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#### **Basic Concepts in Epidemiology**

• What is Epidemiology? IT IS NOT THE STUDY OF THE SKIN!!

It is the study of the distribution and determinants of health-related states or events in specified populations, and the application of this study to control of health problems.

#### How do diseases evolve in individuals over time?

• Disease states:

Time -

- Many diseases afflict an individual over time.
- Some are <u>recurrent</u> person is diseased for a while, then recovers, but may experience the disease again.
- Examples: Urinary tract infection, depression, fracture of radius

#### How do diseases evolve in individuals over time?

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• Disease states:

Time -

Some are <u>non-recurrent</u> – an individual can develop the disease at most once

Diseased

- Diseases may be irreversible
- Examples: Alzheimer's Disease, Osteoarthritis, Coronary Heart Disease

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#### How do diseases evolve in individuals over time?

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- Disease states:
  - Some are <u>non-recurrent</u> an individual can develop the disease at most once
    - Recovery may occur with development of permanent immunity.

Immune

Time ------

• Examples: Measles, Appendicitis

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Diseased

#### How do diseases evolve in individuals over time?

General Model:

- Above patterns of disease behavior can be regarded as special cases of a more general model.
- At any given time, an individual can be in one of three states:
  - 1. Diseased: Currently afflicted.
  - 2. <u>Susceptible</u>: not currently afflicted, but capable of developing the disease in the next moment of time.
  - 3. <u>Immune</u>: not currently afflicted, and incapable of developing the disease in the next moment (and possible longer).

#### **How do diseases evolve in individuals over time?** General Model:

– Examples:

- Uterine cancer men are immune.
- Benign prostatic hypertrophy women are immune.
- Measles those who become ill (or immunized) develop protective antibodies.
- Polio oral vaccines confer immunity.
- "At-risk"
  - Susceptible individuals are termed "at-risk" (there may be wide variability in risk among them, however).
  - Immune and diseased individuals are termed "not atrisk" – they have no chance of developing the disease anew in the next moment of time.

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#### How do diseases evolve in individuals over time?

Disease events:

Time

How

n

- Some diseases are most naturally regarded as instantaneous occurrences at a particular point in time. Unlike states, they have negligible durations. Examples:
  - Rupture of aortic aneurysm.

Dise

occurs

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- Spontaneous abortion.
- Fall in an elderly person.







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Diseases in Populations
are populations "defined"?
by specifying criteria for membership. These criteria nay include:
Geographic boundaries within which individuals must reside.
Membership in a predefined group.
Specification of a time or time period of interest.
Examples:
• Residents of Salt Lake City, UT on January 1, 2005.
• Attendees of a certain church picnic.
• All alumni of the U of U School of Medicine as of

 All alumni of the U of U School of Medicine as of January 1, 2005.

#### **Diseases in Populations**

- In a population that is observed <u>over time</u>, membership in the population may or may not change.
  - If no changes in membership occur, it is called a <u>Closed Population</u> (or "fixed cohort"). Examples:
    - Persons involved in a foodborne illness outbreak that occurs over a short period of time.
    - Clinical trials participants whose mortality experience is followed for the duration of the trial, with no losses to follow-up.

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#### **Diseases in Populations**

- If membership of the population changes during the observation period, it is called an <u>Open</u> <u>Population</u> (or "dynamic population").
  - People may join e.g., births, in-migration.
  - People may leave e.g., deaths, out-migrants.
  - Examples:
    - Population of Salt Lake City during 2004.
    - Employees of the state government during 2000-2004.
    - Enrollees of Intermountain Health Care during 2000-2004.

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 Measurements in Epidemiology

 • Define and use

 - Ratio

 - Proportion

 - Odds

 - Rate

 - Nate

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 Measurements in Epidemiology

 • Define and use

 - Proportion

 - Odds

 - Rate

 Measurements in Epidemiology

 • Define and use

 - Prevalence

 - Incidence

 - Cumulative incidence (CI), Incidence proportion

 - Attack rate (AR)

 - Incidence density (ID), Incidence rate

## Measurements in Epidemiology

- **Count** The number of individuals with the outcome of interest
- **Proportion** The number of individuals within a defined group with the outcome of interest divided by the number of individuals enumerated in the population
- **Rate** The number of individuals within a defined group with the outcome of interest divided by the number of individuals enumerated in the population who are followed for a specified period of time

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• Ratio - One number divided by another number











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Suppose someone says: "Seven children in the city have developed leukemia in the last year."

- Is seven too many?
- It depends on the size of the city!
- The proportion of children with leukemia gives more information than the number of children with leukemia.

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## Proportion

- The quotient of 2 numbers
- Numerator NECESSARELY INCLUDED in the denominator
- Quantities have to be of same nature
- Proportion always ranges between 0 and 1
- Percentage = proportion x 100



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#### **Proportion, Examples**

- Proportion of male in 8 classes
  - 101 students, 9 male
  - Proportion of male student = 0.09
  - Percentage of male student = 9%



#### **Proportions and Probabilities**

- We often interpret proportions as probabilities. If the **proportion** with a disease is 1/10 then we also say that the **probability** of getting the disease is 1/10, or 1 in 10.
- Proportions are usually quoted for **samples** probabilities are usually quoted for **populations**.

**Uses of Proportion** 

As a descriptive statistic:

- The proportion of women in college.
- The percent of economy cars.

As a measure of morbidity

- Disease prevalence
- Disease incidence
- Mortality rate (death rate)

As parameters to characterize screening tests:

- Sensitivity of a screening test
- Specificity of a screening test

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#### Case-Control Design: An Example

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• A case-control study undertaken to identify the reasons for the exceptionally high rate of lung cancer among male residents of coastal Georgia.

Cases (identified from these 3 sources) a. Diagnosis since 1970 at the single large

hospital in Brunswick.

b. Diagnosis during 1975-1976 at three major hospitals in Savannah

c. Death certificates for the period 1970-1974 in the area

#### Proportion as a Descriptive Statistic: Case-Control Design

Controls – selected from admissions to four hospitals and from death certificates from the same period for diagnosis other than lung cancer, bladder cancer or chronic lung cancer.

Risk Factor – "shipbuilding", employment in the shipyards during World War II.

Confounding Factor - an exposure itself, not under investigation but related to the disease and the exposure. In this case, smoking is a confounding factor.

Proportion as a Descriptive Statistic: Case-Control Design

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- There was a high rate of lung cancer among male residents of coastal Georgia.
- Cases (people with lung cancer) were identified from hospital records.
- Controls (people without lung cancer or bladder cancer) were selected from the same hospital.
- Risk factor: exposure to shipyards during World War II.

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• Confounding variable: smoking (effect modifier)

	n						
	Smokers						
Smoking	Shipbuilding	Cases	Controls				
No	Yes	11	35				
	No	50	203				
Yes	Yes	84	45				
	No	313	270				
For the cases:							
– Propor	tion of exposure	=84/397=0	0.212 or 21.				
For the controls:							
- Propor	tion of exposure	=45/315=0	0.143 or 14.				

#### Shipbuilding Example Smokers

The results for smokers reveal different exposure histories for cases and controls.

The proportion of exposure to shipbuilding among cases (21.2%) was higher than among controls (14.3%). The difference is 6.9%.

It is not conclusive *proof*; however, it is a *good clue* indicating a possible relationship between disease and exposure.

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Shipbuilding Example Nonsmokers						
Smoking	Shipbuilding	Cases	Controls			
No	Yes	11	35			
	No	50	203			
Yes	Yes	84	45			
	No	313	270			
For the cases:						
<ul> <li>Proportion of exposure=11/61=0.180 or 18.0%</li> </ul>						
For the co	ontrols:					
– Propor	tion of exposure=	=35/238=0	.147 or 14.'			

#### Shipbuilding Example Nonsmokers

The results for nonsmokers reveal different exposure histories for cases and controls.

The proportion of exposure to shipbuilding among cases (18.0%) was higher than among controls (14.7%). The difference is 3.3%.

It is not conclusive *proof*; however, it is a *good clue* indicating a possible relationship between disease and exposure.

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### Shipbuilding Example: Terminology Review

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- The term *exposure* refers to a suspected *risk factor*. The term exposure is also used when the factor has beneficial effects. This is called a *protective factor* (flouride).
- The *confounder* is not under investigation, but may be related to both disease and exposure. It may act as an *effect modifier*, altering the effect of an exposure.

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• The main effects were different: 6.9% among smokers versus 3.3% among nonsmokers.

 Example

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#### Rates

- The term *rate* is often used interchangeably with the term proportion in epidemiology although sometimes it refers to a quantity of a very different nature.
- Types of rates we will cover:
- Change rates
- Death rate
- Incidence rate
- Follow-up death rate

### **Change Rates**

• These types of rates are used to describe changes after a certain period of time.

changerate(%) =  $\frac{\text{new value-old value}}{\text{old value}} X 100$ 

- Example: A total of 35,238 new AIDS cases were reported in 1989 compared to 32,196 reported during 1988.
  - The change rate for new AIDS cases:

 $\left(\frac{35,238-32,196}{32,196}\right)100 = 9.4\%$ 

Rate

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Infant mortality rate in one year

# of Infant deaths at a defined stage of gestation
# of live births in one year

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## **Rates vs. Proportions**

We must have a clear definition of the denominator to determine if we have a rate or and proportion.

- Rate : denominator contains units of exposure (person-years).
- Proportion: denominator contains a count or the number of individuals at risk (or exposed).

Difference Difference between Two Probability				
Baseball Games	Won	Lost	Total	
Faiwan Team China Team	14 8	6 12	20 20	
Proba	bility Diffe	rence =	: 0.7- 0.4 = 0	).3
	,			_



	Udds Katio					
	Odds_1 / Odds_2					
Baseball Games	Won	Lost	Total			
Taiwan Team China Team	14 8	6 12	20 20			
(14 Odds Ratio =	/ 20) / ( 6	/20)	= 3.5			
	20)/(12	2/20)				



# **Measurements in Epidemiology** Epidemiology involves estimating frequency and distribution of diseases in populations and comparing the effect of suspected risk factors on the frequency of disease

#### **Measuring Disease Frequency**

- Measures of disease frequency are tools used to describe how common an illness (or other outcome) is with reference to the size of the population (population at risk).
- Used to quantify cases in relation to the population and a measure of time.
- 2 main measures:
  - Prevalence
  - Incidence

#### **Measures of Disease Occurence**

- 1. Prevalence
  - Measures population disease status proportion
  - (Point) Prevealence
  - Lifetime prevalence
  - Period Prevalence
- 2. Incidence

Assess frequency of disease onset

- Cumulative incidence (incidence proportion)
- Incidence density (incidence rate)

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#### **Two Measures of Incidence**

- The **proportion** of individuals who experience the event (E) in a defined time period (E/N during some time T) = **cumulative incidence**
- The number of events divided by the amount of person-time observed (E/NT)
   = incidence rate or density (not a proportion)

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# Measures of Disease Frequency - Prevalence Measures frequency of disease in a defined population at a *specified point in time*. Fundamentally a static measure of disease frequency – a "snapshot" view, with time frozen.

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#### **Measures of Disease Frequency - Prevalence**

- The "Point in time" at which prevalence is determined may refer to any of several time scales. Examples:
  - Calendar time e.g., prevalence of HIV infections in Utah on January 1, 2005.
  - Age e.g., prevalence of HIV infections among 20-year-old military recruits (regardless of whether they achieved age 20 in calendar time).
  - Time since some event e.g., prevalence of depression among widows/widowers 6 months after the death of a spouse.

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(	(Point) Prevalence	
Number o	of cases of disease at a specific time	
Рор	oulation exposed at that time t	-
• Proportion of	a population affected by a	
disease at	a given time	
• Expressed as a	ı percentage	
Example of lung	g cancer in a community, Jan 1, 19	80:
Population	3,500,000	
Cases	95,000	
Prevalence	2.71%	
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#### **Prevalence of URT infection in class**

- 42 students
- 7 with URT
- Prevalence: 7/42 = 0.17 or 17%





- Applies to events...
  - The disease may itself be an event (e.g. aneurysm rupture)
  - Or the onset of a disease state may be treated as the event of interest.











#### **Measures of Disease Frequency - Incidence** Provides an estimate of the probability (or risk) that a person selected at random from this population time period. develops this disease during the specified time period Can be measured directly only in a closed population. Example: "43% of persons who ate the potato salad And. became ill within 6 hours thereafter." Aliases: attack rate, incidence proportion, risk. Then, Jeff Lin. MD., PhI



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The probability of occurrence of an outcome in an outcome free population during a specified

If d = number of new cases

N = population initially at risk,

Risk (over a defined period) = d/N

Risk is usually applied to non-recurrent diseases or to the first episode of a disease.

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## **Incidence Density (incidence Rate)**

Number of NEW cases of disease during a period

Total person-time of observation

Rate

Instantaneous concept (like speed)

Denominator:

- is a measure of time
- the sum of each individual's time at risk
  - and free from disease

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#### **Person-Years**

- Person-years: the sum of the amount of time each individual is observed while *free of disease*.
- Each subject may contribute a different amount of person-years.

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#### **Person-Years**

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Time begins when they are enrolled in study.

- Time end with one of three events:
  - 1. Subject still alive on the analysis date.
  - 2. Subjects who died on a known date within the study period.

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3. Subjects who are lost to follow-up after a certain date.

#### **Person-Time at Risk**

- Person-time at risk is the denominator for rates of disease
- 1000 person-years at risk =
  - 100,000 people for 1/100 years

10,000 people for 1/10 years

- 1000 people for 1 years
- 100 people for 10 years
- 20 people for 50 years



#### **Smoking : Pack-Year**

- Pack-year smoking for a subject at risk
- 1 x 365 pack-year =

0.5 x 365 for 2 years

2 x 365 for 0.5 years



#### **Measures of Disease Frequency - Incidence**

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• Sometimes detailed information about time at risk is unavailable for each member of the population. It may nevertheless be possible to estimate total persontime at risk by multiplying *the average size of the population at risk* by *duration of the observation period*.

Total person-time ~ (Avg. size of population at risk) X (Length of observation period)

 Often the mid-period size of the population at risk is used as an estimate of the average population at risk.





#### **Incidence Rate**

• Incidence Rate (Incidence Density): the number of new cases of disease during a defined period of time, divided by the total person-time of observation.

IR = Number of new cases of disease during

Total person-time of observation

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#### Attack Rate (Cumulative Incidence, Incidence Proportion)

• Cumulative incidence during an outbreak Usually expressed for the entire epidemic period, from the first to the last case

Ex: Outbreak of cholera in country X in March 1999

- Number of cases = 490
- Population at risk = 18,600
- Attack rate = 2.6%

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#### **Cumulative Incidence and Incidence Rate**

- CI can be calculated from IR
- When incidence is very low or the time of observation is very short

CI = IR x Duration Time

- Assumes the PAR is constant
- Homicide in NY: 8.7/100,000 person-years
- Annual risk: 8.7/100,000

#### Relationship Between Incidence and Prevalence

Prevalence depends on the duration of the disease (T) and the rate of disease (ID). When the disease is stable (both incidence rate and duration are stable) then prevalence approximates the product of mean duration and incidence (CI).

Prevalence =  $CI \times T$ 

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#### Proportion as a Measure of Morbidity: Prevalence

*Disease Prevalence* = the proportion of people with a given disease at a given time.

Disease Prevalence =

Number of diseased persons at a given time Total number of persons examined at that time

Prevalence is usually quoted as per 100,000 people so the above proportion is generally multiplied by 100,000 (in particular, for rare outcomes).

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#### Proportion as a Measure of Morbidity: Incidence

**Disease Incidence** = the number of "new cases" developed in a defined period of time divided by the number of individuals at risk for the disease at the beginning of the time period.

Disease Incidence =

Number of "new cases" in a period of time

Number of individuals at risk for the disease (at the beginning of the time period).

Incidence is aimed to investigate possible time trends. Multiply incidence by 100,000 for rare outcomes.

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#### Proportion as a Measure of Mortality: Death Rates

*Follow-up Death Rate* = the number of deaths in a year divided by the total follow-up time (usually in person-years).

Follow-up Death Rate =

Number of deaths

Total number of person-years Follow-up death rates are used to measure the effectiveness of medical treatment programs.

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#### **Traditionally Used Measures of Frequency**

**Age-specific rates:** a rate used for a specific age group. Numerator and denominator refer to the same age group.

Rates that are not age specific are called **crude rates** (one example of a crude rate).

**Case fatality rate:** proportion of cases of a specified cause which die in a specified period of time. (Usually expressed as a percentage).

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#### **Traditionally Used Measures of Frequency**

**Cause specific mortality rate:** number of deaths by a specific cause in a defined population in a defined period of time.

**Infant mortality rate (IMR):** number of deaths in children under one year of age divided by the number of live births in the same period, in a specified population.

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#### **Traditionally Used Measures of Frequency**

**Maternal mortality rate:** The number of maternal deaths by the number of women in the reproductive age (15-49 y.o.) in a given year.

**Perinatal mortality rate:** Number of fetal deaths (28 weeks gestation or more) and early neonatal deaths in a year, in a defined population, divided by the total number of live births and fetal deaths in the same population and time period.

**Survival rate:** The proportion of survivors in a group, usually patients with a disease, who survived in a specified period of time.

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#### **Cumulative Incidence versus Prevalence**

- Temporality
- Prediction
  - CI implies probability that similar individuals will develop condition in future
  - prevalence describes current situation among group of individuals
- Probability of having a disease
  - prevalence guides clinical decision making

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interpretation of diagnostic tests



	Measures of Disease Frequency					
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## Proportions as Parameters for Screening Tests

- Through screening tests people are classified as healthy or as falling into one or more disease categories.
- Examples: HIV test, Colon cancer, Skin tests for TB
- These tests are not 100% accurate and therefore misclassification is unavoidable.
- There are 2 proportions that are used to evaluate these types of diagnostic procedures.

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## Sensitivity and Specificity

- Sensitivity and specificity are terms used to describe the effectiveness of screening tests. They describe how good a test is in two ways - finding false positives and finding false negatives
- *Sensitivity* is the Proportion of diseased who screen positive for the disease
- *Specificity* is the Proportion of healthy who screen healthy

## Sensitivity and Specificity: An Example

A cytologic test was undertaken to screen women for cervical cancer.

	Test Positive	Test Negative	Total
Actually Positive	154	225	379
Actually Negative	362	23,362	23,724

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- Sensitivity =?
- Specificity = ?

### **Example Continued**

- Sensitivity = proportion of people who have the disease who test positive
  - Number of people who have the disease = a?
  - Number of people who have the disease who test positive = b ?
  - Sensitivity=b/a = 154/379 = 40.6%

### **Example Continued**

- Specificity = proportion of people who do not have the disease who test negative
  - Number of people without the disease = a?
  - Number of people without the disease who test negative = b?
  - Specificity=b/a = 23,362/23,724 = 98.5%

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## **Example Continued**

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- This cytologic test is highly specific (specificity = 98.5%), but not very sensitive (sensitivity = 40.6%).
- If a healthy person is tested, the result will almost always be negative.
- If a woman with cancer is tested, there is a 59.4% (1- sensitivity) chance that the disease will not be detected!

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#### False Negative and False Positive

• False Negative: the conditional probability that a person had a negative test result given they have the disease.

False negative = 1 -sensitivity.

• False Positive: the conditional probability that a person has a positive test result given they do not have the disease.

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False positive = 1 -specificity.

