







theory the frequencies should be in proportions 9:3:3:1.

Are the data in agreement with Mendel's theory?



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Goodness-of-fit test

Test of the hypothesis that the probabilities of the categories agree with specified values.

Test statistic:

$$X^{2} = \sum_{i} \frac{(\text{observed}_{i} - \text{expected}_{i})^{2}}{\text{expected}_{i}}$$

where the sum is over all categories.

- Small values of X^2 : data and hypothesis agree well.
- Large values of X^2 : data are in disagreement with the hypothesis.

It may be shown that if the hypothesis is true then X^2 follows (approximately) a so-called $\chi^2(r)$ -distribution with r degrees of freedom, where

r =degrees of freedom = number of categories - 1



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Mendel's hypothesis

$$H_0: p_{ry} = \frac{9}{16}, \quad p_{rg} = \frac{3}{16}, \quad p_{wy} = \frac{3}{16}, \quad p_{wg} = \frac{1}{16}.$$

 $X^2 = \ldots = 0.470$

This value is to be compared with a $\chi^2(3)$ -distribution. The

P-value is

$$1 - P(\chi_3^2 \ge 0.470) = 1 - 0.075 = 0.925,$$

which is far from significant. The data are in excellent agreement with Mendel's hypothesis.

Rule of thumb

To calculate the P-value the χ^2 -distribution may be used if the expected number of observations for each category is at least 5.

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The
$$\chi^2(r)$$
-distribution



$_{\rm JNIVERSITY}$ of copenhagen faculty of life sciences 2×2 count tables

With two binary classification criteria the results may be summarized in a 2 by 2 table:

	Response 1	Response 2	Total
Pop 1	<i>y</i> ₁₁	<i>y</i> ₁₂	$n_1 = y_{11} + y_{12}$
Pop 2	<i>y</i> ₂₁	<i>y</i> ₂₂	$n_2 = y_{21} + y_{22}$
Total	$c_1 = y_{11} + y_{21}$	$c_2 = y_{12} + y_{22}$	$n = n_1 + n_2 = c_1 + c_2$

Estimates

$$p_1 = \frac{y_{11}}{n_1}, \qquad \hat{p}_{21} = \frac{y_{21}}{n_2}.$$

Question: Is $p_{11} = p_{21}$?

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 \hat{p}_1

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Example: Avadex and mice

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Example: Avadex and mice (cont.)

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Test in 2×2 count tables

		Tumor	No tumor	Total	Europeter Leonante
	Avadex	4	12	16	
	Control	5	74	79	The expected counts in cell (i,j) in a two-way count table is
	Total	9	86	95	
Does the pes	ticide Avac	lex increa	se the occur	rence of	mors? Expected count = $\frac{(Row total) \cdot (Column total)}{Grand total}$
Hypothesis <i>p</i>	$t_t = p_c$. Est	imates ar	e		Test statistic: as with goodness-of-fit test we use the "chi-squared"statistic
	$\hat{p}_t = \frac{4}{16}$	= 0.25	$\hat{p}_c = \frac{5}{79} = 0$.0633.	$\mathbf{v}^2 - \mathbf{\nabla} (\text{observed}_i - \text{expected}_i)^2$
and a 95% co	onfidence i	nterval fo	$p_t - p_c$ is		$-\sum_{i} - \sum_{j} - \sum_{i} - \sum_{i} - \sum_{j} - \sum_{$
		(-0.032)	1,0.4056)		and the test statistic follows a $\chi^2(1)$ -distribution! [Notice that df $=1$]
which contair	ıs 0.				
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Row 1

Row 2

Row r

Tota

Test statistic

 $r \times k$ count tables

*Y*₁₁

*Y*21

Yr1

*C*₁

Total

 n_1

 n_2

n_r

n

Column k

 y_{1k}

Y2k

÷

Yrk

Cı

. . .

. . .

. . .

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Example: 1111 cats

1111 cats received at an animal shelter





The method applies to an $r \times k$ -table as well as a 2 \times 2 table.

*Y*₁₂

Y22

yr2

 C_2

Column 1 Column 2 ···



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One-sided hypotheses

For 2×2 -tables the alternative hypothesis may occasionally be one-sided, that is,

$$H_0: p_{11} = p_{21}$$
 vs. $H_A: p_{11} > p_{21}$

meaning that we regard it as unthinkable that $p_{11} < p_{21},$ whether the hypothesis is true or not.

Test method of for one-sided test:

- Check whether the estimates are in the direction of alternative hypothesis.
- If no then there is no indication of disagreement with the null-hypothesis and the *p*-value is 1.
- **(3)** If yes then the *p*-value is half of the *p*-value, you would get by a two sided test using the χ^2 distribution.

One-sided hypothesis: diabetes

Diabetes

26

12

38

 $H_0: p_t = p_c \mod H_A: p_t > p_c.$

 $X^2 = 8.3192.$

No diabetes

24

38

62

Tota

50

50 100

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Example: Avadex and mice

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	Tumor	No tumor	Total
Avadex	4	12	16
Control	5	74	79
Total	9	86	95

Does the pesticide Avadex increase the occurrence of tumors?

Expected values under the hypothesis $H_0: p_t = p_c$:

	Tumor	No tumor	Total
Avadex	1.52	14.48	16
Control	7.48	71.52	79
Total	9	86	95

What is the conclusion?

Neutered group

Total

Null-hypothesis

Test statistic:

Non-neutered group





Exact test in 2×2 -tables. No approximations needed.

Advanced: The idea is that the row- and column-totals tells nothing about the relation. Hence we consider those as given in advance and compute the conditional probability of the test statistic given the marginal totals.

What to do? Compute the p-value as the sum of probabilities of tables that are equal to or more extreme than the observed.



	OPEN	HAGE	N		F.	ACUL	TY OF	LIFE SC
Other 1	tabl	es v	vith sa	me marg	inal tot	als		
Augdov	+	-	Total		A 1	+	-	Total
Avadex	4	12	10		Avadex	2	14	16
Control	5	74	79		Control	1	72	79
Total	9	86	95		Total	9	86	95
	+	-	Total			+	-	Total
Avadex	3	13	16		Avadex	1	15	16
, waach	-	72	70		Control	8	71	79
Control	6	15	19		00	-	• -	

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The probability of a table

$$Pr(Y = 3) = \frac{\binom{16}{3}\binom{79}{6}}{\binom{99}{9}} = 0.1325$$

$$Pr(Y = 2) = \frac{\binom{16}{2}\binom{79}{7}}{\binom{99}{9}} = 0.2960$$

$$Pr(Y = 1) = \frac{\binom{16}{1}\binom{79}{8}}{\binom{99}{9}} = 0.3553$$

$$Pr(Y = 0) = \frac{\binom{16}{10}\binom{79}{9}}{\binom{99}{9}} = 0.1752$$

$$Pr(Y = 4) = 0.0349 + 0.1325 + 0.2960 + 0.3553 + 0.1752 = 0.994$$

$$Pr(Y \ge 4) = 1 - 0.994 + 0.0349 = 0.041$$

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Comparison of risks

Three common ways to compare probabilities, for example probability of a disease for two age groups:

- Difference in probability: $p_1 p_2$
- Relative risk: p_1/p_2
- Odds ratio: $Odds_1(A)/Odds_2(A) = \frac{p_1}{1-p_1} / \frac{p_2}{1-p_2}$.

The hypotheses that the two probabilities are the same $(p_1 = p_2)$ may be written

probability difference = 0, RR = 1, OR = 1,

in terms of probabilities, relative risk and odds ratio, respectively, but the magnitude of the deviation depends on which of the three representations is used.

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Odds

Odds: a different way to express a probability.

Odds 1:1 for the event A corresponds to P(A) = 1/2.

Odds 3:1 for the event A corresponds to P(A) = 3/4.

Odds 1:7 for the event A corresponds to P(A) = 1/8.

Odds for an event A are defined as

 $Odds(A) = \frac{P(A)}{1 - P(A)}$

Notice that while the probability is between 0 and 1,

 $0 \leq \text{Odds}(A) \leq \infty, \qquad 0 \leq P(A) \leq 1$

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Comparison of t	wo risks		
Example: Risk of findir • before wash: p ₁ = • after wash: p ₂ = 0	ng pesticide res : 0.04).01	sidues in strawl	perries
Probability difference	= 0.03, RR = 0	0.25, OR = 0.2	42.
Compare the cases			
$(p_1,p_2) =$	(0.04. 0.01)	(0.40, 0.10)	(0.99, 0.96)
Difference, $p_1 - p_2$	0.03	0.30	0.03
RR	4.00	4.00	1.03
OR	4.13	6.00	4.13

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Odds-ratio

Example: Avadex and mice (cont.)

	Tumor	No tumor	Total
Avadex	4	12	16
Control	5	74	79
Total	9	86	95

The estimate of OR is

$$OR = \frac{4 \cdot 74}{5 \cdot 12} = 4.93, \quad In(OR) = 1.596$$

The standard error of the *logarithm* of OR is

$$\mathsf{SE}(\mathsf{ln}(\mathsf{OR})) = \sqrt{\frac{1}{4} + \frac{1}{12} + \frac{1}{5} + \frac{1}{74}} = 0.74$$

The 95%-confidence interval for ln(OR) then becomes $1.60\pm1.96\cdot0.74=(0.15,3.05)$, which is translated back to OR by the exponential function:

$$\exp(0.15) < OR < \exp(3.05).$$

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Case-control and odds-ratio		
Row- and column probabilities are not the sar odds-ratio is the same if rows and columns ar	me $(rac{10}{14} eq rac{10}{16})$, but re switched.	
Intected Not infected		

	Infected	Not infected	10.41
yes	10	4	$OR = \frac{10^{\circ}41}{6}$
no	6	41	6 - 4

Thus, the OR for infection among the "yes" group vs. the "no" group is identical to the OR for "yes" among infected vs. non-infected. $OR\!=\!17.1$ seems to indicate that the product has to do with the risk of infection.

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Odds-ratio is ratio between the odds in two groups:

$$\mathsf{OR} = \frac{\frac{p_1}{1-p_1}}{\frac{p_2}{1-p_2}} = \frac{p_1 \cdot (1-p_2)}{p_2 \cdot (1-p_1)}$$

The value 1 means that the two odds are the same. Hence, a test for H_0 : OR = 1 is a test of equality of odds (and hence probabilities in the two groups!



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Case-control investi	gatio	'n				
		Infected	Not infected	d S	Sum	1
Have eaten the product	yes	Infected	Not infected	d S 4	Sum 14	
Have eaten the product	yes no	Infected 10 6	Not infected	d S 4	Sum 14 47	

After a number of cases of *Salmonella Manhattan* 16 persons with *Salmonella* infection were ask whether they had eaten sliced smoked fillet from a certain slaughterhouse. A control group was asked the same question.

