## Observational Epidemiology

## Small pox epidemic

Developed disease

## Did not developed

 disease

Milkmaid women
had contact Cow pox

Hypothesis- Cow pox offers immunity to Small Pox.

- Is hypothesis true?
- Presence of co- factor Factorls other than cow pox infection present......
- Host factors--- only women are protected?
- What percentage of protection does the cow pox offer?
- Whether or not a statistical association exists between a disease and suspected factor
- If it does exist, the Strength of Association
- Descriptive study- population
- Analytical- Individual



## ANALYTICAL EPIDEMIOLOGY

## Descriptive studies

Hypothesis-
Prevalence of lung CA is higher among smokers

- Hypothesis is a supposition arrived at from observation or reflection.
- It can be accepted or rejected using the technique of analytical epidemiology.


## Cigarette smoking causes lung cancer"

- How many times smokers are at risk of getting lung cancer in comparision with non-smokers?
- Prevalence Rate of lung cancer- what percentage of lung cancer can be attributable to smoking.
- If we prevent smoking from the community- what percentage of decrease in prevalence of lung cancer we can predict?
- The smoking of $30-40$ cigarettes per day causes lung cancer in 10 percent of smokers after 20 years of exposure



## Case- Control Study

Smokers
/

Do not Develop

Develop Cancer

Cohort Study

# Descriptive v/s Analytical studies 

- Instead of Entire Population Individual subjects are studied.
- However inference is applicable to the population from which subjects are selected.


## Case-Control Studies

Alternative names

- Retrospective studies
- Comparison studies
- Case reference studies

Common first approach to test the Hypothesis

## Characteristics

- Both exposure and outcome have occurred before the start of the study
- The study proceeds backwards from effect to cause
- It uses a control or comparison group to accept or reject a hypothesis


## Case



## Smokers

## Control



## Smokers Non

Smokers

## Design of case control study

## Case



## Design of case control study



## Design of case control study



## Direction of inquiry

## Design of case control study



## Direction of inquiry

Time

## Framework

| Case <br> control | Disease <br> yes | Disease <br> no | Total |
| :--- | :--- | :--- | :--- |
| Exposure | $\mathbf{a}$ | $\mathbf{b}$ | $\mathrm{a}+\mathrm{b}$ |
| No <br> exposure | $\mathbf{C}$ | $\mathbf{d}$ | $\mathrm{c}+\mathrm{d}$ |
| Total | $\mathrm{a}+\mathrm{c}$ | $\mathrm{b}+\mathrm{d}$ | $\mathrm{a}+\mathrm{b}+\mathrm{c}+\mathrm{d}$ |

## Process

- People with disease (cases) are selected from amongst the population
- People without the disease (controls) are selected from comparable population and they must be free from disease
- Data are collected
- Occurrence of possible cause is compared between cases and controls
- Analysis of data to find out association between the suspected cause and the effect


## Applications

- Suitable for rare diseases
- Suitable for chronic diseases
- Common first approach to test the hypothesis


## Basic steps

- Selection of cases and controls
- Matching
- Measurement of exposure
- Analysis


## Selection of cases and controls

- Case selection - easy
- Define case
- Define diagnostic criteria
- Define eligibility criteria ( Incident case/ Prevalent case)
- Source of cases (Hospital / Population )


## Selection of cases and controls

e.g.- DM

- Define case- at what bd. sugar level\ urine sugar
- Define diagnostic criteria- RBs\fasting\GTT
- Define eligibility criteria ( Incident case/ Prevalent case)
- Source of cases (Hospital / Population )


# Selection of cases and controls Control selection- Difficult 

-Hospital controls
-Relatives
-Neighborhood (Residence/ Workplace)
-General population
-Control from more than one source

## How many controls

- At least 1: 1
- If few cases- more than one control study
- "Choice of controls and cases must not be influenced by exposure status"


## Basic steps

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- Measurement of exposure
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## Asbestos- $\rightarrow$ CA lung

## Case

## Control group

- Mean age- 30 years
- 50\% males
- $10 \%$-smokers


## Matching

Process by which we select controls in such a way that they are similar to cases with regard to variables which are known to influence the outcome of disease and subsequently confound the results

## Confounding factor

- One which is associated both with exposure and disease and is distributed unequally in study and control group
- One that, although associated with exposure under investigation, is itself independently a risk factor for disease
- E.g.- AgelSex\ Marital status.....


## Matching procedure

- Group matching
- Pairs


## Basic steps

- Selection of cases and controls
- Matching
- Measurement of exposure
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## Control


-\% of smokers in case vs. control gp

- Average duration of smoking
-Average- no. of cigarette/ day.....


## Measurement of exposure

- Occurred or not occurred
- Duration, severity and frequency
- Measured in same manner in both cases and controls
- Rule out any possible bias


## Basic steps

- Selection of cases and controls
- Matching
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## Analysis

- Exposure rate
- Estimation of exposure risk
- To ascertain the association- Test of significance SE of proportions and means
- Association does not imply causation


## Framework

| Case <br> control | Disease <br> yes | Disease <br> no | Total |
| :--- | :---: | :---: | :--- |
| Exposure | $a$ | $b$ | $a+b$ |
| No <br> exposure | $\mathbf{C}$ | $d$ | $c+d$ |
| Total | a+c <br> (Diseased) | $\mathrm{b}+\mathrm{d}$ <br> (Healthy) | $\mathrm{a}+\mathrm{b}+\mathrm{c}+\mathrm{d}$ |

## Framework

| Case control | Disease <br> yes | Disease <br> no | Total |
| :--- | :---: | :---: | :--- |
| Exposure | a | b | $\mathrm{a}+\mathrm{b}$ |
| No exposure | c | d | $\mathrm{c}+\mathrm{d}$ |
| Total | $\mathrm{a}+\mathrm{c}$ <br> (Diseased) | $\mathrm{b}+\mathrm{d}$ <br> (Healthy) | $\mathrm{a}+\mathrm{b}+\mathrm{c}+\mathrm{d}$ |

Exposure rate amongst cases $=\mathrm{a} / \mathrm{a}+\mathrm{c}$
Exposure rate amongst controls $=\mathrm{b} / \mathrm{b}+\mathrm{d}$

# Odds ratio (Risk Ratio/ Cross Product ratio) 

- Measure of strength of association and closely related to RR
- OR = ad / bc (Key parameter in case control study)


## Framework

| Case control | Disease <br> yes | Disease <br> no | Total |
| :--- | :---: | :---: | :--- |
| Exposure | a | b | $\mathrm{a}+\mathrm{b}$ |
| No exposure | c | d | $\mathrm{c}+\mathrm{d}$ |
| Total | $\mathrm{a}+\mathrm{c}$ <br> (Diseased) | $\mathrm{b}+\mathrm{d}$ <br> (Healthy) | $\mathrm{a}+\mathrm{b}+\mathrm{c}+\mathrm{d}$ |

$O R=a d / b c$
(Key parameter in case control study)

- Out of 300 study subjects-
- Cases-100, -- 50 are smokers
- Controls- 200 --- 10 are smokers


## Framework

| Case <br> control | Disease <br> yes | Disease <br> no | Total |
| :--- | :--- | :--- | :--- |
| Smoker | 50 | 10 | 60 |
| Non <br> smoker | 50 | 90 | 140 |
| Total | 100 | 200 | 300 |

## Framework

| Case control | Disease <br> yes | Disease <br> no | Total |
| :--- | :--- | :--- | :--- |
| Smoker | 50 | 10 | 60 |
| Non smoker | 50 | 90 | 140 |
| Total | 100 | 200 | 300 |

$\mathrm{OR}=\mathrm{ad} / \mathrm{bc}$

$$
=\frac{50 \times 90}{50 \times 10}
$$

- $O R=a d / b c$
$=\underline{50 \times 90}$
$50 \times 10$
$=9$
- Smokers are at 9 times higher risk of getting lung cancer than non smoker.
-Disease being investigated is relatively rare
-Case must be representative of those who have the disease
-Controls must be representative of those who do not have the disease


## Types of bias

- Bias due to confounding
- Memory or recall bias
- Selection bias
- Berksonian bias
- Interviewer's bias


## Advantages

- Easy, inexpensive, relatively quick
- Fewer subjects
- Suitable for rare diseases
- No risk for subjects
- Simultaneous study of multiple etiological factors
- Risk factor can be identified
- No attrition problem
- Ethical problems are minimum
- Easy analysis


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## Disadvantages

- Problem of bias specially recall bias
- Selection of controls is difficult
- Incidence can not be calculated
- Only estimate of RR
- Only disease under study is studied
- Only association is measured
- Not suitable for evaluation of therapy or prophylaxis


## THANK YOU

