## Slides 10: Comparing two populations

- Comparing two independent samples
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- Paired samples
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- Paired samples: $H_{0}: \mu_{1}=\mu_{2}$
- Paired binary samples: $H_{0}: p_{1}=p_{2}$
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Comparing two independent samples
Suppose we wish to compare two population distributions with means and standard deviations ( $\mu_{1}, \sigma_{1}$ ) and ( $\mu_{2}, \sigma_{2}$ ) based on two random samples $\left(x_{1}, \ldots, x_{n}\right)$ and $\left(y_{1}, \ldots, y_{m}\right)$ from these two populations.

The difference $\mu_{1}-\mu_{2}$ is estimated by $\bar{x}-\bar{y}$, where

$$
\begin{array}{lll}
\bar{x}=\frac{x_{1}+\ldots+x_{n}}{n}, & s_{\bar{x}}=\frac{s_{1}}{\sqrt{n}}, & s_{1}^{2}=\frac{1}{n-1} \sum_{i=1}^{n}\left(x_{i}-\bar{x}\right)^{2}, \\
\bar{y}=\frac{y_{1}+\ldots+y_{m}}{m}, & s_{\bar{y}}=\frac{s_{2}}{\sqrt{m}}, & s_{2}^{2}=\frac{1}{m-1} \sum_{i=1}^{m}\left(y_{i}-\bar{y}\right)^{2} .
\end{array}
$$

If $\left(X_{1}, \ldots, X_{n}\right)$ is independent from $\left(Y_{1}, \ldots, Y_{m}\right)$, then

$$
\operatorname{Var}(\bar{X}-\bar{Y})=\operatorname{Var}(\bar{X})+\operatorname{Var}(\bar{Y})=\frac{\sigma_{1}^{2}}{n}+\frac{\sigma_{2}^{2}}{m}
$$

and we may compute the standard error of $\bar{x}-\bar{y}$ as

$$
s_{\bar{x}-\bar{y}}=\sqrt{\frac{s_{1}^{2}}{n}+\frac{s_{2}^{2}}{m}}
$$

Question. Is $\bar{x}-\bar{y}$ an unbiased estimate of $\mu_{1}-\mu_{2}$ ?

Large sample test for two means
If $n$ and $m$ are large, we can use a normal approximation

$$
\frac{(\bar{X}-\bar{Y})-\left(\mu_{1}-\mu_{2}\right)}{S_{\bar{X}}-\bar{Y}} \approx \mathrm{~N}(0,1)
$$

Under the hypothesis of no difference $H_{0}: \mu_{1}=\mu_{2}$ the distribution of the test statistic $z=\frac{\bar{x}-\bar{y}}{s_{\bar{x}-\bar{y}}}$ is approximated by the standard normal.

Approximate confidence interval $I_{\mu_{1}-\mu_{2}} \approx \bar{x}-\bar{y} \pm z_{\alpha / 2} \cdot s_{\bar{x}-\bar{y}}$
Example: iron retention
Percentage of $\mathrm{Fe}^{2+}$ and $\mathrm{Fe}^{3+}$ retained by mice data at concentration 1.2 millimolar. From the summary of the data:

$$
\begin{aligned}
& \mathrm{Fe}^{2+}: n=18, \bar{x}=9.63, s_{1}=6.69, s_{\bar{x}}=1.58 \\
& \mathrm{Fe}^{3+}: m=18, \bar{y}=8.20, s_{2}=5.45, s_{\bar{y}}=1.28
\end{aligned}
$$

we obtain


$$
\bar{x}-\bar{y}=1.43, \quad s_{\bar{x}-\bar{y}}=\sqrt{s_{\bar{x}}^{2}+s_{\bar{y}}^{2}}=2.03
$$

According to the large sample test we cannot reject $H_{0}: \mu_{1}=\mu_{2}$.

Large sample test for two proportions
For the binomial model $X \sim \operatorname{Bin}\left(n, p_{1}\right), Y \sim \operatorname{Bin}\left(m, p_{2}\right)$, two independently generated values $(x, y)$ give sample proportions

$$
\hat{p}_{1}=\frac{x}{n}, \quad \hat{p}_{2}=\frac{y}{m},
$$

which are unbiased estimates of $p_{1}, p_{2}$ and have standard errors

$$
s_{\hat{p}_{1}}=\sqrt{\frac{\hat{p}_{1}\left(1-\hat{p}_{1}\right)}{n-1}}, \quad s_{\hat{p}_{2}}=\sqrt{\frac{\hat{p}_{2}\left(1-\hat{p}_{2}\right)}{m-1}} .
$$

If the samples sizes $m$ and $n$ are large, then an approximate $95 \%$ confidence interval for the difference $p_{1}-p_{2}$ is given by

$$
I_{p_{1}-p_{2}} \approx \hat{p}_{1}-\hat{p}_{2} \pm 1.96 \sqrt{\frac{\hat{p}_{1}\left(1-\hat{p}_{1}\right)}{n-1}+\frac{\hat{p}_{2}\left(1-\hat{p}_{2}\right)}{m-1}} .
$$

With help of this formula we can test the null hypothesis of equality

$$
H_{0}: p_{1}=p_{2} .
$$

Question. Would you reject $H_{0}: p_{1}=p_{2}$ given a $95 \%$ confidence interval $I_{p_{1}-p_{2}} \approx(-0.3,-0.1)$, at what significance level?

Consider two consecutive monthly poll results $\hat{p}_{1}$ and $\hat{p}_{2}$ with $n \approx m \approx 5000$ interviews. A change in support to a major political party from $\hat{p}_{1}$ to $\hat{p}_{2}$ (with both numbers being close to $40 \%$ ) is deemed significant at $5 \%$ level, if

$$
\left|\hat{p}_{1}-\hat{p}_{2}\right|>1.96 \cdot \sqrt{2 \cdot \frac{0.4 \cdot 0.6}{5000}} \approx 1.9 \%
$$

This should be compared to comparing one poll result with the previous election result $p_{0}=0.4$. Here we apply the one-sample hypothesis for testing $H_{0}: p=0.4$ vs $H_{0}: p \neq 0.4$. In view of

$$
I_{p} \approx \hat{p} \pm 1.96 \cdot \sqrt{\frac{\hat{p}(1-\hat{p})}{n-1}}
$$

with $\hat{p} \approx 0.4$, we conclude that the difference from the election result is significant if

$$
|\hat{p}-0.4|>1.96 \cdot \sqrt{\frac{0.4 \cdot 0.6}{5000}} \approx 1.3 \%
$$

Question. Where the difference between two margines of error $1.9 \%$ vs $1.3 \%$ come from?

Examples of paired observations
different drugs for two patients matched by age, sex,
a fruit weighed before and after shipment, two types of tires tested on the same car.

Two paired samples can be viewed as one 2D random sample

$$
\left(x_{1}, y_{1}\right), \ldots,\left(x_{n}, y_{n}\right)
$$

Two estimate $\mu_{1}-\mu_{2}$, turn to a 1D sample of differences

$$
\left(d_{1}, \ldots, d_{n}\right), \quad d_{i}=x_{i}-y_{i} .
$$

Its sample mean is $\bar{d}=\bar{x}-\bar{y}$. It is an unbiased estimate of $\mu_{1}-\mu_{2}$ whose standard error is computed based on

$$
\operatorname{Var}(\bar{X}-\bar{Y})=\frac{1}{n}\left(\sigma_{1}^{2}+\sigma_{2}^{2}-2 \sigma_{1} \sigma_{2} \rho\right)
$$

taking into account the correlation coefficient $\rho=\frac{\operatorname{Cov}(X, Y)}{\sigma_{1} \sigma_{2}}$.
Question. Why pairing should ensure $\rho>0$ ?

Paired samples: large sample test of no difference
To study the effect of cigarette smoking on platelet aggregation, Levine (1973) drew blood samples from $n=11$ individuals before and after they smoked a cigarette and counted the platelets.

Sample correlation coefficient
$r=\frac{\left(x_{1}-\bar{x}\right)\left(y_{1}-\bar{y}\right)+\ldots+\left(x_{n}-\bar{x}\right)\left(y_{n}-\bar{y}\right)}{(n-1) s_{1} s_{2}}=0.90$
We test $H_{0}: \mu_{1}=\mu_{2}$ by applying the large sample test for the mean to
$H_{0}: \mu=0$ against $H_{1}: \mu \neq 0$
where $\mu=\mu_{1}-\mu_{2}$. The test statistic

| Before $y_{i}$ | After $x_{i}$ | $d_{i}=x_{i}-y_{i}$ |
| :---: | :---: | :---: |
| 25 | 27 | 2 |
| 25 | 29 | 4 |
| 27 | 37 | 10 |
| 28 | 43 | 15 |
| 30 | 46 | 16 |
| 44 | 56 | 12 |
| 52 | 61 | 9 |
| 53 | 57 | 4 |
| 53 | 80 | 27 |
| 60 | 59 | -1 |
| 67 | 82 | 15 |

$z_{\text {obs }}=\frac{\bar{d}}{s_{\bar{d}}}=\frac{10.27}{2.40}=4.28$
gives a very small two-sided p-value, $2 \cdot(1-\Phi(4.28)=0.00002$, showing that smoking has a significant health effect.

Question. Where the value $r=0.9$ was used?

Suppose we have two dependent random variables

$$
X \sim \operatorname{Bin}\left(1, p_{1}\right), \quad Y \sim \operatorname{Bin}\left(1, p_{2}\right)
$$

Vector $(X, Y)$ takes one of the four possible values $(0,0),(0,1),(1,0)$, $(1,1)$ with probabilities $\pi_{00}, \pi_{01}, \pi_{10}, \pi_{11}$. Observe that

$$
p_{1}=\pi_{10}+\pi_{11}, \quad p_{2}=\pi_{01}+\pi_{11}, \quad p_{1}-p_{2}=\pi_{10}-\pi_{01}
$$

With $n$ independent paired observations, we count ( $w_{00}, w_{01}, w_{10}, w_{11}$ ) the numbers of different outcomes.

An unbiased point estimate of $p_{1}-p_{2}$ is given by

$$
\hat{p}_{1}-\hat{p}_{2}=\hat{\pi}_{10}-\hat{\pi}_{01}, \quad \hat{\pi}_{10}=\frac{w_{10}}{n}, \quad \hat{\pi}_{01}=\frac{w_{01}}{n} .
$$

Example. In terms of opinion polls, paired sampling corresponds to asking the same $n$ individuals in January and then in February about their opinion towards a certain political party. The important counts are $w_{01}$ and $w_{10}$ of how many people have changed their preferences.

Paired binary samples: confidence intervall for $p_{1}-p_{2}$
Using the multinomial $\operatorname{Mn}\left(n, \pi_{00}, \pi_{01}, \pi_{10}, \pi_{11}\right)$ distribution, we find

$$
\operatorname{Var}\left(W_{10}-W_{01}\right)=n\left(\pi_{10}+\pi_{01}-\left(\pi_{10}-\pi_{01}\right)^{2}\right)
$$

This yields a formula for the standard error

$$
s_{\hat{p}_{1}-\hat{p}_{2}}=\sqrt{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}-\left(\hat{\pi}_{10}-\hat{\pi}_{01}\right)^{2}}{n-1}} .
$$

Again, referring to the CLT we arrive at a $95 \%$ confidence interval

$$
I_{p_{1}-p_{2}} \approx \hat{\pi}_{10}-\hat{\pi}_{01} \pm 1.96 \sqrt{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}-\left(\hat{\pi}_{10}-\hat{\pi}_{01}\right)^{2}}{n-1}}
$$

Example. Suppose the same $n=2000$ individuals were asked first in January and then in February about their opinion towards a certain political party. The counts were $w_{01}=100$ and $w_{10}=60$. In this case, $\hat{\pi}_{01}=0.05$ and $\pi_{10}=0.03$, so that

$$
I_{p_{2}-p_{1}} \approx \hat{\pi}_{01}-\hat{\pi}_{10} \pm 1.96 \sqrt{\frac{\hat{\mu}_{10}+\hat{\pi}_{01}-\left(\hat{\mu}_{10}-\hat{\pi}_{01}\right)^{2}}{n-1}}=0.03 \pm 0.012
$$

Significant difference at $5 \%$ level.

The hypothesis of no difference $H_{0}: p_{1}=p_{2}$ is equivalent to
$H_{0}: \pi_{10}=\pi_{01}$. According to

$$
I_{p_{1}-p_{2}} \approx \hat{\pi}_{10}-\hat{\pi}_{01} \pm 1.96 \sqrt{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}-\left(\hat{( }_{10}-\hat{\pi}_{01}\right)^{2}}{n-1}}
$$

the rejection region for against $H_{0}: \pi_{10} \neq \pi_{01}$ has the form

$$
\mathcal{R}=\left\{\frac{\left|\hat{\pi}_{10}-\hat{\pi}_{01}\right|}{\sqrt{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}-\left(\hat{\pi}_{10}-\hat{\pi}_{01}\right)^{2}}{n-1}}}>1.96\right\}
$$

Now notice that the squared left hand side equals

$$
\frac{\left(\hat{\pi}_{10}-\hat{\pi}_{01}\right)^{2}}{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}-\left(\hat{\pi}_{10}-\hat{\pi}_{01}\right)^{2}}{n-1}} \approx \frac{1}{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}}{n\left(\hat{\pi}_{10}-\hat{\pi}_{01}\right)^{2}}-\frac{1}{n}} \approx \frac{1}{\frac{\hat{\pi}_{10}+\hat{\pi}_{01}}{n\left(w_{10}-\hat{\pi}_{01}\right)^{2}}}=\frac{\left.w_{01}\right)^{2}}{w_{10}+w_{01}} .
$$

This leads to the McNemar test statistic

$$
\mathrm{X}^{2}=\frac{\left(w_{10}-w_{01}\right)^{2}}{w_{10}+w_{01}}
$$

whose null distribution is approximately $\chi_{1}^{2}$-distribution.

## Controlled experiments

Double-blind, randomised controlled experiments are used to
balance out such external factors as

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placebo effect,
time factor,
background variables like temperature,
location factor.
```

Example. Portocaval shunt is an operation used to lower blood pressure in the liver. People believed in its high efficiency until the controlled experiments were performed.

| Enthusiasm level | Marked | Moderate | None |
| :--- | :---: | :---: | :---: |
| No controls | 24 | 7 | 1 |
| Nonrandomized controls | 10 | 3 | 2 |
| Randomized controls | 0 | 1 | 3 |

Simpson's paradox
Hospital A has higher overall death rate than hospital B.
However, if we split the data in two parts, patients in good $(+)$ and bad (-) conditions, for both parts hospital A performs better.

| Hospital: | A | B | $\mathrm{A}+$ | $\mathrm{B}+$ | $\mathrm{A}-$ | $\mathrm{B}-$ |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: |
| Died | 63 | 16 | 6 | 8 | 57 | 8 |
| Survived | 2037 | 784 | 594 | 592 | 1443 | 192 |
| Total | 2100 | 800 | 600 | 600 | 1500 | 200 |
| Death Rate | .030 | .020 | .010 | .013 | .038 | .040 |

Here, the external factor, patient condition, is an example of a confounding factor:

Hospital performance $\leftarrow$ Patient condition $\rightarrow$ Death rate
Always remember that correlation does not imply causation.

