D/\sim E} = 0$ [for case-control study: $p_{e/d} - p_{e/\sim d} = 0$] Test Statistic:

$$\frac{\hat{p}_{d/e} - \hat{p}_{d/e}}{\sqrt{\frac{(\vec{p}_{d/e})(1 - \vec{p}_{d/e})}{n_e} + \frac{(\vec{p}_{d/e})(1 - \vec{p}_{d/e})}{n_{ee}}}} \sim Z$$

95% CI:
$$(\hat{p}_{d/e} - \hat{p}_{d/e}) \pm 1.96 * \sqrt{\frac{(\hat{p}_{d/e})(1 - \hat{p}_{d/e})}{n_e} + \frac{(\hat{p}_{d/e})(1 - \hat{p}_{d/e})}{n_{ee}}}$$

SAS CODE:

proc freq data=yourdata; tables yourExposure*yourOutcome /measures cl; weight counts; *if you have grouped data; run;

2x2xK Contingency Tables

<u>Notation</u>: outcome=*d*; predictor=*e*; categorical covariate=k

<u>Steps</u>

- 1. Calculate crude OR_{d-e} (or RR_{d-e})
- 2. Calculate stratum-specific OR's: OR_{d-e/k=K}
- 3. If crude OR and stratum-specific OR's are all similar \rightarrow **STOP**. *k* is unlikely to be a confounder or an effect modifier, and you may use usual methods for 2x2 table (see page i), ignoring *k*.
- 4. If crude OR and stratum-specific OR's differ \rightarrow proceed to (a) or (b) below:
 - a) If stratum-specific OR's are similar to each other → suspect confounding
 (i) Apply Cochran-Mantel-Haenszel test of conditional independence:
 - H₀: *e* and *d* are conditionally independent.
 - Test Statistic:

Exposed Unexposed
$$c$$
 d $k_{i=1}^{k} (a_k - E(a_k))]^2$ χ_1^2 $E(a_k) = \frac{(a+b)_k(a+c)_k}{n_k}$; where n_k = total in stratum k $Var(a_k) = \frac{(a+b)_k(a+c)_k}{n_k}$; where n_k = total in stratum k $Var(a_k) = \frac{(a+b)_k(a+c)_k(b+d)_k(c+d)_k}{n_k^2(n_k-1)}$

(ii) Calculate Mantel-Haenszel (MH) summary OR (adjusted for confounding by k). If you rejected the null in (i), then MH OR should be ≠ 1.0. If you did not reject null in (i), then MH OR should be ≅1.0.

$$OR_{d-e/k} = \frac{\sum_{i=1}^{k} \frac{a_i d_i}{T_i}}{\sum_{i=1}^{k} \frac{b_i c_i}{T_i}} \qquad \qquad \frac{\text{For RR:}}{RR_{d-e/k}} = \frac{\sum_{i=1}^{k} \frac{a_i (c_i + d_i)}{T_i}}{\sum_{i=1}^{k} \frac{c_i (a_i + b_i)}{T_i}}$$

 b) If stratum-specific OR's differ from each other→ suspect interaction (effect modification)

(i) Apply Breslow-Day test of homogeneity of the OR's:

- H₀: stratum-specific OR's are equal (homogenous)
- Test Statistic: complex \rightarrow use computer software
- ➢ If you reject null→ effect modification is present. Report stratumspecific OR's
- ➢ If you fail to reject null → insufficient evidence of effect modification; calculate Mantel-Haenszel summary OR, as above.

```
SAS CODE:
proc freq data=yourdata;
    tables yourCovariate*yourExposure*yourOutcome /cmh;
    weight counts; *if you have grouped data;
run;
```

<u>Multi-way Contingency Tables:</u> *Notation: x, y, and z are categorical variables*

<u>Chi-square test of independence (*RxC* table):</u> H₀: *x*, *y*, and *z* are independent Test Statistic:

$$\sum_{cells} \frac{(Observed - Expected)^2}{Expected} \sim \chi^2_{(rows-1)(columns-1)}$$

Log-Linear Model

 $log(counts) = \alpha + \beta_x x + \beta_y y + \beta_z z + interactions if appropriate$

Odds ratio interpretation (if 2x2 table):

$$\log(OR_{d-e}) = \log(\frac{ad}{bc}) = \log a + \log d - \log b - \log c$$
$$\log(\frac{ad}{bc}) = (\alpha + \beta_d + \beta_e + \beta_{d^*e}) + (\alpha) - (\alpha + \beta_d) - (\alpha + \beta_e) = \beta_{d^*e}$$
$$\therefore OR = e^{\beta_{d^*e}}$$

<u>SAS CODE:</u> <u>Chi-square test</u> proc freq data=yourdata; tables yourX*yourY*yourK / chisq; weight counts; *if you have grouped data; run; <u>Log-linear models</u> proc genmod data=yourdata;

```
proc genmod data=yourdata;
    model total = yourX yourY yourK YourInteractions /
    dist=poisson link=log pred;
run;
```

Logistic Regression

Model:

$$\ln(\frac{p}{1-p}) = \alpha + \beta_1 x_1 + \beta_2 x_2 + \dots$$

Tests of Model Fit:

• Wald Test

Simplest form of the logistic likelihood (from 2x2):					
$l(\beta) = \left(\frac{e^{\alpha+\beta_e}}{1+e^{\alpha+\beta_e}}\right)^a x\left(\frac{1}{1+e^{\alpha+\beta_e}}\right)^b x\left(\frac{e^\alpha}{1+e^\alpha}\right)^c x\left(\frac{1}{1+e^\alpha}\right)^d$					
	Disease	No Disease			
Exposed	а	b			
Unexposed	с	d			

$$Z = \frac{\hat{\beta} - 0}{\text{asymptotic standard error } (\hat{\beta})}$$

• <u>Likelihood Ratio Test:</u> where full model has *n* parameters and reduced model has *n*-*p*

$$-2\ln\frac{L(reduced)}{L(full)} =$$
$$-2\ln(L(reduced)) - [-2\ln(L(full))] \sim \chi_p^2$$

Interpretation of Estimated Coefficients in Odds and Probabilities:

OR interpretation: $OR_{exposure} = e^{\beta_{exp}}$

OR interpretation in the presence of interaction (two binary predictors):

OR_{exposure/interacting factor present} =
$$e^{\beta_e + \beta_k + \beta_{e \times k}}$$

OR_{exposure/interacting factor absent} = e^{β_e}

Probability interpretation: P(D/E) = $\frac{e^{\alpha + \beta_e}}{1 + e^{\alpha + \beta_e}}$

```
SAS CODE:
proc logistic data = analysis;
    class YourPredictor_c (ref= "YourBaseline");
        * automatic dummy coding to get >1 OR against the reference group;
    model YourOutcome_c (event = "Yes") = YourPredictor / lackfit;
        *lackfit gives Hosmer and Lemeshow goodness of fit chi-square test;
    output out = OutDataSet p = Predicted;
        *outputs predicted probabilities to new dataset;
run;
```

Pair-Matched Data

Case	Matched-control		
	Exposed	Unexposed	
Exposed	а	b	
Unexposed	С	d	

Odds Ratio

$$OR = \frac{b}{c}$$

McNemar's Test

- H₀: exposure and disease are independent
- Test Statistic:

$$\frac{(b-c)^2}{b+c} \sim \chi_1^2$$

1:M Matched Data

<u>Matched Data</u> Unrestricted Predictors and Covariates Binary Outcome

Conditional Logistic Regression:

The simplest likelihood (from 2x2): $l(\beta) = \left(\frac{e^{\beta_e}}{1 + e^{\beta_e}}\right)^b x \left(\frac{1}{1 + e^{\beta_e}}\right)^c$

OR interpretation: $OR_{exposure} = e^{\beta_{exp}}$

SAS CODE:** SAS V9 ONLY
proc logistic data = yourData ;
 model YourOutcpme_c (event= "Yes") = YourPredictor1
 YourPredictor2 YourPredictor3...;
 strata MatchingVariable1 MatchingVariable2 / info;
 output out = OutDataset p = Predicted;
run;