# TAMS38 – Experimental Design and Biostatistics, 6 hp January 17 2019, 14–18

The collection of the formulas in mathematical statistics prepared by Department of Mathematics LiU and calculator with empty memory are allowed on the exam. Dictionary English-other language are allowed. No extra notes in the formula collection is allowed.

Score limits: 7-9 points gives grade 3, 9.5-12 gives 4 and 12.5-15 gives 5.

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The result will *normally* be published via LADOK within 12 working days.

Clear answers and justifications are required for each problem.

1. The tensile property of a metal wire depends on different treatments. Three different treatments  $A_1 - A_3$  were tested and compared to a non-treated raw wire  $A_4$ . The tensile property for 11 different wires were

	$y_{ij}$							
$A_1$	9.07	8.75						
$A_2$	9.76	9.88						
$A_3$	9.28	9.15	9.21					
$A_4$	8.47	7.38	7.78	8.72				

with  $SS_A = 4.6098$  and  $SS_E = 1.2083$ .

- (a) Given normal distribution, write out the one-factor model and test the hypothesis of equal means for the four wires at level 1%. (1.5p)
- (b) Assume we can NOT use normal distribution, use a non-parametric method to test the hypothesis of equal medians for the four wires at level 1%. (1.5p)
- 2. Burning cigarettes produce appreciable quantities of carbon monoxide. When cigarette smoke is inhaled, carbon monoxide is combined with carboxyhemoglobin. In a study (*Carbon Monoxide and Exercise in Chronic Bronchitis and Emphysema*, Brit. Med. J., Vol. 283, pp. 877-880, 1981) researchers wanted to determine whether an appreciable concentration of carboxyhemoglobin reduces the exercise tolerance of patients suffering from chronic bronchitis and emphysema. Seven such person was randomly selected and asked to walk for 12 minutes breathing one of four gas mixtures: air, oxygen, air plus CO, oxygen plus CO. The walked distances (in meters) are given in the table below.

	Gas mixture					
Subjets	Air	Oxygen	Air+CO	Oxygen+CO	$\bar{y}_{i}$ .	$s_i$ .
1	835	874	750	854	828.25	54.5
2	787	827	755	829	799.50	35.4
3	724	738	698	726	721.50	16.84
4	336	378	210	279	300.75	72.8
5	252	315	168	336	267.75	75.5
6	560	672	558	642	608.00	57.9
7	336	341	260	336	318.25	38.9
$ar{y}_{\cdot j}$	547.1	592.1	485.6	571.7		
$s_{.j}$	240.9	240.8	265	248.8		$\bar{y}_{} = 549.1$

(a) Give an appropriate model and choose one table below (i, ii, iii or iv) to use in an analysis. (0.5p)

i - Two	-way	ANOVA: y	versus	subj,	gas	ii - Two	-way	ANOVA: y	versus	subj,	gas
Source	d.f	Sum Sq.				Source	d.f	Sum Sq.			
subj	7	1471772				subj	6	1471772			
gas	4	44827				gas	3	44827			
Error	18	16316				Error	18	16316			
Total	29	1532915				Total	27	1532915			
iii - T	wo-wa	y ANOVA:	y versu	us sub	j, gas	iv - Two	-way	ANOVA: y	versus	subj,	gas
Source	d.f	Sum Sq.				Source	d.f	Sum Sq.			
subj	6	1471772				subj	7	1471772			
gas	3	44827				gas	4	44827			
Error	20	16316				Error	16	16316			
Totol	07					<b>—</b> · ¬	07	4 5 9 9 9 4 5			
IOUAL	27	1532915				Total	27	1532915			

- (b) Use an appropriate test to test at level 5% the null hypothesis: the distance does not depend on the gas mixture. Give both  $H_0$  and  $H_1$ . (1p)
- (c) Which gas mixture will give the shortest distance? Answer the question with appropriate intervals with simultaneous significance level at most 5%. (1.5p)
- 3. A canning factory uses many different machines to fill their cans. They suspected that there is great variation in how much each machine fills. They select four machines randomly to test this. Result:

	Machine						
	1	2	3	4			
	1.24	1.20	1.19	1.18			
	1.22	1.20	1.20	1.18			
$y_{ij}$	1.22	1.21	1.19	1.19			
	1.23	1.22	1.20	1.18			
	1.23	1.20	1.21	1.20			
$\bar{y}_{i}$ .	1.228	1.206	1.198	1.186			

**Model:**  $y_{ij} = \mu + \tau_i + \varepsilon_{ij}$ , where  $\tau_i \sim N(0, \sigma_{\tau})$  and  $\varepsilon_{ij} \sim N(0, \sigma)$ ,  $i = 1, \ldots, 4$  and  $j = 1, \ldots, 5$ , independent.

Analysis of Variance for y

Source	DF	SS
М	3	0.0047
Error	16	0.0012
Total	19	0.0059

- (a) Test if the machines are homogeneous with an F-test at level 5%. Write out both the null hypothesis  $H_0$  and the alternative  $H_1$ . (1.5p)
- (b) Give an interval for  $\mu$  with confidence level 95%. Conclusion? (1.5p)
- 4. In nitride etching,  $C_2F_6$  is used as the acting gas. It is possible to vary the flow of the gas, the effect on the cathode, the pressure in the etching chamber and the distance between the cathode and the anode. We will use a 2<sup>4</sup> design to analyze the process. The factors are varied at the following levels.

		Factors		
	Distance (A)	Pressure (B)	$C_2 F_6$ flow (C)	Effect $(D)$
Level	$(\mathrm{cm})$	(mTorr)	(SCCM)	(w)
_	0.80	450	125	275
+	1.20	550	200	325

Many different response variables would be interesting to analyze, but in our experiment we measure the etching rate of silicon nitride. High speed is good and we get

(1)	550	d	1037
a	669	ad	749
b	604	bd	1052
$^{\mathrm{ab}}$	650	abd	868
с	633	cd	1075
ac	642	acd	860
$\mathbf{b}\mathbf{c}$	601	bcd	1063
abc	635	abcd	729

with the following Minitab analysis where the  $2^4$  design matrix is in c1-c16 and the response variable in c17.

MTB > copy c1-c16 m1 MTB > trans m1 m2 MTB > copy c17 m3 MTB > mult m2 m3 m4 MTB > copy m4 c18 MTB > set c19 DATA> 1:16 DATA> end MTB > sort c18 c19 c20 c21; SUBC> by c18. MTB > print c20 c21 Data Display



(a) What effects appear to be of the greatest importance based on the Minitab analysis above? (1p)

Given the result in a), another Minitab analysis is made.

ANOVA: y versus A, D Analysis of Variance for y Source DF SS MS F Ρ 41311 41311 0.000 1 23.77 А D 1 374850 374850 215.66 0.000 A\*D 1 94403 94403 54.31 0.000 20858 Error 12 1738 531421 Total 15 S = 41.6911R-Sq = 96.08%R-Sq(adj) = 95.09% Means А Ν у 826.88 -1 8 1 8 725.25 D Ν у -1 8 623.00 1 8 929.13

А	D	Ν	У
-1	-1	4	597.0
-1	1	4	1056.8
1	-1	4	649.0
1	1	4	801.5

- (b) Which model has been used in the second Minitab analysis? Is it possible to find the best combination of the factors A and D? Motivate your answer using confidence intervals with simultaneous confidence level 95%. (2p)
- 5. In one study, two different methods would be compared  $A_1$  and  $A_2$  to determine the sulfur content of carbon. In total, 14 equivalent samples were distributed to 7 randomly selected laboratories and each laboratory  $(B_1, ..., B_7)$  analyzed on sample with  $A_1$  and one with  $A_2$ . Result:

	$B_1$	$B_2$	$B_3$	$B_4$	$B_5$	$B_6$	$B_7$
$A_1$	0.106	0.124	0.113	0.108	0.096	0.116	0.150
$A_2$	0.111	0.125	0.115	0.116	0.105	0.119	0.162

Since the laboratories were chosen at random, a random effects model should be used. Method no i and laboratory no j gives the observation  $y_{ij}$  from the random variable

$$Y_{ij} = \mu + \tau_i + \beta_j + \varepsilon_{ij},$$

where the  $\tau_i$ :s are fix,  $\sum \tau_i = 0$ ,  $\beta_j \sim N(0, \sigma_\beta)$ ,  $\varepsilon_{ij} \sim N(0, \sigma)$  with all the random variables  $\beta_j$  and  $\varepsilon_{ij}$  independently distributed.

- (a) Estimate  $\tau_2 \tau_1$  and show that the estimator is unbiased. (1p)
- (b) The usual sum of squares for an additive model are given by

$$SS_A = 0.0001143, \quad SS_B = 0.003856, \quad SS_E = 0.00004971.$$

Estimate  $\sigma_{\beta}^2$  and show that the estimator also is unbiased. (2p) Hint:  $\frac{SS_E}{\sigma^2} \sim \chi^2(6)$ 

# TAMS38 Some extra formulas

# Some nonparametric tests

### Tukey-Duckworth's quick test

If  $4 \le n_1 \le n_2 \le 30$ , and  $n_2 \le \frac{4n_1}{3} + 3$  then we can test the null hypothesis  $H_0$  with the test:

- 1. Find the smallest and largest observation in each sample, respectively.
- 2. For the sample with the largest observation, count how many observation that is larger in that sample than in the other sample.
- 3. For the other sample, count how many observations that is smaller than the smallest observation in the first sample.
- 4. Let C be the sum of the number of observations in 2. and 3. For  $\alpha = 0.05$ , 0.01 or 0.001, reject  $H_0$  if  $C \ge 7$ , 10, or 13, respectively.

#### The Wilcoxon signed rank test

Let  $r_i$  be the rank for the observations  $|y_i| \neq 0, i = 1, ..., n$ . Let  $T_+ = \sum_{\{y_i > 0\}} r_i$  and  $T_- = \sum_{\{y_i < 0\}} r_i$ . When  $H_0$  is true and n > 15 us that  $T_+$  and  $T_- \approx N\left(\frac{n(n+1)}{4}, \sqrt{\frac{n(n+1)(2n+1)}{24}}\right)$ . For  $n \leq 15$  us table for Wilcoxons signed rank distribution.

For confidence interval us the N = n(n+1)/2 ordered pairwise means  $A_i$  and  $P(A_{(k)} < \mu < A_{(N-k+1)}) = 1 - 2P(W_S \le k - 1)$ , where  $W_S$  is Wilcoxons signed rank distributed.

## The Wilcoxon-Mann-Whitney test

Let  $d_{ij}$  be the differences  $d_{ij} = x_i - y_j$ ,  $i = 1, ..., n_1$ ,  $j = 1, ..., n_2$  and  $d_{(k)}$ ,  $k = 1, ..., n_1 n_2$ , the ordered differences. The confidence interval for the difference in mean/median is given by

$$I = (d_{(c+1)}, d_{(n_1 n_2 - c)}),$$

where  $c = T_l - \frac{n_1(n_1+1)}{2}$  and  $T_l$  is from the Wilcoxon table for the rank sum test.

#### The Kruskal-Wallis test

Assume a treatments. Let  $r_{ij}$  be the rank for the observation  $y_{ij}$ . Test statistic

$$T = \begin{cases} \frac{12S_a}{N(N+1)} - 3(N+1), & \text{if no ties,} \\ \\ \frac{(N-1)(S_a - C)}{S_r - C}, & \text{if ties,} \end{cases}$$

where  $s_i = \sum_{j=1}^{n_i} r_{ij}$ ,  $S_a = \sum_{i=1}^{a} \frac{s_i^2}{n_i}$ ,  $S_r = \sum_{i=1}^{a} \sum_{j=1}^{n_i} r_{ij}^2$ ,  $C = \frac{1}{4}N(N+1)^2$  and  $N = \sum_{i=1}^{a} n_i$ . For small values of  $n_1, \ldots, n_a$  ( $a \le 3$  and  $n_i \le 5$ ) use table and for large values of  $n_1, \ldots, n_a$  use that  $T \approx \chi^2(a-1)$  when there is no treatment effect.

## The Friedman test

Assume t treatments and b blocks. Let  $r_{ij}$  be the rank of  $y_{ij}$  within each block, i.e., for each j,  $r_{ij} = 1, \ldots, t$ . Test statistic for the treatments is given by

$$T = \begin{cases} \frac{12S_t}{t(t+1)} - 3b(t+1), & \text{if no ties,} \\ \\ \frac{b(t-1)(S_t - C)}{S_r - C}, & \text{if ties,} \end{cases}$$

where  $s_i = \sum_{j=1}^{b} r_{ij}$ ,  $S_t = \frac{1}{b} \sum_{i=1}^{t} s_i^2$ ,  $S_r = \sum_{i=1}^{t} \sum_{j=1}^{b} r_{ij}^2$  and  $C = \frac{1}{4} bt(t+1)^2$ . For small values of b and t (t = 3, b \le 15 and t = 4, b \le 8) use table and for large values of b and t use that  $T \approx \chi^2(t-1)$  when there is no treatment effect.