#### **Test of No Exposure-Disease Association**

$$Z^{2} = \frac{\left[X - \widehat{E}(X \mid H_{0})\right]^{2}}{V\widehat{a}r(X \mid H_{0})} \sim \chi_{1}^{2}$$

Hο

 $C_1=C_0$ 

 $\mathbf{X}$   $\widehat{\mathbf{E}}(\mathbf{X} | \mathbf{H}_0)$   $\mathbf{Var}(\mathbf{X} | \mathbf{H}_0)$ 

# Methods for Count Data (closed cohort and cross-sectional studies)

$$\begin{array}{c|cccc} & E & \overline{E} & \\ cases & a & b & M_1 \\ Non-cases & N_1-a & N_0-b & M_0 \end{array}$$

Unstratified

# Methods for Person-Time Data (open cohort and closed cohort studies)

	Е	Ē	,		а	$\frac{N_1M_1}{T}$	$\frac{N_1N_0M_1}{T^2}$	Unstratified
cases	a	b	$\mathbf{M}_{1}$	<sub>1</sub> =  <sub>0</sub>   <sub>1</sub> /  <sub>0</sub> =1	-	•	•	
PT	N <sub>1</sub>	$N_0$	Т	I <sub>1</sub> -I <sub>0</sub> =0	$\sum a_{i}$	$\sum \frac{N_{1i}M_{1i}}{T_i}$	$\sum \frac{N_{1i}N_{0i}M_{1i}}{T_i^2}$	Stratified

### **Methods for Case-control Data**

	$E \overline{E}$			$N_1M_1$	$\underline{M_{\scriptscriptstyle 1}M_{\scriptscriptstyle 0}N_{\scriptscriptstyle 1}N_{\scriptscriptstyle 0}}$	
cases	a b M <sub>1</sub>	OR=1	a	T	$T^2(T-1)$	Unstratified
controls	$\begin{bmatrix} c & d \\ N_1 & N_0 \end{bmatrix}$	I <sub>1</sub> /I <sub>0</sub> =1	$\sum a_i$	$\sum \frac{N_{1i}M_{1i}}{T}$	$\sum \frac{M_{1i} M_{0i} N_{1i} N_{0i}}{T_{i}^{2} \big( T_{i} - 1 \big)}$	Stratified

# **Test of Homogeneity of Effect Measures**

$$H = \sum_{i=1}^{l} \frac{\left[\hat{X}_i - \hat{X}_{summary}\right]^2}{V_0^2 r_i \left[\hat{X}_i\right]} \sim \chi_{l-1}^2$$

	H <sub>0</sub>	$\mathbf{\hat{X}}_{i}$	$\hat{X}_{summary}$	$V\widehat{a}r_{i}(\widehat{X}_{i})$			
Methods for Count Data (closed cohort and cross-sectional studies)							
Difference measure	$CID_1=CID_2==CID_i$ $CID_i=CID_j$ for all i,j	$CID_{i}$	$CID_{summary}$	$\frac{a_{i}c_{i}}{N_{1i}^{3}}+\frac{b_{i}d_{i}}{N_{0i}^{3}}$			
Ratio measure	$CIR_1$ = $CIR_2$ == $CIR_i$ $CIR_i$ = $CIR_j$ for all i,j	ln (CIR <sub>i</sub> )	<i>ln</i> (CIR <sub>MH</sub> )	$\frac{c_{i}}{a_{i}N_{1i}} + \frac{d_{i}}{b_{i}N_{0i}}$			
Methods for Person-	Methods for Person-Times Data (open cohort and closed cohort studies)						
Difference measure	IRD <sub>1</sub> =IRD <sub>2</sub> ==IRD <sub>i</sub> IRD <sub>i</sub> =IRD <sub>j</sub> for all i,j	$IRD_{i}$	$IRD_{summary}$	$\frac{a_{i}}{N_{1i}^{2}} + \frac{b_{i}}{N_{0i}^{2}}$			
Ratio measure	IRR <sub>1</sub> =IRR <sub>2</sub> ==IRR <sub>i</sub> IRR <sub>i</sub> =IRR <sub>j</sub> for all i,j	<i>ℓn</i> (IRR <sub>i</sub> )	<i>ln</i> (IRR <sub>MH</sub> )	$\frac{1}{a_i} + \frac{1}{b_i}$			
Methods for Case-control Data							
Ratio measure	OR <sub>1</sub> =OR <sub>2</sub> ==OR <sub>i</sub> OR <sub>i</sub> =OR <sub>j</sub>	<b>ln</b> (OR <sub>i</sub> )	<i>ln</i> (ОRмн)	$\frac{1}{a_i}+\frac{1}{b_i}+\frac{1}{c_i}+\frac{1}{d_i}$			

#### **Relative Excess Risk due to Interaction**

for all i,j

$$RERI = CIR_{11}-CIR_{10}-CIR_{01}+1$$

#### E-value

E-value = 
$$RR + \sqrt{RR(RR - 1)}$$

If RR < 1, take the reciprocal before computing the E-value

#### **Confidence Intervals for Ratio and Difference Measures**

$$X\pm Z_{1-\alpha/2}\sqrt{V\widehat{a}r(X)}$$

 $\mathbf{X}$   $\mathbf{w}_i$   $\mathbf{V}\widehat{\mathsf{ar}}(\mathsf{X})$ 

# Methods for Count Data (closed cohort and cross-sectional studies)

$$\frac{a}{N_1} - \frac{b}{N_0}$$

$$\frac{ac}{N_1^3} + \frac{ba}{N_0^3}$$

Summary cumulative incidence difference

$$\frac{\sum w_i \! \left[ \frac{a_i}{N_{1i}} \! - \! \frac{b_i}{N_{0i}} \right]}{\sum w_i}$$

$$\frac{N_{1i}N_{0i}}{T_i}$$

$$\frac{\sum \left(\frac{a_i c_i N_{0i}^2}{T_i^2 (N_{1i} - 1)} + \frac{b_i d_i N_{1i}^2}{T_i^2 (N_{0i} - 1)}\right)}{\left(\sum \frac{N_{1i} N_{0i}}{T_i}\right)^2}$$

Cumulative incidence ratio ( $\ell n$ )

$$\ln \left\{ \frac{a}{N_1} / \frac{b}{N_0} \right\}$$

$$\frac{c}{aN_1} + \frac{d}{bN_0}$$

Summary cumulative incidence ratio (*ln*)

$$\ln \left\{ \frac{\sum w_i \left[ \frac{a_i}{N_{1i}} \middle/ \frac{b_i}{N_{0i}} \right]}{\sum w_i} \right.$$

$$\frac{b_i N_{1i}}{T_i}$$

$$\frac{\sum (M_{1i}N_{1i}N_{0i} - a_ib_iT_i)/T_i}{\left[\sum \frac{a_iN_{0i}}{T_i}\right]\sum \frac{b_iN_{1i}}{T_i}}$$

# Methods for Person-Time Data (open cohort and closed cohort studies)

Rate difference

$$\frac{a}{N_1} - \frac{b}{N_0}$$

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$$\frac{a}{N_1^2} + \frac{b}{N_0^2}$$

Summary rate difference

$$\frac{\sum w_i \left[ \frac{a_i}{N_{1i}} - \frac{b_i}{N_{0i}} \right]}{\sum w_i}$$

 $\frac{N_{1i}N_{0i}}{T_i}$ 

$$\frac{\sum \left(\frac{a_i N_{0i}^2 + b_i N_{1i}^2}{T_i^2}\right)}{\left(\sum \frac{N_{1i} N_{0i}}{T_i}\right)^2}$$

Rate ratio (*ln*)

$$\ln \left\{ \frac{a}{N_1} / \frac{b}{N_0} \right\}$$

--

$$\frac{1}{a} + \frac{1}{b}$$

Summary rate ratio (*ln*)

$$\ln \left\{ \frac{\sum w_i \left[ \frac{a_i}{N_{1i}} \middle/ \frac{b_i}{N_{0i}} \right]}{\sum w_i} \right]$$

 $\frac{b_{_{i}}N_{_{1i}}}{T_{_{i}}}$ 

$$\frac{\sum (M_{1i}N_{1i}N_{0i})/T_i^2}{\left[\sum \frac{a_iN_{0i}}{T_i}\right]\!\!\left[\sum \frac{b_iN_{1i}}{T_i}\right]}$$

### **Methods for Case-control Data**

Odds ratio (*ln*)

$$\ln \left\{ \frac{\mathsf{ad}}{\mathsf{bc}} \right\}$$

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$$\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}$$

Summary odds ratio (*ln*)

$$\ln \left\{ \frac{\sum w_i \frac{a_i d_i}{b_i c_i}}{\sum w_i} \right\}$$

 $\frac{b_i c_i}{T_i}$ 

RGB variance

# **Computational Form of the Mantel Haenszel Estimators**

#### Methods for Count Data (closed cohort and cross-sectional studies)

$$\hat{CIR}_{MH} = \frac{\sum_{i=1}^{I} \frac{a_{i} N_{0i}}{T_{i}}}{\sum_{i=1}^{I} \frac{b_{i} N_{1i}}{T_{i}}}$$

$$\frac{\sum \left(\frac{a_i N_{0i} - b_i N_{1i}}{T_i}\right)}{\sum \frac{N_{1i} N_{0i}}{T_i}}$$

# Methods for Person-Time Data (open cohort and closed cohort studies)

$$\hat{IRR}_{MH} = \frac{\displaystyle\sum_{i=1}^{I} \frac{a_{i}N_{0i}}{T_{i}}}{\displaystyle\sum_{i=1}^{I} \frac{b_{i}N_{1i}}{T_{i}}} \qquad \qquad \text{Summary rate} \\ \text{difference}$$

$$\frac{\sum \left(\frac{a_i N_{0i} - b_i N_{1i}}{T_i}\right)}{\sum \frac{N_{1i} N_{0i}}{T_i}}$$

#### **Methods for Case-control Data**

$$\hat{O}R_{MH} = \frac{\sum_{i=1}^{I} \frac{a_{i}d_{i}}{T_{i}}}{\sum_{i=1}^{I} \frac{b_{i}c_{i}}{T_{i}}}$$

#### Formulae for 1:1 matched case-control studies:

Cases 
$$\begin{bmatrix} & & & & & & & \\ & E & & E & \\ \hline E & & & & & \end{bmatrix}$$

$$OR_{MH} = f_{10}/f_{01}$$

$$H_O$$
:  $OR_{MH} = 1$   
 $H_A$ :  $OR_{MH} \neq 1$ 

$$Z^{2} = \frac{(f_{10} - f_{01})^{2}}{f_{10} + f_{01}} \sim \chi_{1}^{2}$$

Variance formula for confidence interval:

$$Var[\ln(OR_{MH})] = \frac{1}{f_{10}} + \frac{1}{f_{01}}$$



#### **Computational Form for Standardization**

Indirectly standardized rate ratio: 
$$SMR = \frac{\sum a_i}{\sum (N_{1i})I_{0i}}$$

Directly standardized rate ratio: 
$$SRR = \frac{\sum (N_{0i}) I_{1i}}{\sum b_i}$$

Indirectly standardized rate difference: 
$$SMD = \frac{\sum a_i}{\sum N_{1i}} - \frac{\sum (N_{1i})(I_{0i})}{\sum N_{1i}}$$

Directly standardized rate difference: 
$$SRD = \frac{\sum (N_{0i})(I_{1i})}{\sum N_{0i}} - \frac{\sum b_i}{\sum N_{0i}}$$

### **Misclassification Correction**

Observed					
	Е	Ē			
cases	Α	В	m <sub>1</sub>		
controls	С	D	$m_0$		
	N <sub>1</sub>	No	Т		

Truth				
	Е	E		
cases	а	b	m <sub>1</sub>	
controls	С	d	$m_0$	
	N <sub>1</sub>	N <sub>0</sub>	Т	

#### **Truth**

	E	Ē	
cases	specificity *m <sub>1</sub> -B	sensitivity*m <sub>1</sub> – A	m <sub>1</sub>
	$a = \frac{1}{\text{sensitivity} + \text{specificity} - 1}$	$b = \frac{1}{\text{sensitivity} + \text{specificity} - 1}$	
controls	specificity * m <sub>0</sub> – D	sensitivity * m <sub>0</sub> – C	m <sub>0</sub>
	$c = \frac{1}{1}$ sensitivity + specificity $-1$	$u - \frac{1}{\text{sensitivity} + \text{specificity} - 1}$	
	N <sub>1</sub>	N <sub>0</sub>	Т