This analysis report is concerned with the relationship between working conditions and smoking on biological levels of cotinine. The variables "priming", "barning", "topping", and "other" are indicator variables describing work tasks. The test of whether all cell means for task are equal in the model

log salivary cotinine =  $\beta_0 \cdot \text{priming} + \beta_1 \cdot \text{barning} + \beta_2 \cdot \text{topping} + \beta_3 \cdot \text{other}$ 

is equivalent to testing  $H_0$ :  $\beta_0 = \beta_1 = \beta_2 = \beta_3$ . We failed to reject the null (F-stat = 116.20, df = 3, p-value < 0.0001) and conclude the cell means for the tasks are not equal. Pairwise comparisons were done using Scheffe's method, the results of which are tabulated below.

	Least Squares Means for Effect task							
i	j	Difference Between Means	Simultaneous 95% Confidence Limits for LSMean(i)-LSMean(j)		Degrees of freedom	F-statistic	p-value	
1	2	0.920781	0.337572	1.503991	1, 690	19.57599	0.0002	
1	3	1.673848	1.265368	2.082328	1, 690	131.8714	< 0.0001	
1	4	2.699252	2.284530	3.113975	1, 690	332.6837	< 0.0001	
2	3	0.753067	0.167517	1.338616	1, 690	12.98968	0.0049	
2	4	1.778471	1.188549	2.368393	1, 690	71.37796	< 0.0001	
3	4	1.025404	0.607397	1.443411	1, 690	47.25875	< 0.0001	

Table 1

As can be seen in Table 1, the differences in mean values between each group are all highly significant, where group numbers correspond to the order of tasks listed in the model above.

Parameter	Estimate	Standard Error	Parameter	Estimate	Standard Error
priming	4.508556597	0.10222010	Intercept	4.508556597	0.10222010
barning	3.587775112	0.18127655	barning	-0.920781485	0.20811087
topping	2.834708541	0.10390980	topping	-1.673848056	0.14576075
other	1.809304310	0.10701227	other	-2.699252287	0.14798843
Table 2				Tahla 3	

Tables 2 and 3 display the parameter estimates using cell mean coding and reference cell coding respectively, with task priming as the reference category.

If we wished to test the hypothesis that the average cotinine level for priming workers exceeds that of all other workers, the contrast matrix used would have to account for the different sizes of the barning (n = 69), topping (n = 210) and other (n = 198) task groups so works out to be C = [1 -0.145 -0.440 -0.415] with  $\theta_0$ =0 under cell mean coding, while under reference cell coding C = [0 -0.145 -0.440 -0.415] with the same  $\theta_0$ .

A two-way ANOVA model was fit using a reference cell coding scheme, adding a binary variable "wet" to the previous model, including interaction terms:

log salivary cotinine =  $\beta_0$  +  $\beta_1$ •barning +  $\beta_2$ •topping +  $\beta_3$ •other +  $\beta_4$ •wet +  $\beta_5$ •barning•wet +  $\beta_6$ •topping•wet +  $\beta_7$ •other•wet

The interpretation of the estimate of  $\beta_0$  is the mean log salivary cotinine level of a priming worker in dry conditions. The estimate of  $\beta_1$  is the change in mean log salivary cotinine level comparing a barning worker in dry conditions to a priming worker in dry conditions. Similarly, the estimates of  $\beta_2$  and  $\beta_3$  are analogous for topping and other respectively. The estimate of  $\beta_4$  is the change in mean log salivary cotinine level comparing a priming worker in wet conditions to one in dry conditions. Similarly, the estimates of  $\beta_5$ ,  $\beta_6$  and  $\beta_7$  are analogous for barning, topping and other respectively.

The validity of the HILE Gauss assumptions were confirmed primarily by residual analysis. The histogram and QQ plot confirmed the Gaussian assumption and a scatterplot confirmed homoscedacity. Independence was assumed by study design, existence is guaranteed by finite sample and linearity is implicit in ANOVA.

Table 4 shows the mean values for each of the eight combinations of task and wet.

Parameter	Estimate	Standard Error	Coefficients
Marg Mean: Priming Dry	4.26933727	0.18547121	β <sub>0</sub>
Marg Mean: Barning Dry	3.54274757	С	$\beta_0 + \beta_1$
Marg Mean: Topping Dry	2.68818516	C	$\beta_0 + \beta_2$
Marg Mean: Other Dry	1.80888875	C	$\beta_0 + \beta_3$
Marg Mean: Priming Wet	4.61311604	С	$\beta_0 + \beta_4$
Marg Mean: Barning Wet	3.66702358	С	$\beta_0 + \beta_1 + \beta_4$
Marg Mean: Topping Wet	2.87930261	C	$\beta_0 + \beta_2 + \beta_4$
Marg Mean: Other Wet	1.81053437	C	$\beta_0 + \beta_3 + \beta_4$

Table 4

Another model was fit using a reference cell coding scheme, adding a continuous variable "Innsmoke" (defined as log[1 + # of cigarettes smoked]) to the previous model, including interaction terms:

$$\label{eq:continuous} \begin{split} &\log \ salivary \ cotinine = \beta_0 + \beta_1 \bullet barning + \beta_2 \bullet topping + \beta_3 \bullet other + \beta_4 \bullet wet + \beta_5 \bullet lnnsmoke + \\ &\beta_6 \bullet barning \bullet wet + \beta_7 \bullet topping \bullet wet + \beta_8 \bullet other \bullet wet + \beta_9 \bullet barning \bullet smoke + \beta_{10} \bullet topping \bullet smoke + \\ &\beta_{11} \bullet other \bullet smoke \\ &\beta_{12} \bullet wet \bullet smoke + \beta_{13} \bullet barning \bullet wet \bullet smoke + \beta_{14} \bullet topping \bullet wet \bullet smoke + \\ &\beta_{15} \bullet other \bullet wet \bullet smoke \end{split}$$

The 16 parameter estimates are listed in Table 5 below

Parameter	Estimate	Parameter	Estimate
Intercept	3.883414621	other_wet	-0.327038865
barning	-0.888324180	barning_smoke	0.287856212
topping	-2.066780957	topping_smoke	0.823713934
other	-3.050019611	other_smoke	1.335608893
wet	0.611992088	wet_smoke	-0.283108376
Innsmoke	0.510018497	barning_wet_smoke	0.024005208
barning_wet	-0.266139213	topping_wet_smoke	0.148209637
topping_wet	-0.355995374	other_wet_smoke	0.050316835

Rather than interpret each parameter, a representative one is selected, with other interpretations somewhat analogous given the full model listed above. The estimate of  $\beta_{14}$  is the expected increase in log salivary cotinine with one unit increase in log(1 + # cigarettes smoked) for a topping worker in wet conditions beyond the contributions due to smoking, the interaction of smoking and topping, and the interaction between smoking and wet.

For the ANCOVA model to hold, each interaction term with Innsmoke would have to be insignificant but the interactions with task are highly significant (example is estimate of  $\beta_{11}$ , t-stat = 6.89, p-value < 0.0001). Therefore the ANCOVA model does not hold.

Group-wise backwards selection was chosen as the method to optimize the model. The first group tested were the 3-way interactions, followed by 2-way interactions involved "wet". The selected "best" model was

log salivary cotinine =  $\beta_0$  +  $\beta_1$ •barning +  $\beta_2$ •topping +  $\beta_3$ •other +  $\beta_4$ •wet +  $\beta_5$ •Innsmoke +  $\beta_6$ •barning•smoke +  $\beta_7$ •topping•smoke +  $\beta_8$ •other•smoke

Based on this model, the test for whether wet is significant is equivalent to testing  $H_0$ :  $\beta_4$ =0. We reject the null hypothesis (t-stat = 2.65, df = 1, p-value = 0.0083) and conclude that wet conditions have an effect on log salivary cotinine. Since there are no interactions, the differences lie only between workers in dry conditions and workers in wet conditions.

Based on this model, the test for whether task is significant is equivalent to testing  $H_0$ :  $\beta_1 = \beta_2 = \beta_3 = \beta_6 = \beta_7 = \beta_8 = 0$ . We reject the null hypothesis (F-stat = 18.59, df=6, p-value < 0.0001) and conclude that task has an effect on cotinine level. Furthermore, to determine whether task is significant in regards to 2-way interactions, the test  $H_0$ :  $\beta_6 = \beta_7 = \beta_8 = 0$  was performed. Again, we reject the null (F-stat = 20.27, df=3, p-value < 0.0001). To determine where the differences lie, contrasts were conducted looking at all pairwise comparisons of tasks. Bonferroni correction to p-values was utilized as it is more conservative than Scheffe and less ambiguous in this situation, however given the p-values obtained correction was unneccesary.

The contrasts are displayed in Table 6 below

Contrast	DF	Contrast SS	Mean Square	F Value	Pr > F
Barning Topping	1	47.7708168	47.7708168	37.40	<.0001
Barning Other	1	164.2166069	164.2166069	128.57	<.0001
Topping Other	1	57.3655073	57.3655073	44.91	<.0001

Table 6

For these contrasts,  $H_0$ :  $\beta_i = \beta_j$  where the index corresponds to a task. For comparing barning, topping and other to the reference (priming), the t-values from the full model glm results were used (-5.84, -17.25 and -23.49 respectively, all p-values < 0.0001). As can be seen, the differences lies between each of the task categories.

In summary, cotinine levels are related to working task, wet conditions and cigarettes smoked. For tasks, priming contributes the most to cotinine, followed by barning, topping and other, in that order. Wet conditions also contribute to an increase in cotinine, but no interaction with task or smoking is observed. The greater the number of cigarettes smoked, the higher the cotinine levels observed, which is expected as cotinine is a nicotine metabolite. In addition though, an interaction between smoking and task is also observed, increasing the effect of each task with increasing number of cigarettes smoked.