

Case-Control Studies

Objectives

- How to design a case-control study?
- When to use case-control study?
- How to minimize bias in case-control study?
- How to establish association between exposure and outcome?

Case-Control

- It is observational analytic epidemiologic investigation
- Subjects are grouped based on their outcome
- The groups are then compared with respect to the proportion having history of an exposure or characteristic of interest

Case control studies

- Are good for long latency period (efficient in time and costs)
- Are good for rare diseases
- allow for evaluation of a wide range of potential etiologic exposures

- Major problem with this design:
 - Both exposure and disease have already occurred at the time the participants enter the study

Aspects to be considered

- You have to:
 - Guarantee comparability between cases and controls
 - Do your best to obtain accurate and complete data
 - Have different sources of information (about exposure and disease) and insure the validity of these sources
 - Clear definition of the outcome (homogenous disease entity) e.g. congenital malformation
 - Definite
 - Probable
 - possible

Selection of cases

- Hospital-based case-control study
- Population-based case-control study

Validity should not be compromised in an attempt to achieve generalizability

Selection of controls

- The controls must be selected to represent **not** the entire nondiseased population
- But they should represent the population of individuals who would have been identified and included as cases had they developed the outcome
 - Sources of controls could be:
 - Hospital controls
 - General population controls
 - Special controls such as:
 - Friends
 - Neighbors
 - Relatives
- How many controls per case should be used (1:1 to 4(controls):1(case))

Advantages of using hospitalized controls

- Easily identified
- Readily available in sufficient numbers
- Reduce recall bias (how?)
- Minimizing bias due to nonresponse (how?)

What is (are) the disadvantage(s) of having hospitalized controls

Ascertainment of Disease and Exposure Status

- Any potential source of information must be carefully considered (accurate and comparable for all study groups)
- Possible sources of information:
 - Death certificates
 - Case registries
 - Office records of physicians
 - Hospital admission or discharge records
 - Log books
 - HIS

Exposure

- From study subjects
- Surrogate
- Medical records
- HIS
 - IT IS ADVISED TO HAVE THE INTERVIEWER UNAWARE OF THE HYPOTHESIS, WHY?
 - IT IS RECOMMENDED TO OBTAIN EXPOSURE INFORMATION FROM RECORDS COMPLETED BEFORE THE OCCURRENCE OF THE OUTCOME, WHY?

The role of bias

- Selection bias:

Can occur whenever the inclusion of cases or controls into the study depends in some way on the exposure of interest

- Observation bias:

This type of bias may arise because of the prior knowledge of the disease status by the participant or by the investigator

- Recall bias:

Relates to differences in the ways exposure information is remembered by cases and by controls.

- Misclassification:

Refers to error in the categorization of either exposure or disease status
Differential vs NonDifferential

Strength and limitations of case-control study

- Strengths:

- Relatively quick and inexpensive
- Well-suited to the evaluation of diseases with long latent periods.
- Optimal for the evaluation of rare diseases
- Can examine multiple etiologic factors for a single disease

Strength and limitations of case-control study

- Limitations:
 - Insufficient for the evaluation of rare exposures
 - In some situations, the temporal relationship between exposure and disease may be difficult to establish
 - It is particularly prone to bias (selection and recall)^{exposure and disease}

Thank you