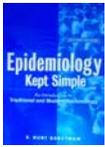


## Epidemiology Kept Simple



Chapter 12  
Error in Epidemiologic Research

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## §12.1 Introduction

- Analytic studies are exercises in **measurement**
- All measurements are **prone to error**
- Understanding **measurement error** begins by distinguishing between parameters and estimates



Train wreck

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## Parameters and Estimates

<p><b>Parameters</b></p> <ul style="list-style-type: none"> <li>- Error-free</li> <li>- Hypothetical</li> <li>- Causal effect</li> </ul>	<p><b>Statistical estimates</b></p> <ul style="list-style-type: none"> <li>- Error-prone</li> <li>- Calculated</li> <li>- Association</li> </ul>
--	--

Epidemiologic statistics are merely approximations of the parameters they wish to estimate.

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## Target Analogy

Let  $\phi$  represent the RR parameter.

Replicate the study many times

Let  $\hat{\phi}_i$  represent the RR estimate from study  $i$

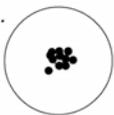
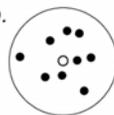
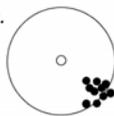
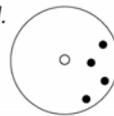


How closely do the estimates reflect the parameter? How accurate is a given estimate?

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## Target Analogy

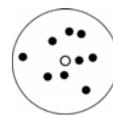
Random Error (Imprecision)

	Low	High
Low	a. 	b. 
High	c. 	d. 

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## Random Error (Imprecision)

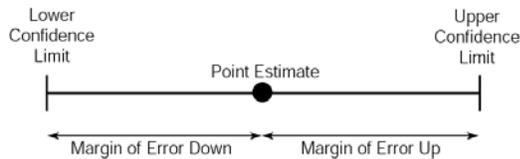
- Balanced "scatter"
- Use statistical methods such as confidence intervals and hypothesis tests of significance to address random error




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## Confidence Intervals (CI)

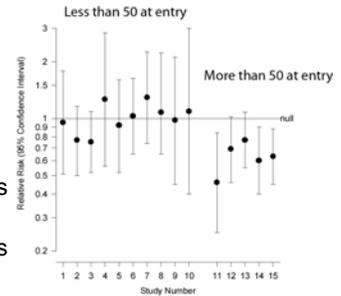
- Surround point estimate with **margin of error**
- Locates** parameter with given level of confidence (e.g., 95% confidence)
- CI width** quantifies precision of the estimate (narrow CI → precise estimate)



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## Mammographic Screening and Breast Cancer Mortality

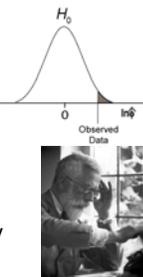
- E = mammographic screening
- D = Breast CA death
- First series: women in their 40s
- Second series: women in their 50s



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## P-values In a Nutshell

- Start with this null assumption:  $H_0$ : RR = 1 (no association)
- Small  $P$ -value provide **evidence against  $H_0$**  (do no use  $P$ -value cutoffs -- surely, G-d loves  $P = .06$ )
- By **convention**
  - $P \leq .10$  is considered marginally significant evidence against  $H_0$
  - $P \leq .01$  is considered very significant evidence against  $H_0$



R.A. Fisher

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## Childhood SES and Stroke

Factor	RR	P-value
Crowding (persons/room)	< 1.5 = 0.4	trend $P = 0.53$
	1.5 – 2.49 = 1.0 (ref.)	
	2.5 – 3.49 = 0.6	
Tap water	0.73	$P = 0.53$
Toilet type	flush/not shared = 1.3	trend $P = 0.67$
	flush/shared = 1.0 (referent)	
	no flush = 1.0	
Ventilation	good = 1.0 (ref.)	trend $P = 0.08$
	fair = 1.7	
	poor = 1.7	
Cleanliness	good = 1.1	trend $P = 0.07$
	fair = 1.0 (ref.)	
	poor = 0.5	

Nonsignif. evidence against  $H_0$

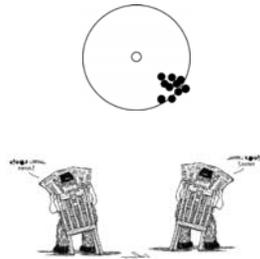
Marginally significant evidence against  $H_0$

Source: Galobardes et al., *Epidemiologic Reviews*, 2004, p. 14

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## Systematic Error (Bias)

- Bias** = systematic error in inference (*not* an imputation of prejudice)
- Direction** of bias
- Toward the null** (underestimates risks or benefits)
- Away from null** (overestimates risks or benefits)

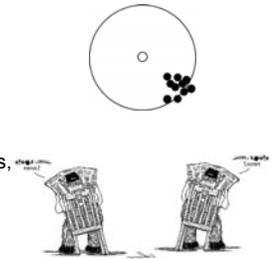


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## Categories of Bias

- Selection bias:** participants selected in a such a way as to favor a certain outcome
- Information bias:** misinformation favoring a particular outcome
- Confounding:** extraneous factors unbalanced in groups, favoring a certain outcome



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### Examples of Selection Bias

- Berkson's bias
- Prevalence-incidence bias
- Publicity bias
- Healthy worker effect
- Convenience sample bias
- Read about it: pp. 229 – 231



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### Examples of Information Bias

- Recall bias
- Diagnostic suspicion bias
- Obsequiousness bias
- Clever Hans effect
- Read about it: p. 231



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### The Misinformation Effect

- Memory is constructed rather than played back
- Social contagion of memory
- The misinformation effect: Elizabeth Loftus ([http://en.wikipedia.org/wiki/Misinformation\\_effect](http://en.wikipedia.org/wiki/Misinformation_effect))



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### Recall Bias in Case-Control Studies

	Cases	Controls
Exposure reported	a ↑	b
Exposure not reported	c	d ↓

Cases recall past exposure better than controls & controls tend to underreport their past exposures: ↑ a or ↓ d in case-control studies → ↑ OR

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### Differential & Nondifferential Misclassification

- **Non-differential misclassification:** groups equally misclassified ⇒ biases results toward null
- **Differential misclassification:** groups unequally classified ⇒ biases in either direction



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**Table 17.6** Examples of nondifferential and differential misclassification, and their effect on relative risk estimates.

**Scenario A:** Nondifferential misclassification: 10% of the cases in the exposed and nonexposed groups are misclassified as noncases.

	Accurate data		Total	→	Misclassified data		Total
	+	-			+	-	
Exposed	100	900	1000		90	910	1000
Nonexposed	50	450	500		45	455	500

$\widehat{RR} = \frac{100/1000}{50/500} = 1.00$        $\widehat{RR} = \frac{90/1000}{45/500} = 1.00$

**Scenario B:** Differential misclassification: none (0%) of the cases are misclassified in the exposed group, while 10% are misclassified in the nonexposed group.

	Accurate data		Total	→	Misclassified data		Total
	+	-			+	-	
Exposed	100	900	1000		100	900	1000
Nonexposed	50	450	500		45	455	500

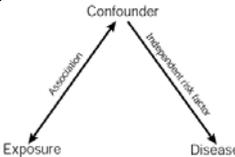
$\widehat{RR} = \frac{100/1000}{50/500} = 1.00$        $\widehat{RR} = \frac{100/1000}{45/500} = 1.11$

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## Confounding



- **Confounding** ("to mix together")  $\equiv$  a distortion brought about by extraneous variables
- Properties of confounder:
  - Confounder is associated with the exposure
  - Confounder is an independent risk factor
  - Confounder is *not* in causal pathway



## Accident Evacuation Crude Analysis

	Died	Survived	Total
Helicopter	64	136	200
Road	260	840	1100

- $R_1 = 64 / 200 = .3200$
- $R_0 = 260 / 1100 = .2364$
- $RR = .3200 / .2364 = 1.35$
- 35% increase in risk of death? This makes no sense!

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## Accident Evacuation Serious Accidents (Stratum 1)

	Died	Survived	Total
Helicopter	48	52	100
Road	60	40	100

- $R_1 = 48 / 100 = .4800$
- $R_0 = 60 / 100 = .6000$
- $RR_1 = .48 / .60 = 0.80$
- Helicopter evacuation cut down on risk of death by 20% among serious accidents

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## Accident Evacuation Minor Accidents (Stratum 2)

	Died	Survived	Total
Helicopter	16	84	100
Road	200	800	1000

- $R_1 = 16 / 100 = .16$
- $R_0 = 200 / 1000 = .20$
- $RR_2 = .16 / .20 = 0.80$
- Helicopter evacuation cut down on risk of death by 20% among minor accidents

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## Accident Evacuation Properties of Confounding

- Seriousness of accident (C) associated with method of evacuation (E)
- Seriousness of accident (C) is an independent risk factor for the outcome (D)
- Seriousness of accident (C) *not* in the causal pathway (the helicopter evaluation does not cause the accident to become more serious)

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