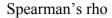
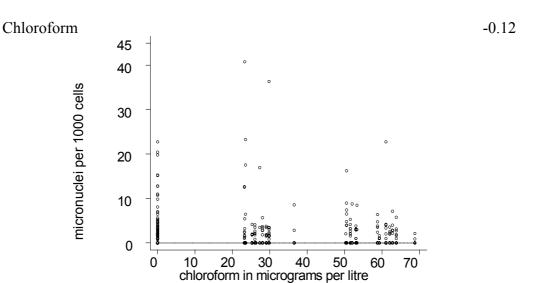
## Appendices

Appendix 1: Scatter plots (with smoothers) examining the relationship between available dose and frequency of micronuclei in bladder epithelial cells	163
Appendix 1a: Scatter plots (with smoothers) examining the relationship between available dose and DNA index	165
Appendix 2: Scatter plots examining the relationship between intake dose and frequency of micronuclei in bladder epithelial cells	167
Appendix 3: Scatter plots examining the relationship between internal dose and frequency of micronuclei in bladder epithelial cells	169
Appendix 3a: Scatter plots examining the relationship between internal dose (excluding outliers) and DNA index from flow cytometry	171
Appendix 4: Correlation coefficients between available dose of THMs and potentially confounding variables	173
Appendix 5: Correlation coefficients between intake dose of THM (from diary) and potentially confounding variables	174
Appendix 6: Correlation coefficients between intake dose of THMs (from questionnaire) and potentially confounding variables	175
Appendix 7: Correlation coefficients between internal dose of THMs and potentially confounding variables	176
Appendix 8: Relative risks for the associations between the potentially confounding variables and frequency of micronuclei	178
Appendix 9: Relative risks in the assessment of confounding in the association between available dose of chloroform and bromoform, and frequency of micronuclei	180
Appendix 10: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by fluid intake diary, and frequency of micronuclei	182
Appendix 11: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by questionnaire, and frequency of micronuclei	184
Appendix 12: Relative risks in the assessment of confounding in the association between internal dose of chloroform, bromodichloromethane, and bromoform, and frequency of micronuclei	186
Appendix 12a: Relative risks in the assessment of confounding in the association between internal dose (excluding outliers) of chloroform, bromodichloromethane, and bromoform, and frequency of micronuclei	188
Appendix 13: Relative risks in the assessment of confounding in the association between available dose of total THM and frequency of micronuclei	190
Appendix 14: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by fluid intake diary, and frequency of micronuclei	191
Appendix 15: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by questionnaire, and frequency of micronuclei	

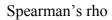
Appendix 16: Relative risks in the assessment of confounding in the association between internal dose of total THM and frequency of micronuclei	. 193
Appendix 16a: Relative risks in the assessment of confounding in the association between internal dose of total THM (excluding outliers) and frequency of micronuclei	. 194
Appendix 17: Relative risks in the assessment of confounding in the association between available dose of chloroform and bromoform, and DNA index from flow cytometry	. 195
Appendix 18: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by fluid intake diary, and DNA index from flow cytometry	. 197
Appendix 19: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by questionnaire, and DNA index from flow cytometry	. 198
Appendix 20: Relative risks in the assessment of confounding in the association between internal dose of chloroform, bromodichloromethane, and bromoform, and DNA index from flow cytometry	. 199
Appendix 21: Relative risks in the assessment of confounding in the association between available dose of total THM and DNA index from flow cytometry	. 201
Appendix 22: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by diary, and DNA index from flow cytometry	. 202
Appendix 23: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by questionnaire, and DNA index from flow cytometry	. 203
Appendix 24: Relative risks in the assessment of confounding in the association between internal dose of total THM and DNA index from flow cytometry	204
Appendix 25: Environmental Health Questionnaire 1	205
Appendix 25a: Environmental Health Questionnaire 2	230
Appendix 26: Fluid intake diary	241
Appendix 27: Introductory letter	246
Appendix 28: Information brochure	248
Appendix 29: Consent form	250
Appendix 30: Interviewers declearation	251
Appendix 31: Instructions for collecting urine sample 1	252
Appendix 32: Instructions for collecting urine sample 2	253
Appendix 33: Randomised controlled trial	254
Appendix 34: Abbreviations	260
Appendix 35: Glossary	261

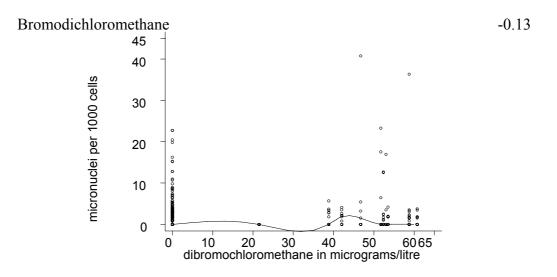
Appendix 1: Scatter plots (with smoothers) examining the relationship between available dose and frequency of micronuclei in bladder epithelial cells

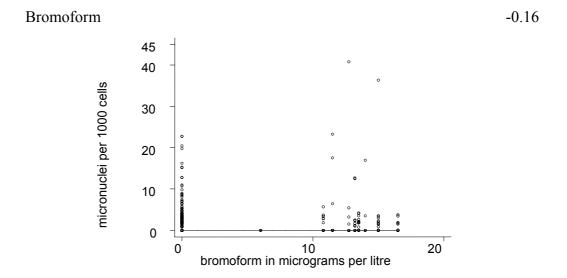


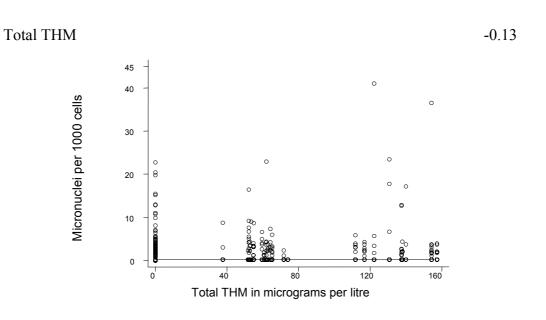


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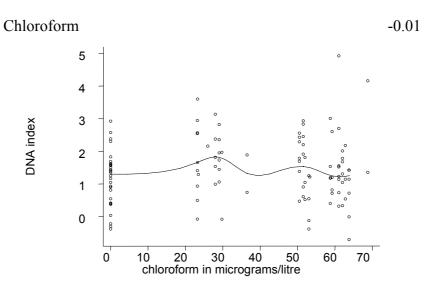


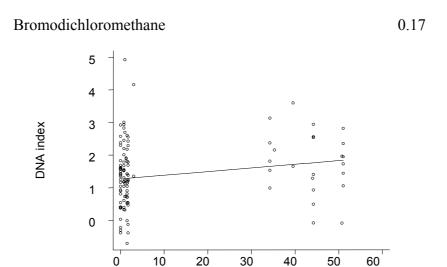




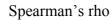
# Appendix 1a: Scatter plots (with smoothers) examining the relationship between available dose and DNA index

Spearman's rho



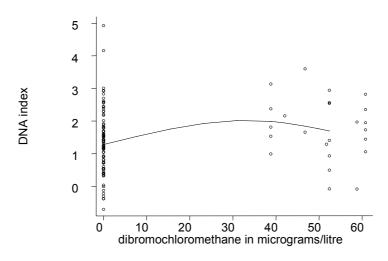


bromodichloromethane in micrograms/litre

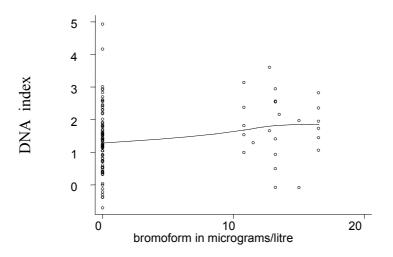


### Dibromochloromethane

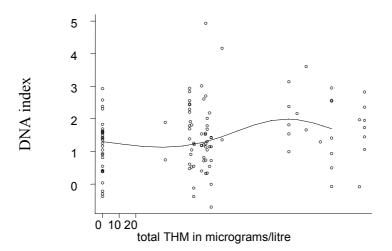
0.20



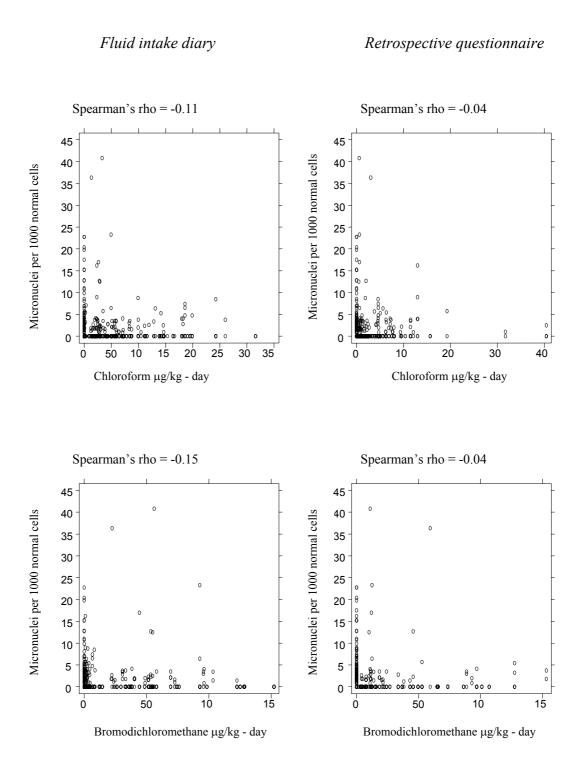
Bromoform 0.21



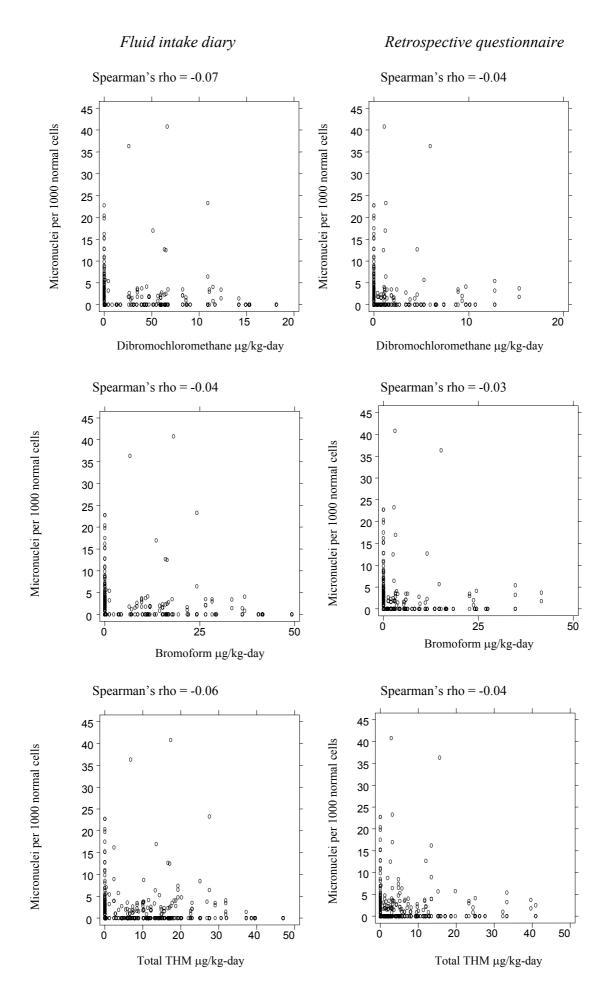
Total THM 0.15



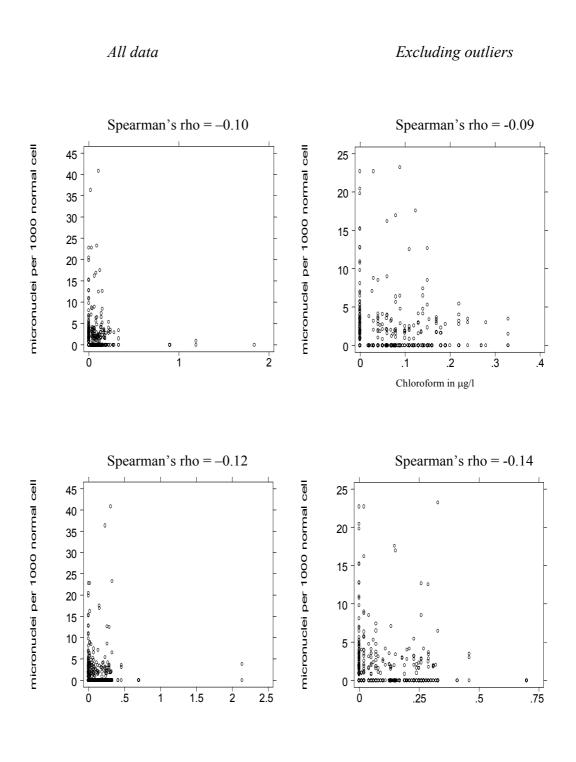
Appendix 2: Scatter plots examining the relationship between intake dose and frequency of micronuclei in bladder epithelial cells



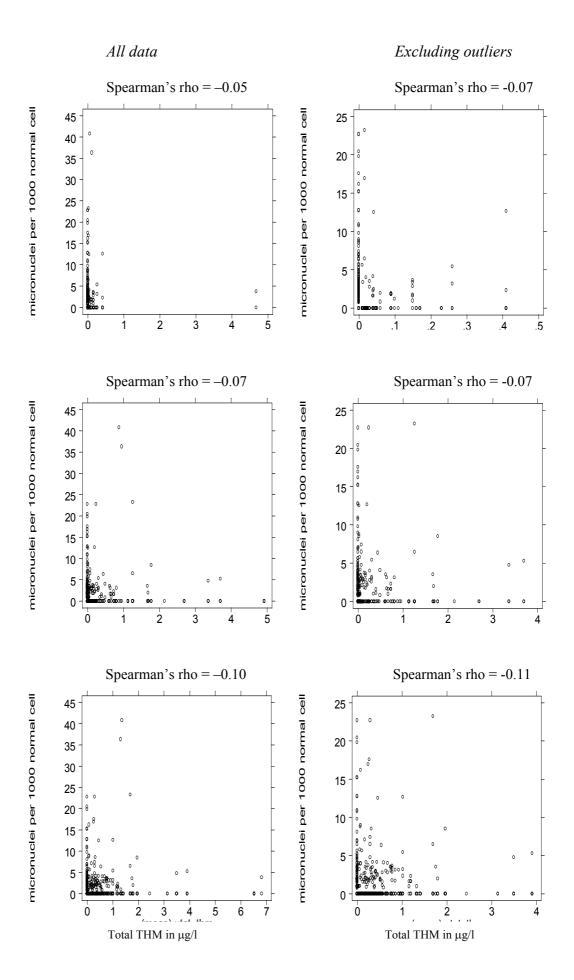
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Appendix 3: Scatter plots examining the relationship between internal dose and frequency of micronuclei in bladder epithelial cells

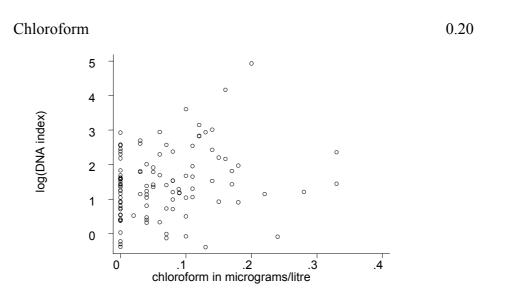


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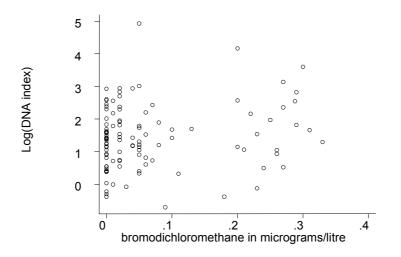


# Appendix 3a: Scatter plots examining the relationship between internal dose (excluding outliers) and DNA index from flow cytometry

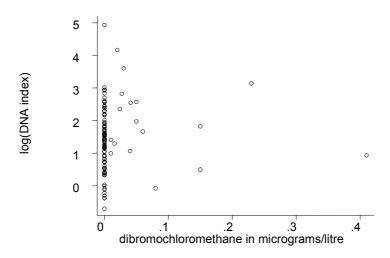
Spearman's rho



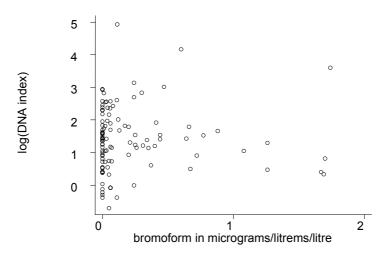
Bromodichloromethane 0.11



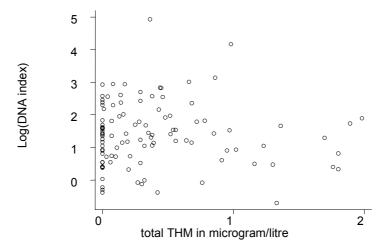
0.17



Bromoform 0.70



Total THM 0.82



Appendix 4: Correlation coefficients between available dose of THMs and potentially confounding variables

	Available dose of			
	chloroform	bromoform	total THM	
Bladder infection prior to last 12 months	0.02	0.01	0.01	
Kidney infection prior to the last 12 months	-0.07	0.00	-0.04	
Family history of bladder cancer	0.11	0.00	0.05	
Worked with dyes - ever	-0.10	-0.03	-0.08	
Worked with dyes- last year	-0.09	-0.06	-0.09	
Worked with chemicals - ever	-0.14	0.03	-0.03	
Worked with electrical cables or making rubber	-0.03	-0.10	-0.10	
Worked with leather - ever	0.07	-0.06	-0.04	
Worked with paint - ever	-0.10	0.08	0.03	
Worked with paint - last year	-0.13	0.17	0.11	
Worked with paint - fortnight	-0.05	0.19	0.15	
Worked as a truck driver - ever	-0.24	-0.07	0.18	
Worked as a truck driver - last year	-0.10	-0.06	-0.10	
Worked as a truck driver - last fortnight	-0.06	-0.08	-0.10	
Used dyes in hobbies - ever	0.04	0.01	0.01	
Used chemicals in hobbies - ever	-0.09	0.04	0.01	
Worked with leather in hobbies - ever	-0.14	-0.04	-0.10	
Worked with paint in hobbies - ever	-0.03	0.12	0.09	
Worked with paint in hobbies - last year	-0.12	0.18	0.10	
Worked with paint in hobbies - fortnight	-0.08	0.10	0.07	
Other hobbies	0.08	0.08	0.09	
Gardening or farming (recreational)	0.03	0.09	0.11	
Use permanent hair dyes - ever	0.02	0.02	0.02	
Used semi-permanent hair dyes - ever	0.00	0.12	0.11	
Used hair rinses - ever	0.10	0.10	0.13	
Smoke tobacco - ever	-0.03	-0.06	-0.08	
Smoke tobacco - now	-0.15	-0.02	-0.11	
Passive smoking	-0.21	-0.02	-0.13	
Passive smoking - workplace	0.08	-0.01	0.02	
Worked with dyes	-0.13	-0.02	-0.07	
Worked with chemicals	-0.02	0.12	0.10	
Worked with paint	-0.03	0.15	0.11	
Worked as a truck driver	-0.10	-0.08	-0.13	
Used paint in hobby	-0.20	-0.04	-0.09	
Other hobbies	-0.08	-0.01	-0.04	
Swimming in pool / chlorinated water	0.02	0.01	0.02	
Smoked cigarettes	-0.14	-0.01	-0.09	
Passive smoking	-0.09	0.02	-0.03	
Serum folate level	0.40	0.09	0.28	
Serum vitamin B12 level	0.09	0.16	0.19	
Age	0.01	0.26	0.24	

Appendix 5: Correlation coefficients between intake dose of THM (from diary) and potentially confounding variables

	Intake	dose from dia	ary of
	chloroform	bromoform	total THM
Bladder infection prior to last 12 months	-0.03	0.01	-0.01
Kidney infection prior to the last 12 months	0.02	0.05	0.04
Family history of bladder cancer	0.18	-0.04	0.07
Worked with dyes - ever	-0.07	-0.06	-0.09
Worked with dyes- last year	-0.02	-0.06	-0.06
Worked with chemicals - ever	-0.03	-0.02	-0.03
Worked with electrical cables or making rubber	-0.09	-0.06	-0.10
Worked with leather - ever	-0.03	-0.05	-0.06
Worked with paint - ever	-0.08	-0.02	-0.06
Worked with paint - last year	-0.04	0.08	0.03
Worked with paint - fortnight	0.01	0.10	0.08
Worked as a truck driver - ever	-0.08	-0.04	-0.06
Worked as a truck driver - last year	-0.06	-0.01	-0.04
Worked as a truck driver - last fortnight	-0.06	-0.11	-0.12
Used dyes in hobbies - ever	0.08	0.01	0.05
Used chemicals in hobbies - ever	-0.02	-0.03	-0.03
Worked with leather in hobbies - ever	-0.10	-0.06	-0.10
Worked with paint in hobbies - ever	-0.03	0.07	0.04
Worked with paint in hobbies - last year	-0.06	0.12	0.06
Worked with paint in hobbies - fortnight	0.00	0.16	0.12
Other hobbies	0.13	0.11	0.15
Gardening or farming (recreational)	0.12	0.16	0.20
Use permanent hair dyes - ever	0.01	-0.06	-0.04
Used semi-permanent hair dyes - ever	0.09	0.02	0.07
Used hair rinses - ever	0.15	0.23	0.25
Smoke tobacco - ever	-0.03	-0.01	-0.03
Smoke tobacco - now	-0.10	0.00	-0.06
Passive smoking	0.14	-0.02	-0.10
Passive smoking - workplace	0.01	-0.03	-0.02
During the study pariod:			
During the study period: Worked with dwar	-0.07	-0.05	-0.07
Worked with dyes			
Worked with noint	0.08 0.03	0.07	0.09
Worked with paint Worked as a truck driver		0.09	0.08
	-0.05	0.01	-0.03
Used paint in hobby	-0.05	-0.05	-0.07
Other hobbies	0.05	0.05	0.07
Swimming in pool / chlorinated water	-0.05	0.04	0.00
Smoked cigarettes	-0.09	0.02	-0.04
Passive smoking	-0.10	-0.06	-0.10
Serum folate level	0.29	0.03	0.20
Serum vitamin B12 level	0.01	0.09	0.08
Age	0.12	0.29	0.29

Appendix 6: Correlation coefficients between intake dose of THMs (from questionnaire) and potentially confounding variables

Intake dose from questionnaire of				
	chloroform	bromoform	total THM	
Bladder infection prior to last 12 months	0.00	0.14	0.03	
Kidney infection prior to the last 12 months	-0.08	0.00	-0.09	
Family history of bladder cancer	0.00	0.00	0.04	
Worked with dyes - ever	-0.08	-0.03	-0.08	
Worked with dyes- last year	-0.04	-0.06	-0.04	
Worked with chemicals - ever	-0.01	0.03	0.09	
Worked with electrical cables or making rubber	-0.06	-0.10	-0.08	
Worked with leather - ever	-0.05	-0.16	-0.06	
Worked with paint - ever	0.03	0.08	0.12	
Worked with paint - last year	-0.01	0.03	0.12	
Worked with paint - fast year  Worked with paint - fortnight	0.03	0.17	0.14	
Worked as a truck driver - ever	-0.16	-0.07	<b>-</b> 0.11	
Worked as a truck driver - last year	-0.10	-0.07	-0.11	
Worked as a truck driver - last fortnight	-0.05	-0.08	-0.03	
Used dyes in hobbies - ever	-0.03	0.01	0.02	
Used chemicals in hobbies - ever	0.00	0.01	0.02	
Worked with leather in hobbies - ever	-0.04	-0.04	<b>-</b> 0.11	
Worked with paint in hobbies - ever	0.00	0.12	0.05	
Worked with paint in hobbies - last year	-0.04	0.12	0.03	
Worked with paint in hobbies - fortnight	-0.04	0.10	-0.05	
Other hobbies	-0.09	0.10	0.05	
Gardening or farming (recreational)	0.05	0.09	0.03	
Use permanent hair dyes - ever	-0.02	0.02	-0.02	
Used semi-permanent hair dyes - ever	0.07	0.02	0.14	
Used hair rinses - ever	-0.04	0.12	-0.05	
Smoke tobacco - ever	-0.07	-0.06	0.00	
Smoke tobacco - now	-0.07	-0.00	0.00	
Passive smoking	-0.11	-0.02	-0.02	
Passive smoking - workplace	-0.11	-0.02	-0.02	
1 assive smoking - workplace	-0.03	-0.01	-0.10	
During the study period:				
Worked with dyes	-0.05	-0.02	-0.06	
Worked with chemicals	0.05	0.12	0.17	
Worked with paint	0.04	0.15	0.14	
Worked as a truck driver	-0.10	-0.08	-0.13	
Used paint in hobby	-0.09	-0.04	-0.10	
Other hobbies	-0.05	-0.01	-0.03	
Swimming in pool / chlorinated water	0.02	0.01	-0.02	
Smoked cigarettes	-0.08	-0.01	0.02	
Passive smoking	-0.08	0.02	-0.05	
Serum folate level	0.10	0.09	0.07	
Serum vitamin B12 level	0.04	0.16	0.16	
Age	-0.04	0.26	0.02	

Appendix 7: Correlation coefficients between internal dose of THMs and potentially confounding variables

• •		•	•	
		Internal dose	of	
	chloroform	bromodichloromethane	bromoform	Total THM
Bladder infection prior to last 12 months	-0.04	0.02	-0.05	-0.04
Kidney infection prior to the last 12 months	-0.03	-0.05	0.07	0.03
Family history of bladder cancer	-0.01	0.17	0.12	0.20
Worked with dyes - ever	-0.02	-0.06	-0.04	-0.05
Worked with dyes- last year	-0.06	-0.06	0.01	-0.02
Worked with chemicals - ever	-0.09	-0.05	-0.06	-0.05
Worked with electrical cables or making rubber	-0.04	-0.01	-0.05	-0.04
Worked with leather - ever	-0.01	0.01	-0.04	-0.03
Worked with paint - ever	0.04	-0.03	-0.11	-0.08
Worked with paint - last year	0.04	0.06	-0.07	-0.03
Worked with paint - fortnight	0.06	0.07	-0.05	-0.01
Worked as a truck driver - ever	-0.14	-0.07	-0.10	-0.08
Worked as a truck driver - last year	-0.04	-0.07	-0.06	-0.07
Worked as a truck driver - last fortnight	-0.01	-0.05	-0.05	-0.05
Used dyes in hobbies - ever	0.10	0.12	-0.04	0.07
Used chemicals in hobbies - ever	0.11	0.12	0.08	0.12
Worked with leather in hobbies - ever	-0.01	0.04	-0.08	-0.02
Worked with paint in hobbies - ever	0.08	0.01	0.03	0.01
Worked with paint in hobbies - last year	0.09	0.04	0.05	0.05
Worked with paint in hobbies - fortnight	0.11	0.02	0.00	0.01
Other hobbies	0.17	0.12	0.24	0.26
Gardening or farming (recreational)	0.00	0.26	0.09	0.18

		Internal dose of		
	chloroform	bromodichloromethane	bromoform	total THM
Use permanent hair dyes - ever	-0.01	-0.04	-0.04	-0.04
Used semi-permanent hair dyes - ever	0.02	0.04	0.09	0.08
Used hair rinses - ever	0.02	0.03	0.04	0.04
Smoke tobacco - ever Smoke tobacco - now	-0.11 -0.02	-0.14 -0.12	-0.10 -0.12	-0.14 -0.14
Passive smoking	-0.07	-0.14	-0.13	-0.16
Passive smoking - workplace	0.01	-0.02	-0.09	-0.07
During the study period:				
Worked with dyes	-0.03	-0.05	-0.04	-0.04
Worked with chemicals	0.12	0.08	0.12	0.13
Worked with paint	0.02	0.06	-0.04	-0.01
Worked as a truck driver	-0.06	-0.09	-0.06	-0.08
Used paint in hobby	-0.08	-0.08	-0.08	-0.09
Other hobbies	0.06	0.06	0.17	0.14
Swimming in pool / chlorinated water	0.00	-0.02	-0.04	-0.04
Smoked cigarettes	-0.08	-0.11	-0.11	-0.12
Passive smoking	-0.02	-0.05	-0.05	-0.05
Serum folate level	0.13	0.10	0.18	0.17
Serum vitamin B12 level	0.01	0.08	0.03	0.04
Age	0.06	0.12	0.14	0.12

Appendix 8: Relative risks for the associations between the potentially confounding variables and frequency of micronuclei

	n positive / n*	relative risk	95% CI	p value
Bladder infection prior to last 12 months	16/228	0.87	0.50 to 1.49	0.6
Kidney infection prior to the last 12 months	12/228	0.99	0.51 to 1.90	0.9
Family history of bladder cancer	8/226	1.16	0.51 to 2.64	0.7
Worked with dyes - ever	20/228	0.94	0.57 to 1.55	0.8
Worked with dyes- last year	9/228	1.22	0.58 to 2.58	0.6
Worked with fulchisin - ever	6/228	1.26	0.60 to 2.63	0.5
Worked with chemicals - ever	106/228	0.80	0.60 to 1.06	0.1
Worked with electrical cables or making rubber	5/228	0.54	0.18 to 1.58	0.3
Worked with leather - ever	3/228	0.78	0.22 to 2.76	0.7
Worked with paint - ever	38/228	0.61	0.41 to 0.92	0.02
Worked with paint - last year	19/228	0.72	0.42 to 1.22	0.2
Worked with paint - fortnight	12/228	0.70	0.35 to 1.39	0.3
Worked as a truck driver - ever	65/228	1.18	0.88 to 1.59	0.3
Worked as a truck driver - last year	22/228	0.91	0.57 to 1.46	0.7
Worked as a truck driver - last fortnight	18/228	1.09	0.66 to 1.81	0.7
Used dyes in hobbies - ever	16/228	0.99	0.58 to 1.67	1.0
Used chemicals in hobbies - ever	114/228	1.07	0.80 to 1.43	0.7
Worked with leather in hobbies - ever	28/228	1.55	1.08 to 2.24	0.02
Worked with paint in hobbies - ever	190/228	1.02	0.68 to 1.52	0.9
Worked with paint in hobbies - last year	120/228	1.29	0.97 to 1.72	0.1
Worked with paint in hobbies - fortnight	31/228	1.10	0.73 to 1.67	0.6
Other hobbies	72/228	1.27	0.94 to 1.72	0.1
Gardening or farming (recreational)	26/228	1.33	0.74 to 2.38	0.3
Use permanent hair dyes - ever	12/228	1.14	0.64 to 2.05	0.7
Used semi-permanent hair dyes - ever	8/228	1.46	0.64 to 3.31	0.4
Used hair rinses - ever	5/228	0.52	0.10 to 2.58	0.4
Smoke tobacco - ever	151/228	1.15	0.85 to 1.57	0.4
Smoke tobacco - now	53/228	1.04	0.74 to 1.44	0.9
Passive smoking	10/228	0.24	0.14 to 1.14	0.6
Passive smoking - workplace	23/228	1.13	0.72 to 1.78	0.6

	n positive / n*	relative risk	95% CI	p value
During the study period:				
Worked with dyes	5/228	1.23	0.46 to 3.24	0.7
Worked with chemicals	22/228	1.16	0.70 to 1.92	0.6
Worked with paint	10/228	0.60	0.25 to 1.43	0.3
Worked as a truck driver	21/228	1.01	0.63 to 1.63	1.0
Used paint in hobby	26/228	1.39	0.91 to 2.10	0.1
Other hobbies	31/228	1.37	0.94 to 1.99	0.1
Swimming in pool / chlorinated water	16/228	0.88	0.46 to 1.69	0.7
Smoked cigarettes	54/228	1.04	0.75 to 1.45	0.8
Passive smoking	17/228	0.85	0.49 to 1.45	0.5
Serum folate level	208/228	1.00	0.98 to 1.01	0.7
Serum vitamin B12 level	208/228	1.00	0.998 to 1.001	0.8
Age	221/228	1.00	0.98 to 1.02	0.9

<sup>\*</sup> number of events reported / total number of respondents

Appendix 9: Relative risks in the assessment of confounding in the association between available dose of chloroform and bromoform, and frequency of micronuclei

Variable	n	n	Relative risk	(%	Relative risk	(%	Relative risk	(%
	observations	participants	Chloroform	change)	Bromoform	change)	AOX	change)
Baseline	605	226	0.985	(0)	0.968	(0)	1.0029	(0)
Bladder infection prior to last 12 months	605	226	0.985	(<1)	0.971	(<1)	1.0028	(<1)
Kidney infection prior to the last 12 months	605	226	0.985	(0)	0.968	(0)	1.0029	(0)
Family history of bladder cancer	605	226	0.985	(<1)	0.970	(<1)	1.0027	(<1)
Worked with dyes - ever	605	226	0.985	(<1)	0.970	(<1)	1.0028	(<1)
Worked with dyes- last year	605	226	0.985	(<1)	0.968	(0)	1.0029	(0)
Worked with fulchisin - ever	605	226	0.985	(<1)	0.969	(<1)	1.0029	(0)
Worked with chemicals - ever	605	226	0.985	(<1)	0.974	(<1)	1.0026	(<1)
Worked with electrical cables or making	601	225	0.986	(<1)	0.973	(<1)	1.0026	(<1)
rubber								
Worked with leather - ever	601	225	0.985	(<1)	0.968	(0)	1.0029	(0)
Worked with paint - ever	605	226	0.986	(<1)	0.977	(<1)	1.0024	(<1)
Worked with pain - last year	605	226	0.985	(0)	0.973	(<1)	1.0027	(<1)
Worked with paint - fortnight	605	226	0.985	(0)	0.972	(<1)	1.0028	(<1)
Worked as a truck driver – ever	605	226	0.985	(0)	0.965	(<1)	1.0031	(<1)
Worked as a truck driver - last year	605	226	0.985	(<1)	0.969	(<1)	1.0028	(<1)
Worked as a truck driver - last fortnight	605	226	0.984	(<1)	0.966	(<1)	1.0030	(<1)
Used dyes in hobbies - ever	605	226	0.985	(<1)	0.968	(0)	1.0029	(0)
Used chemicals in hobbies - ever	605	226	0.985	(<1)	0.968	(0)	1.0029	(0)
Worked with leather in hobbies – ever	605	226	0.987	(<1)	0.977	(<1)	1.0025	(<1)
Worked with paint in hobbies - ever	605	226	0.985	(0)	0.968	(0)	1.0029	(<1)
Worked with paint in hobbies - last year	603	225	0.985	(0)	0.962	(<1)	1.0030	(<1)
Worked with paint in hobbies – fortnight	603	225	0.984	(<1)	0.966	(<1)	1.0030	(<1)
Other hobbies	605	226	0.984	(<1)	0.963	(<1)	1.0030	(<1)

Variable	n	n	Relative risk	(%	Relative risk	(%	Relative risk	(%
	observations	participants	Chloroform	change)	Bromoform	change)	AOX	change)
Gardening or farming (recreational)	605	226	0.985	(<1)	0.969	(<1)	1.0028	(<1)
Use permanent hair dyes - ever	605	226	0.984	(<1)	0.966	(<1)	1.0030	(<1)
Used semi-permanent hair dyes – ever	605	226	0.985	(<1)	0.969	(<1)	1.0028	(<1)
Used hair rinses - ever	605	226	0.985	(<1)	0.968	(0)	1.0030	(<1)
Smoke tobacco - ever	605	226	0.985	(<1)	0.969	(<1)	1.0028	(<1)
Smoke tobacco - now	605	226	0.985	(0)	0.968	(0)	1.0029	(0)
Passive smoking	605	226	0.984	(<1)	0.967	(<1)	1.0030	(<1)
Passive smoking - workplace	577	217	0.983	(<1)	0.962	(<1)	1.0033	(<1)
During the study period:								
Worked with dyes	605	226	0.985	(<1)	0.968	(0)	1.0029	(0)
Worked with chemicals	605	226	0.985	(0)	0.967	(<1)	1.0029	(0)
Worked with paint	605	226	0.985	(<1)	0.975	(<1)	1.0027	(<1)
Worked as a truck driver	605	226	0.985	(<1)	0.967	(<1)	1.0030	(<1)
Used paint in hobby	605	226	0.985	(<1)	0.965	(<1)	1.0031	(<1)
Other hobbies	605	226	0.985	(0)	0.964	(<1)	1.0030	(<1)
Swimming in pool / chlorinated water	605	226	0.984	(<1)	0.966	(<1)	1.0030	(<1)
Smoked cigarettes	605	226	0.985	(0)	0.968	(0)	1.0029	(<1)
Passive smoking	605	226	0.985	(0)	0.970	(<1)	1.0028	(0)
Serum folate level	605	226	0.987	(<1)	0.972	(<1)	1.0026	(<1)
Serum vitamin B12 level	605	226	0.987	(<1)	0.973	(<1)	1.0026	(<1)
Age	604	225	0.985	(0)	0.969	(<1)	1.0026	(<1)

### Notes:

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = available dose for chloroform, bromoform, and AOX (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.

Appendix 10: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by fluid intake diary, and frequency of micronuclei

Variable	n	n	Relative risk	(% change)	Relative risk	(%
	observations	participants	chloroform		Bromoform	change)
Baseline	605	226	1.000	(0)	1.004	(0)
Bladder infection prior to last 12 months	605	226	1.000	(0)	1.005	(<1)
Kidney infection prior to the last 12 months	605	226	1.000	(0)	1.004	(0)
Family history of bladder cancer	605	226	0.999	(<1)	1.004	(0)
Worked with dyes – ever	605	226	1.000	(0)	1.004	(0)
Worked with fulchisin – ever	605	226	1.000	(0)	1.004	(0)
Worked with chemicals – ever	605	226	1.000	(0)	1.004	(0)
Worked with electrical cables or making rubber	601	225	0.999	(<1)	1.004	(0)
Worked with leather – ever	601	225	1.000	(0)	1.004	(0)
Worked with paint – ever	605	226	0.999	(<1)	1.003	(<1)
Worked with paint – last year	605	226	0.999	(0)	1.005	(<1)
Worked with paint – fortnight	605	226	1.000	(0)	1.005	(<1)
Worked as a truck driver – ever	605	226	1.000	(0)	1.005	(<1)
Worked as a truck driver - last year	605	226	1.000	(0)	1.004	(0)
Worked as a truck driver - last fortnight	605	226	1.000	(0)	1.004	(0)
Used dyes in hobbies – ever	605	226	1.000	(0)	1.004	(0)
Used chemicals in hobbies – ever	605	226	1.000	(0)	1.004	(0)
Worked with leather in hobbies – ever	605	226	1.000	(0)	1.007	(<1)
Worked with paint in hobbies – ever	605	226	1.000	(0)	1.004	(0)
Worked with paint in hobbies - last year	603	225	1.000	(0)	1.002	(<1)
Worked with paint in hobbies – fortnight	603	225	1.000	(0)	1.003	(<1)

Variable	n	n	Relative risk	(% change)	Relative risk	(%
	observations	participants	chloroform		Bromoform	change)
Other hobbies	605	226	1.000	(0)	1.002	(<1)
Gardening or farming (recreational)	605	226	0.999	(<1)	1.002	(<1)
Use permanent hair dyes – ever	605	226	1.000	(0)	1.004	(0)
Used semi-permanent hair dyes – ever	605	226	1.000	(0)	1.004	(0)
Used hair rinses - ever	605	226	1.000	(0)	1.006	(<1)
Smoke tobacco - ever	605	226	1.000	(0)	1.004	(0)
Smoke tobacco – now	605	226	1.000	(0)	1.004	(0)
Passive smoking	605	226	1.000	(0)	1.004	(0)
Passive smoking – workplace	577	217	1.000	(0)	1.009	(<1)
During the study period:						
Worked with dyes	605	226	1.000	(0)	1.004	(0)
Worked with chemicals	605	226	1.000	(0)	1.004	(0)
Worked with paint	605	226	1.000	(0)	1.006	(<1)
Worked as a truck driver	605	226	1.000	(0)	1.004	(0)
Used paint in hobby	605	226	1.000	(0)	1.005	(<1)
Other hobbies	605	226	1.000	(0)	1.002	(<1)
Swimming in pool / chlorinated water	605	226	1.000	(0)	1.004	(0)
Smoked cigarettes	605	226	1.000	(0)	1.004	(0)
Passive smoking	605	226	1.000	(0)	1.004	(0)
Serum folate level	605	226	1.000	(0)	1.007	(<1)
Serum vitamin B12 level	605	226	1.000	(0)	1.006	(<1)
Age	604	225	1.000	(0)	1.004	(0)

<sup>•</sup> Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model

<sup>•</sup> Baseline model = intake dose of chloroform and bromoform (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model – i.e. baseline model + listed variable + outcome.

Appendix 11: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by questionnaire, and frequency of micronuclei

Variable	n	n	Relative risk	(% change)	Relative risk	(% change)
	observations	participants	chloroform		Bromoform	
Baseline	605	226	1.000	(0)	1.009	(0)
Bladder infection prior to last 12 months	605	226	1.000	(0)	1.008	(<1)
Kidney infection prior to the last 12 months	605	226	1.000	(0)	1.008	(<1)
Family history of bladder cancer	605	226	1.000	(0)	1.009	(0)
Worked with dyes – ever	605	226	1.000	(0)	1.009	(0)
Worked with dyes- last year	605	226	1.000	(0)	1.008	(<1)
Worked with fulchisin – ever	605	226	1.000	(0)	1.008	(<1)
Worked with chemicals – ever	605	226	1.000	(0)	1.009	(0)
Worked with electrical cables or making rubber	601	225	1.000	(0)	1.009	(0)
Worked with leather – ever	601	225	1.000	(0)	1.009	(0)
Worked with paint – ever	605	226	1.000	(0)	1.012	(<1)
Worked with pain - last year	605	226	1.000	(0)	1.012	(<1)
Worked with paint – fortnight	605	226	1.000	(0)	1.008	(<1)
Worked as a truck driver – ever	605	226	1.000	(0)	1.009	(0)
Worked as a truck driver – last year	605	226	1.000	(0)	1.009	(0)
Worked as a truck driver – last fortnight	605	226	1.000	(0)	1.009	(0)
Used dyes in hobbies – ever	605	226	1.000	(0)	1.009	(0)
Used dyes in hobbies - last year	605	226	1.000	(0)	1.013	(<1)
Used chemicals in hobbies – ever	605	226	1.000	(0)	1.008	(<1)
Worked with leather in hobbies - ever	605	226	1.000	(0)	1.009	(0)
Worked with paint in hobbies – ever	605	226	1.000	(0)	1.009	(0)
Worked with paint in hobbies - last year	603	225	1.000	(0)	1.007	(<1)
Worked with paint in hobbies – fortnight	603	225	1.000	(0)	1.009	(0)

Variable	n	N	Relative risk	(% change)	Relative risk	(% change)
	observations	participants	chloroform	200	Bromoform	
Other hobbies	605	226	1.000	(0)	1.006	(<1)
Gardening or farming (recreational)	605	226	1.000	(0)	1.009	(0)
Use permanent hair dyes – ever	605	226	1.000	(0)	1.009	(0)
Used semi-permanent hair dyes – ever	605	226	1.000	(0)	1.008	(<1)
Used hair rinses – ever	605	226	1.000	(0)	1.009	(0)
Smoke tobacco – ever	605	226	1.000	(0)	1.008	(<1)
Smoke tobacco - now	605	226	1.000	(0)	1.009	(0)
Passive smoking	605	226	1.000	(0)	1.009	(0)
Passive smoking - workplace	577	217	1.000	(0)	1.000	(<1)
During the study period:						
Worked with dyes	605	226	1.000	(0)	1.009	(0)
Worked with chemicals	605	226	1.000	(0)	1.008	(<1)
Worked with paint	605	226	1.000	(0)	1.012	(<1)
Worked as a truck driver	605	226	1.000	(0)	1.009	(0)
Used paint in hobby	605	226	1.000	(0)	1.010	(<1)
Other hobbies	605	226	1.000	(0)	1.006	(<1)
Swimming in pool / chlorinated water	605	226	1.000	(0)	1.009	(0)
Smoked cigarettes	605	226	1.000	(0)	1.009	(0)
Passive smoking	605	226	1.000	(0)	1.009	(0)
Serum folate level	605	226	1.000	(0)	1.011	(<1)
Serum vitamin B12 level	605	226	1.000	(0)	1.011	(<1)
Age	604	225	1.000	(0)	1.009	(0)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose for chloroform and bromoform (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.

Appendix 12: Relative risks in the assessment of confounding in the association between internal dose of chloroform, bromodichloromethane, and bromoform, and frequency of micronuclei

	Relative risk chloroform	(% change)	Relative risk bromodichloromethane	(% change)	Relative risk chloroform	(% change)
Baseline	0.43		1.28		1.12	
Bladder infection prior to the last 12	0.43	(0)	1.29	(<1)	1.11	(<1)
Kidney infection prior to the last 12 months	0.43	(0)	1.28	(0)	1.12	(0)
Family history of bladder cancer	0.42	(2.3)	1.32	(3.1)	1.12	(0)
Worked with dyes - ever	0.42	(2.3)	1.27	(<1)	1.12	(0)
Worked with dyes- last year	0.44	(2.3)	1.29	(<1)	1.11	(<1)
Worked with fulchisin - ever	0.44	(2.3)	1.3	(1.6)	1.12	(0)
Worked with chemicals - ever	0.4	(7.0)	1.3	(1.6)	1.11	(<1)
Worked with electrical cables or making	0.43	(0)	1.3	(1.6)	1.07	(4.5)
Worked with leather - ever	0.42	(2.3)	1.28	(0)	1.11	(<1)
Worked with paint - ever	0.51	(18.6)	1.21	(5.5)	1.07	(4.5)
Worked with paint - last year	0.44	(2.3)	1.28	(0)	1.1	(1.8)
Worked with paint - fortnight	0.45	(4.7)	1.23	(3.9)	1.11	(<1)
Worked as a truck driver - ever	0.43	(0)	1.29	(<1)	1.12	(0)
Worked as a truck driver - last year	0.43	(0)	1.28	(0)	1.11	(<1)
Worked as a truck driver - last fortnight	0.43	(0)	1.29	(<1)	1.12	(0)
Used dyes in hobbies - ever	0.43	(0)	1.29	(<1)	1.11	(<1)
Used chemicals in hobbies - ever	0.43	(0)	1.26	(1.6)	1.11	(<1)
Worked with leather in hobbies - ever	0.43	(0)	1.16	(9.4)	1.16	(3.6)
Worked with paint in hobbies - ever Worked with paint in hobbies - last year	0.42 0.39	(2.3) (9.3)	1.31 1.33	(2.3) (3.9)	1.12 1.11	(0) (<1)
Worked with paint in hobbies - fortnight	0.4	(7.03)	1.28	(0)	1.12	(0)

	Relative risk chloroform	(% change)	Relative risk bromodichloromethane	(% change)	Relative risk chloroform	(% change)
Other hobbies	0.38	(11.6)	1.2	(6.3)	1.08	(3.6)
Gardening or farming (recreational)	0.46	(7.0)	1.19	(7)	1.11	(<1)
Use permanent hair dyes - ever	0.43	(0)	1.28	(0)	1.12	(0)
Used semi-permanent hair dyes - ever	0.44	(2.3)	1.29	(<1)	1.11	(<1)
Used hair rinses - ever	0.43	(0)	1.29	(<1)	1.12	(0)
Smoke tobacco - ever	0.42	(2.3)	1.34	(4.7)	1.13	(<1)
Smoke tobacco - now	0.43	(0)	1.29	(<1)	1.12	(0)
Passive smoking	0.46	(7.0)	1.29	(<1)	1.11	(<1)
Passive smoking - workplace	0.38	(11.6)	1.25	(2.3)	1.13	(<1)
During the study period: Worked with dyes	0.44	(2.3)	1.29	(<1)	1.12	(0)
Worked with chemicals	0.42	(2.3)	1.29	(<1)	1.11	(<1)
Worked with paint	0.45	(4.7)	1.29	(<1)	1.11	(<1)
Worked as a truck driver	0.43	(0)	1.28	(0)	1.12	(0)
Used paint in hobby	0.45	(4.7)	1.29	(<1)	1.11	(<1)
Other hobbies	0.43	(0)	1.27	(<1)	1.09	(2.7)
Swimming in pool / chlorinated water	0.43	(0)	1.28	(0)	1.11	(<1)
Smoked cigarettes	0.44	(2.3)	1.29	(<1)	1.12	(0)
Passive smoking	0.42	(2.3)	1.27	(<1)	1.11	(<1)
Serum folate level	0.45	(4.7)	1.32	(3.1)	1.11	(<1)
Serum vitamin B12 level	0.44	(2.3)	1.32	(3.1)	1.1	(1.8)
Age	0.43	(0)	1.29	(<1)	1.11	(<1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = internal dose of chloroform, bromodichloromethane and bromoform (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.
- Highlighted are variable that change the exposure-outcome association by five percent of more, therefore included as a confounder in the final model

Appendix 12a: Relative risks in the assessment of confounding in the association between internal dose (excluding outliers) of chloroform, bromodichloromethane, and bromoform, and frequency of micronuclei

	Relative risk chloroform	(% change)	Relative risk bromodichloromethane	(% change)	Relative risk bromoform	(% change)
Baseline	1.23	change)	1.17	change)	1.17	change)
Bladder infection prior to the last 12 months	1.2	(2.4)	1.26	(7.8)	1.17	(0)
Kidney infection prior to the last 12 months	1.24	(<1)	1.15	(1.7)	1.18	(<1)
Family history of bladder cancer	1.24	(<1)	1.16	(<1)	1.18	(<1)
Worked with dyes - ever	1.2	(2.4)	1.17	(0)	1.17	(0)
Worked with dyes- last year	1.31	(6.5)	1.18	(<1)	1.17	(0)
Worked with fulchisin - ever	1.29	(4.9)	1.21	(3.4)	1.18	(<1)
Worked with chemicals - ever	1.15	(6.5)	1	(14.5)	1.18	(<1)
Worked with electrical cables or making rubber	0.98	(20.3)	1.37	(17.1)	1.12	(4.3)
Worked with leather - ever	1.17	(4.9)	1.19	(1.7)	1.17	(0)
Worked with paint - ever	1.42	(15.5)	1.02	(12.8)	1.12	(4.3)
Worked with paint - last year	1.34	(8.9)	1.17	(0)	1.15	(1.7)
Worked with paint - fortnight	1.4	(13.8)	1.18	(<1)	1.15	(1.7)
Worked as a truck driver - ever	1.17	(4.9)	1.19	(1.7)	1.18	(<1)
Worked as a truck driver - last year	1.27	(3.3)	1.15	(1.7)	1.17	(0)
Worked as a truck driver - last fortnight	1.17	(4.9)	1.19	(1.7)	1.18	(<1)
Used dyes in hobbies - ever	1.23	(0)	1.17	(0)	1.17	(0)
Used chemicals in hobbies - ever	1.3	(5.7)	1.11	(5.1)	1.17	(0)
Worked with leather in hobbies - ever	1.53	(24.4)	1.31	(11.9)	1.21	(3.4)
Worked with paint in hobbies - ever	1.24	(<1)	1.15	(1.7)	1.17	(0)
Worked with paint in hobbies - last year	1.55	(26)	0.94	(19.7)	1.17	(0)
Worked with paint in hobbies - fortnight	1.22	(<1)	1.09	(6.8)	1.16	(<1)

	Relative risk chloroform	(% change)	Relative risk bromodichloromethane	(% change)	Relative risk bromoform	(% change)
Other hobbies	0.84	(31.7)	1.29	(10.3)	1.13	(3.4)
Gardening or farming (recreational)	1.33	(8.1)	1.12	(4.3)	1.16	(<1)
Use permanent hair dyes - ever	1.21	(1.6)	1.15	(1.17)	1.18	(<1)
Used semi-permanent hair dyes - ever	1.26	(2.4)	1.17	(0)	1.16	(<1)
Used hair rinses - ever	1.23	(0)	1.23	(5.1)	1.18	(<1)
Smoke tobacco - ever	1.02	(17.1)	1.25	(6.8)	1.19	(1.7)
Smoke tobacco - now	1.29	(4.9)	1.2	(2.6)	1.18	(<1)
Passive smoking	1.2	(2.4)	1.23	(5.1)	1.18	(<1)
Passive smoking - workplace	0.93	(24.4)	1.15	(1.7)	1.18	(<1)
During the study period: Worked with dyes Worked with chemicals	1.28 1.19	(4.1) (3.3)	1.19 1.18	(1.7) (<1)	1.17 1.17	(0) (0)
Worked with paint	1.37	(11.4)	1.22	(4.3)	1.16	(<1)
Worked as a truck driver	1.23	(0)	1.18	(<1)	1.17	(0)
Used paint in hobby	1.37	(11.4)	1.22	(4.3)	1.16	(<1)
Other hobbies	1.44	(17.1)	0.98	(16.2)	1.15	(1.7)
Swimming in pool / chlorinated water	1.25	(1.6)	1.15	(1.7)	1.17	(0)
Smoked cigarettes	1.3	(5.7)	1.19	(1.7)	1.18	(<1)
Passive smoking	1.19	(3.34)	1.18	(<1)	1.17	(0)
Serum folate level	1.4	(13.8)	1.55	(32.5)	1.17	(0)
Serum vitamin B12 level	1.14	(7.3)	1.63	(39.3)	1.16	(<1)
Age	1.23	(0)	1.18	(<1)	1.17	(0)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = internal dose of chloroform, bromodichloromethane and bromoform (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.
- Highlighted are variable that change the exposure-outcome association by five percent of more, therefore included as a confounder in the final model

Appendix 13: Relative risks in the assessment of confounding in the association between available dose of total THM and frequency of micronuclei

Variable	n	n	Relative risk	(%
	observations	participants	total THM	change)
Baseline	605	226	1.0002	(0)
Bladder infection prior to last 12 months	605	226	1.0002	(0)
Family history of bladder cancer	605	226	1.0002	(0)
Worked with dyes – ever	605	226	1.0002	(0)
Worked with dyes- last year	605	226	1.0002	(0)
Worked with chemicals – ever	605	226	1.0002	(0)
Worked with electrical cables/making rubber	601	225	1.0000	(<1)
Worked with leather – ever	601	225	1.0002	(0)
Worked with paint - ever	605	226	1.0001	(<1)
Worked with pain - last year	605	226	1.0002	(0)
Worked with paint - fortnight	605	226	1.0002	(0)
Worked as a truck driver - ever	605	226	1.0002	(0)
Worked as a truck driver - last year	605	226	1.0002	(0)
Worked as a truck driver - last fortnight	605	226	1.0002	(0)
Used dyes in hobbies - ever	605	226	1.0002	(0)
Used chemicals in hobbies - ever	605	226	1.0002	(0)
Worked with leather in hobbies – ever	605	226	1.0001	(<1)
Worked with paint in hobbies – ever	605	226	1.0002	(0)
Worked with paint in hobbies - last year	603	225	1.0002	(0)
Worked with paint in hobbies – fortnight	603	225	1.0002	(0)
Other hobbies	605	226	1.0002	(0)
Gardening or farming (recreational)	605	226	1.0002	(0)
Use permanent hair dyes – ever	605	226	1.0002	(0)
Used semi-permanent hair dyes – ever	605	226	1.0002	(0)
Used hair rinses – ever	605	226	1.0002	(0)
Smoke tobacco – ever	605	226	1.0002	(0)
Smoke tobacco – now	605	226	1.0003	(<1)
Passive smoking	603	225	1.0002	(0)
Passive smoking - workplace	577	217	1.0002	(0)
During the study period:	60.5	226	1.0000	(0)
Worked with dyes	605	226	1.0002	(0)
Worked with chemicals	605	226	1.0002	(0)
Worked with paint	605	226	1.0002	(0)
Worked as a truck driver	605	226	1.0002	(0)
Used paint in hobby	605	226	1.0006	(<1)
Other hobbies	605	226	1.0002	(0)
Swimming in pool / chlorinated water	605	226	1.0002	(0)
Smoked cigarettes	605	226	1.0002	(0)
Passive smoking	605	226	1.0002	(0)
Age	604	225	1.0002	(0)
Serum folate level	556	206	1.0002	(0)
Serum vitamin B12 level	556	206	1.0004	(<1)

<sup>•</sup> Baseline model = available dose total THM (exposure) and frequency of micronuclei per 1000 normal cells (outcome).

<sup>•</sup> Each subsequent line represents a multivariate model – i.e. baseline model + listed variable + outcome.

Appendix 14: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by fluid intake diary, and frequency of micronuclei

Variable	n observations	n participants	Relative risk total THM	(% change)
Baseline	605	226	1.0001	
Bladder infection prior to last 12 months	605	226	1.0001	(<1)
Kidney infection prior to the last 12	605	226	1.0002	(0)
months				. ,
Family history of bladder cancer	605	226	1.0001	(0)
Worked with dyes - ever	605	226	1.0001	(0)
Worked with dyes- last year	605	226	1.0002	(<1)
Worked with fulchisin - ever	605	226	1.0001	(0)
Worked with chemicals - ever	605	226	1.0001	(0)
Worked with electrical cables or making rubber	605	226	1.0000	(<1)
Worked with leather - ever	605	226	1.0001	(0)
Worked with paint - ever	605	226	1.0000	(<1)
Worked with paint - last year	605	226	1.0001	(0)
Worked with paint - fortnight	605	226	1.0001	(0)
Worked as a truck driver - ever	605	226	1.0001	(<1)
Worked as a truck driver - last year	605	226	1.0002	(0)
Worked as a truck driver - last fortnight	605	226	1.0001	(<1)
Used dyes in hobbies - ever	605	226	1.0002	(0)
Used chemicals in hobbies - ever	605	226	1.0001	(0)
Worked with leather in hobbies - ever	605	226	1.0004	(<1)
Worked with paint in hobbies - ever	605	226	1.0004	(0)
Worked with paint in hobbies - last year	605	226	1.0001	(0)
Worked with paint in hobbies - fortnight	605	226	1.0001	(0)
Other hobbies	605	226	1.0001	(<1)
Gardening or farming (recreational)	605	226	1.0000	(<1)
Use permanent hair dyes - ever	605	226	1.0001	(0)
Used semi-permanent hair dyes - ever	605	226	1.0001	(0)
Used hair rinses - ever	605	226	1.0001	` '
Smoke tobacco - ever	605			(<1)
		226	1.0002	(<1)
Smoke tobacco - now	605	226	1.0001	(0)
Passive smoking	605	226	1.0002	(<1)
Passive smoking - workplace	605	226	1.0005	(<1)
During the study period:	605	226	1 0002	(<1)
Worked with dyes	605	226	1.0002	(<1)
Worked with chemicals	605	226	1.0001	(0)
Worked with paint	605	226	1.0002	(<1)
Worked as a truck driver	605	226	1.0001	(0)
Used paint in hobby	605	226	1.0003	(<1)
Other hobbies	605	226	1.0000	(<1)
Swimming in pool / chlorinated water	605	226	1.0001	(0)
Smoked cigarettes	605	226	1.0001	(0)
Passive smoking	605	226	1.0002	(<1)
Serum folate level	605	226	1.0005	(<1)
Serum vitamin B12 level	605	226	1.0003	(<1)
Age	605	226	1.0001	(0)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of total THM (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.

Appendix 15: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by questionnaire, and frequency of micronuclei

	n	n	Relative risk	(%
	observations		total THM	change)
Baseline	605	226	1.0006	
Bladder infection prior to last 12 months	605	226	1.0006	(0)
Kidney infection prior to the last 12 months	605	226	1.0006	(0)
Family history of bladder cancer	605	226	1.0005	(<1)
Worked with dyes - ever	605	226	1.0005	(<1)
Worked with dyes- last year	605	226	1.0006	(0)
Worked with fulchisin - ever	605	226	1.0005	(<1)
Worked with chemicals - ever	605	226	1.0007	(<1)
Worked with electrical cables or making rubber	601	225	1.0005	(<1)
Worked with leather - ever	601	225	1.0005	(<1)
Worked with paint - ever	605	226	1.0007	(<1)
Worked with paint - last year	605	226	1.0007	(<1)
Worked with paint - fortnight	605	226	1.0008	(<1)
Worked as a truck driver - ever	605	226	1.0007	(<1)
Worked as a truck driver - last year	605	226	1.0006	(0)
Worked as a truck driver - last fortnight	605	226	1.0005	(<1)
Used dyes in hobbies - ever	605	226	1.0006	(0)
Used chemicals in hobbies - ever	605	226	1.0005	(<1)
Worked with leather in hobbies - ever	605	226	1.0008	(<1)
Worked with paint in hobbies - ever	605	226	1.0006	(0)
Worked with paint in hobbies - last year	603	225	1.0005	(<1)
Worked with paint in hobbies – fortnight	603	225	1.0005	(<1)
Other hobbies	605	226	1.0004	(<1)
Gardening or farming (recreational)	605	226	1.0005	(<1)
Use permanent hair dyes – ever	605	226	1.0006	(0)
Used semi-permanent hair dyes - ever	605	226	1.0005	(<1)
Used hair rinses – ever	605	226	1.0005	(<1)
Smoke tobacco – ever	605	226	1.0005	(<1)
Smoke tobacco – now	605	226	1.0005	(<1)
Passive smoking	605	226	1.0006	(0)
Passive smoking – workplace	577	217	1.0000	(<1)
During the study period:				
Worked with dyes	605	226	1.0006	(0)
Worked with chemicals	605	226	1.0005	(<1)
Worked with paint	605	226	1.0008	(<1)
Worked as a truck driver	605	226	1.0006	(0)
Used paint in hobby	605	226	1.0008	(<1)
Other hobbies	605	226	1.0005	(<1)
Swimming in pool / chlorinated water	605	226	1.0006	(0)
Smoked cigarettes	605	226	1.0005	(<1)
Passive smoking	605	226	1.0005	(<1)
Serum folate level	605	226	1.0007	(<1)
Serum vitamin B12 level	605	226	1.0007	(<1)
Age	604	225	1.0006	(0)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of total THM (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.

Appendix 16: Relative risks in the assessment of confounding in the association between internal dose of total THM and frequency of micronuclei

	n	n	Relative risk	
	observations		total THM	(% change)
Baseline	605	226	1.05	· · · · · · · · · · · · · · · · · · ·
Bladder infection prior to last 12 months	605	226	1.05	(0)
Kidney infection prior to the last 12 months	605	226	1.05	(0)
Family history of bladder cancer	605	226	1.04	(<1)
Worked with dyes - ever	605	226	1.05	(0)
Worked with dyes- last year	605	226	1.05	(0)
Worked with fulchisin - ever	605	226	1.05	(0)
Worked with chemicals - ever	605	226	1.05	(0)
Worked with electrical cables or making rubber	601	225	1.03	(1.9)
Worked with leather - ever	601	225	1.04	(<1)
Worked with paint - ever	605	226	1.03	(1.9)
Worked with paint - last year	605	226	1.04	(<1)
Worked with paint - fortnight	605	226	1.04	(<1)
Worked as a truck driver - ever	605	226	1.05	(0)
Worked as a truck driver - last year	605	226	1.05	(0)
Worked as a truck driver - last fortnight	605	226	1.05	(0)
Used dyes in hobbies - ever	605	226	1.05	(0)
Used chemicals in hobbies - ever	605	226	1.04	(<1)
Worked with leather in hobbies - ever	605	226	1.04	(<1)
Worked with paint in hobbies - ever	605	226	1.05	(0)
Worked with paint in hobbies - last year	603	225	1.05	(0)
Worked with paint in hobbies - fortnight	603	225	1.04	(<1)
Other hobbies	605	226	1.01	(3.8)
Gardening or farming (recreational)	605	226	1.03	(1.9)
Use permanent hair dyes - ever	605	226	1.05	(0)
Used semi-permanent hair dyes – ever	605	226	1.05	(0)
Used hair rinses – ever	605	226	1.05	(0)
Smoke tobacco – ever	605	226	1.05	(0)
Smoke tobacco – now	605	226	1.05	(0)
Passive smoking	603	225	1.05	(0)
Passive smoking – workplace	577	217	1.05	(0)
During the study period:				
Worked with dyes	605	226	1.05	(0)
Worked with chemicals	605	226	1.04	(0)
Worked with paint	605	226	1.05	(0)
Worked as a truck driver	605	226	1.05	(0)
Used paint in hobby	605	226	1.06	(<1)
Other hobbies	605	226	1.04	(<1)
Swimming in pool / chlorinated water	605	226	1.05	(0)
Smoked cigarettes	605	226	1.05	(0)
Passive smoking	605	226	1.04	(<1)
Serum folate level	556	225	1.05	(0)
Serum vitamin B12 level	556	225	1.05	(0)
Age	604	225	1.05	(0)

<sup>•</sup> Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model

<sup>•</sup> Baseline model = internal dose of total THM (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model – i.e. baseline model + listed variable + outcome.

Appendix 16a: Relative risks in the assessment of confounding in the association between internal dose of total THM (excluding outliers) and frequency of micronuclei

	n	n	Relative risk	(%
	observations	participants	total THM	change)
Baseline	601	220	1.18	
Bladder infection prior to last 12 months	601	220	1.18	(0)
Kidney infection prior to the last 12 months	601	220	1.17	(<1)
Family history of bladder cancer	601	220	1.19	(<1)
Worked with dyes - ever	601	220	1.17	(<1)
Worked with dyes- last year	601	220	1.18	(0)
Worked with fulchisin - ever	601	220	1.19	(<1)
Worked with chemicals - ever	601	220	1.16	(1.17)
Worked with electrical cables or making	597	219	1.13	(4)
Worked with leather - ever	597	219	1.17	(<1)
Worked with paint - ever	601	220	1.14	(3.4)
Worked with paint - last year	601	220	1.17	(<1)
Worked with paint - fortnight	601	220	1.17	(<1)
Worked as a truck driver - ever	601	220	1.20	(1.17)
Worked as a truck driver - last year	601	220	1.17	(<1)
Worked as a truck driver - last fortnight	601	220	1.18	(0)
Used dyes in hobbies - ever	601	220	1.18	(0)
Used chemicals in hobbies - ever	601	220	1.17	(<1)
Worked with leather in hobbies – ever	601	220	1.24	(5.1)
Worked with paint in hobbies – ever	601	220	1.18	(0)
Worked with paint in hobbies - last year	599	219	1.17	(<1)
Worked with paint in hobbies – fortnight	599	219	1.16	(1.17)
Other hobbies	60	220	1.12	(5.1)
Gardening or farming (recreational)	601	220	1.17	(<1)
Use permanent hair dyes - ever	601	220	1.18	(0)
Used semi-permanent hair dyes - ever	601	220	1.17	(<1)
Used hair rinses - ever	601	220	1.19	(<1)
Smoke tobacco - ever	601	220	1.19	(<1)
Smoke tobacco - now	601	220	1.19	(<1)
Passive smoking	569	211	1.18	(0)
Passive smoking – workplace	569	211	1.16	(1.17)
During the study period:				
Worked with dyes	601	220	1.19	(<1)
Worked with chemicals	601	220	1.17	(<1)
Worked with paint	601	220	1.18	(0)
Worked as a truck driver	601	220	1.18	(0)
Used paint in hobby	601	220	1.21	(2.5)
Other hobbies	601	220	1.16	(1.17)
Swimming in pool / chlorinated water	601	220	1.17	(<1)
Smoked cigarettes	601	220	1.19	(<1)
Passive smoking	601	220	1.17	(<1)
Serum folate level	550	201	1.21	(2.5)
Serum vitamin B12 level	550	201	1.20	(1.7)
Age	601	220	1.18	(0)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = internal dose of total THM (exposure), and frequency of micronuclei per 1000 normal cells (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.

Appendix 17: Relative risks in the assessment of confounding in the association between available dose of chloroform and bromoform, and DNA index from flow cytometry

	Relative risk Chloroform	(% change)	Relative risk Bromoform	(% change)	Relative risk AOX	(% change)
Baseline	0.9996		1.012		1.001	
Bladder infection prior to 12 months	0.9989	(<1)	1.008	(<1)	1.002	(<1)
Kidney infection prior to 12 months	0.9983	(<1)	1.005	(<1)	1.002	(<1)
Family history of bladder cancer	0.9995	(<1)	1.014	(<1)	1.001	(<1)
Ever worked with dyes	0.9985	(<1)	1.008	(<1)	1.002	(<1)
Worked with dyes in the last year	0.9995	(<1)	1.010	(<1)	1.001	(<1)
Ever worked with chemicals	0.9997	(<1)	1.009	(<1)	1.001	(<1)
Ever worked making rubber or electrical cables	0.9991	(<1)	1.011	(<1)	1.001	(<1)
Ever worked with leather	0.9996	(<1)	1.012	(<1)	1.001	(<1)
Ever worked with paint	0.9993	(<1)	1.009	(<1)	1.002	(<1)
Ever worked with paint	0.9993	(<1)	1.009	(<1)	1.002	(<1)
Worked with paint in the last year	0.9986	(<1)	1.004	(<1)	1.002	(<1)
Worked with paint in the last fortnight	0.9988	(<1)	1.008	(<1)	1.002	(<1)
Ever worked as a truck driver	0.9999	(<1)	1.017	(<1)	1.001	(<1)
Worked as a truck driver in the last year	0.9997	(<1)	1.012	(<1)	1.001	(<1)
Worked as a truck driver in the last fortnight	0.9995	(<1)	1.012	(<1)	1.001	(<1)
Ever used dyes in hobbies	0.9985	(<1)	1.006	(<1)	1.002	(<1)
Ever worked with leather as part of a hobby	0.9988	(<1)	1.010	(<1)	1.001	(<1)
Ever used paint in a hobby	0.9995	(<1)	1.009	(<1)	1.001	(<1)
Used paint in a hobby in the last year	1.0000	(<1)	1.012	(<1)	1.001	(<1)

	Relative risk chloroform	(% change)	Relative risk Bromoform	(% change)	Relative risk AOX	(% change)
Used paint in a hobby in the last fortnight	0.9984	(<1)	1.011	(<1)	1.001	(<1)
Ever used permanent hair dyes	1.0002	(<1)	1.017	(<1)	1.001	(<1)
Ever used semi-permanent hair dyes	0.9994	(<1)	1.011	(<1)	1.001	(<1)
Ever used hair rinses	0.9991	(<1)	1.011	(<1)	1.001	(<1)
Ever smoked tobacco	1.0005	(<1)	1.018	(<1)	1.001	(<1)
Passive smoking home	1.0030	(<1)	1.016	(<1)	1.001	(<1)
Passive smoking workplace	1.0001	(<1)	1.015	(<1)	1.001	(<1)
During study period:						
Used dyes at workplace	1.0014	(<1)	1.025	(<1)	1.001	(<1)
Used chemicals in workplace	0.9990	(<1)	1.009	(<1)	1.002	(<1)
Used paint in the workplace	0.9996	(<1)	1.012	(<1)	1.001	(<1)
Worked as a truck driver	0.9996	(<1)	1.013	(<1)	1.001	(<1)
Used paint in a hobby	0.9992	(<1)	1.009	(<1)	1.002	(<1)
Swimming in chlorinated water	0.9982	(<1)	1.004	(<1)	1.002	(<1)
Smoke tobacco	0.9991	(<1)	1.010	(<1)	1.002	(<1)
Passive smoking – home	0.9996	(<1)	1.012	(<1)	1.001	(<1)
Passive smoking – workplace	0.9999	(<1)	1.012	(<1)	1.001	(<1)
Vitamin B12	1.0013	(<1)	1.022	(<1)	1.001	(<1)
Folate	0.9999	(<1)	1.019	(<1)	1.001	(<1)
Age	0.9997	(<1)	1.012	(<1)	1.001	(<1)

#### Notes:

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = available dose for chloroform, bromoform, and AOX (exposure), and log transformed DNA index (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.

Appendix 18: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by fluid intake diary, and DNA index from flow cytometry

				<u>.</u>
n=112	Relative risk	(%	Relative risk	(%
	chloroform	change)	bromoform	change)
Baseline	1.0018	(0)	1.01	(0)
Bladder infection prior to 12 months	1.0019	(<1)	1.01	(0)
Kidney infection prior to 12 months	1.0018	(0)	1.01	(0)
Family history of bladder cancer	1.0016	(<1)	1.01	(0)
Ever worked with dyes	1.0019	(<1)	1.00	(<1)
Worked with dyes in the last year	1.0020	(<1)	1.01	(0)
Ever worked with chemicals	1.0021	(<1)	1.00	(<1)
Ever worked making rubber or electrical cables	1.0017	(<1)	1.00	(<1)
Ever worked with leather	1.0018	(0)	1.01	(0)
Ever worked with paint	1.0018	(0)	1.01	(0)
Worked with paint in the last year	1.0017	(<1)	1.00	(<1)
Worked with paint in the last fortnight	1.0017	(<1)	1.00	(<1)
Ever worked as a truck driver	1.0010	(<1)	1.00	(0)
Worked as a truck driver in the last year	1.0017	` ′	1.01	(0)
•		(0)		
Worked as a truck driver in the last fortnight	1.0018	(0)	1.01	(0)
Ever worked in an aluminium smelter	1.0019	(<1)	1.01	(0)
Ever used dyes in hobbies	1.0022	(<1)	1.00	(<1)
Used dyes in hobbies in the last year	1.0021	(<1)	1.00	(<1)
Worked with leather as part of a hobby in the last	1.0015	(<1)	1.00	(<1)
year Ever used paint in a hobby	1.0020	(<1)	1.00	(<1)
Used paint in a hobby in the last year	1.0020	(<1) (<1)	1.00	` /
		` ′		(0)
Used paint in a hobby in the last fortnight	1.0018	(0)	1.01	(0)
Ever used permanent hair dyes	1.0018	(0)	1.01	(0)
Used permanent hair dyes in the last year	1.0019	(<1)	1.01	(0)
Ever used semi-permanent hair dyes	1.0015	(<1)	1.01	(0)
Used semi-permanent hair dyes in the last year	1.0015	(<1)	1.01	(0)
Ever used hair rinses	1.0017	(<1)	1.00	(<1)
Ever smoked tobacco	1.0018	(0)	1.01	(0)
Passive smoking home	1.0000	(<1)	1.00	(<1)
Passive smoking workplace	1.0018	(0)	1.00	(<1)
During study period:				
Used dyes at workplace	1.0017	(<1)	1.00	(<1)
Used chemicals in workplace	1.0017	(<1)	1.00	(0)
Used paint in the workplace	1.0015	(<1)	1.00	(<1)
Worked as a truck driver	1.0018	(<1)	1.01	(0)
Used paint in a hobby	1.0019	(<1)	1.00	(<1)
Swimming in chlorinated water	1.0019	(<1)	1.01	(0)
Smoke tobacco	1.0020	(<1)	1.01	(0)
Passive smoking – home	1.0018	(<1)	1.01	(0)
Passive smoking – workplace	1.0018	(0)	1.01	(0)
Vitamin B12	1.0018	(0)	1.01	(0)
Folate	1.0016	(<1)	1.01	(0)
Age	1.0018	(0)	1.00	(<1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of chloroform and bromoform (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome

Appendix 19: Relative risks in the assessment of confounding in the association between intake dose of chloroform and bromoform as estimated by questionnaire, and DNA index from flow cytometry

n=112	relative risk	(%	relative risk (%
	chloroform	change)	bromoform change)
Baseline	1.0002	(0)	0.999 (<1)
Bladder infection prior to 12 months	1.0003	(<1)	0.999 (<1)
Kidney infection prior to 12 months	1.0002	(0)	0.999 (<1)
Family history of bladder cancer	1.0003	(0)	1.000 (<1)
Ever worked with dyes	1.0001	(<1)	0.997 (<1)
Worked with dyes in the last year	1.0003	(0)	0.999 (<1)
Ever worked with chemicals	1.0004	(<1)	0.996 (<1)
Ever worked making rubber or electrical cables	1.0001	(<1)	0.998 (<1)
Ever worked with leather	1.0002	(0)	0.999 (<1)
Ever worked with paint	1.0002	(0)	0.998 (<1)
Worked with paint in the last year	1.0002	(0)	0.998 (<1)
Worked with paint in the last fortnight	1.0001	(<1)	1.000 (<1)
Ever worked as a truck driver	0.9998	(<1)	0.999 (<1)
Worked as a truck driver in the last year	1.0002	(0)	0.999 (<1)
Worked as a truck driver in the last fortnight	1.0002	(0)	0.999 (<1)
Ever worked in an aluminium smelter	1.0003	(0)	0.999 (0)
Ever used dyes in hobbies	1.0002	(<1)	0.995 (<1)
Used dyes in hobbies in the last year	1.0003	(0)	0.998 (<1)
Ever worked with leather as part of a hobby	1.0002	(<1)	0.995  (<1)
Worked with leather as part of a hobby in the last year	1.0000	(<1)	0.997 (<1)
Ever used paint in a hobby	1.0002	(0)	0.998 (<1)
Used paint in a hobby in the last year	1.0002	(0)	1.001 (<1)
Used paint in a hobby in the last fortnight	1.0002	(<1)	0.998 (<1)
Ever used permanent hair dyes	1.0002	(0)	1.000 (<1)
Used permanent hair dyes in the last year	1.0003	(0)	0.998 (<1)
Ever used semipermanent hair dyes	1.0001	(<1)	1.000 (<1)
Used semipermanent hair dyes in the last year	1.0000	(<1)	1.001 (<1)
Ever used hair rinses	1.0003	(<1)	0.999 (0)
Ever smoked tobacco	1.0006	(<1)	0.997 (<1)
Passive smoking home	1.0000	(<1)	1.000 (<1)
Passive smoking workplace	1.0002	(0)	0.998 (<1)
Desire of Leaving			
During study period:	1 0001	( <1)	0.000 (<1)
Used dyes at workplace	1.0001	(<1)	0.998 (<1)
Used chemicals in workplace	1.0002	(0)	0.998 (<1)
Used paint in the workplace	1.0000	(<1)	1.000 (<1)
Worked as a truck driver	1.0002	(0)	0.999 (<1)
Used paint in a hobby	1.0003	(<1)	1.000 (<1)
Swimming in chlorinated water	1.0003	(0)	1.003 (<1)
Smoke tobacco	1.0005	(<1)	0.998 (<1)
Passive smoking home	1.0002	(0)	0.999 (<1)
Passive smoking workplace	1.0003	(0)	0.998 (<1)
Vitamin B12	1.0001	(<1)	0.998 (<1)
Folate	1.0001	(<1)	1.000 (<1)
Age	1.0003	(0)	0.995 (<1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of chloroform and bromoform (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome

Appendix 20: Relative risks in the assessment of confounding in the association between internal dose of chloroform, bromodichloromethane, and bromoform, and DNA index from flow cytometry

	Relative risk	(% change)	Relative risk	(% change)	Relative risk	(% change
	chloroform		Bromodichloro-methan	ie	Bromoform	
Baseline	1.04		1.11		1.04	
Bladder infection prior to 12 months	1.05	(1.28)	1.09	(1.70)	1.04	(0.13)
Kidney infection prior to 12 months	1.02	(2.13)	1.07	(4.02)	1.05	(1.01)
Family history of bladder cancer	1.06	(1.79)	1.01	(9.04)	1.03	(0.94)
Ever worked with dyes	1.13	(8.45)	1.08	(2.39)	1.03	(0.77)
Worked with dyes in the last year	1.02	(1.63)	1.10	(1.21)	1.05	(1.30)
Ever worked with chemicals	0.96	(8.13)	1.15	(3.42)	1.04	(0.17)
Ever worked making rubber or electrical cables	1.01	(2.57)	1.08	(2.66)	1.03	(1.21)
Ever worked with leather	1.04	(0.40)	1.11	(0.04)	1.04	(0.44)
Ever worked with paint	1.03	(0.86)	1.12	(0.75)	1.04	(0.18)
Worked with paint in the last year	1.05	(1.12)	1.09	(2.14)	1.05	(0.69)
Worked with paint in the last fortnight	1.04	(0.02)	1.09	(1.89)	1.04	(0.01)
Ever worked as a truck driver	0.98	(5.70)	1.23	(10.91)	1.00	(3.60)
Worked as a truck driver in the last year	1.04	(0.36)	1.11	(0.08)	1.04	(0.39)
Worked as a truck driver in the last fortnight	1.04	(0.42)	1.11	(0.12)	1.04	(0.36)
Ever used dyes in hobbies	1.20	(14.98)	1.33	(19.69)	1.01	(3.36)
Ever worked with leather as part of a hobby	1.02	(1.91)	1.33	(19.72)	0.99	(4.76)
Ever used paint in a hobby	1.01	(2.52)	1.15	(3.91)	1.04	(0.42)
Used paint in a hobby in the last year	1.00	(3.39)	1.13	(1.42)	1.04	(0.38)
Used paint in a hobby in the last fortnight	1.10	(5.65)	1.09	(2.22)	1.02	(2.13)

Cont'd

	Relative risk chloroform	(% change)	Relative risk Bromodichloro- methane	(% change)	Relative risk Bromoform	(% change)
Ever used permanent hair dyes	1.03	(0.93)	1.11	(0.44)	1.03	(0.85)
Ever used semipermanent hair dyes	1.05	(1.42)	1.14	(2.93)	1.02	(1.56)
Ever used hair rinses	1.04	(0.08)	1.09	(1.80)	1.03	(1.03)
Ever smoked tobacco	1.06	(2.37)	1.18	(6.51)	1.05	(0.68)
Passive smoking home	1.00	(3.85)	1.00	(9.91)	1.00	(3.85)
Passive smoking workplace	1.05	(0.82)	1.10	(0.87)	1.03	(0.75)
During study period:						
Used dyes at workplace	1.04	(0.39)	1.09	(1.47)	1.03	(1.01)
Used chemicals in workplace	1.03	(0.52)	1.11	(0.00)	1.05	(1.22)
Used paint in the workplace	1.04	(0.09)	1.10	(1.32)	1.03	(0.52)
Worked as a truck driver	1.04	(0.39)	1.11	(0.29)	1.04	(0.31)
Used paint in a hobby	1.06	(1.57)	1.12	(1.24)	1.04	(0.01)
Swimming in chlorinated water	1.06	(2.38)	1.18	(6.55)	1.05	(1.34)
Smoke tobacco	1.01	(2.80)	1.13	(1.71)	1.05	(0.76)
Passive smoking home	1.04	(0.36)	1.11	(0.07)	1.04	(0.40)
Passive smoking workplace	1.07	(2.49)	1.10	(0.99)	1.03	(1.10)
Vitamin B12	1.09	(4.83)	1.15	(3.31)	1.04	(0.34)
Folate	1.03	(1.40)	1.18	(6.42)	1.02	(2.37)
Age	1.04	(0.10)	1.10	(0.62)	1.03	(0.90)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = internal dose for chloroform, bromodichloromethane, and bromoform (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome.
- Highlighted are variable that change the exposure-outcome association by five percent of more, therefore included as a confounder in the final model

Appendix 21: Relative risks in the assessment of confounding in the association between available dose of total THM and DNA index from flow cytometry

(n=112)	Relative risk Total THM	(% change)
Baseline	1.004	
Bladder infection prior to 12 months	1.004	(<1)
Kidney infection prior to 12 months	1.004	(<1)
Family history of bladder cancer	1.004	(0)
Ever worked with dyes	1.004	(<1)
Worked with dyes in the last year	1.004	(0)
Ever worked with chemicals	1.004	(<1)
Ever worked making rubber or electrical cables	1.004	(<1)
Ever worked with leather	1.004	(0)
Ever worked with paint	1.004	(<1)
Worked with paint in the last year	1.004	(<1)
Worked with paint in the last fortnight	1.004	(<1)
Ever worked as a truck driver	1.004	(<1)
Worked as a truck driver in the last year	1.004	(<1)
Worked as a truck driver in the last fortnight	1.004	(<1)
Ever used dyes in hobbies	1.004	(<1)
Ever worked with leather as part of a hobby	1.003	(<1)
Ever used paint in a hobby	1.004	(<1)
Used paint in a hobby in the last year	1.004	(<1)
Used paint in a hobby in the last fortnight	1.004	(<1)
Ever used permanent hair dyes	1.004	(<1)
Ever used semipermanent hair dyes	1.004	(0)
Ever used hair rinses	1.004	(<1)
Ever smoked tobacco	1.005	(<1)
Passive smoking home	1.004	(<1)
Passive smoking workplace	1.004	(<1)
During study period:		
Used dyes at workplace	1.005	(<1)
Used chemicals in workplace	1.004	(0)
Used paint in the workplace	1.004	(<1)
Worked as a truck driver	1.004	(<1)
Used paint in a hobby	1.004	(<1)
Swimming in chlorinated water	1.004	(0)
Smoke tobacco	1.004	(<1)
Passive smoking workplace	1.004	(<1)
Vitamin B12	1.004	(0)
Folate	1.004	(<1)
Age	1.004	(<1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of total THM (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome

Appendix 22: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by diary, and DNA index from flow cytometry

n=112	Relative risk	(% change)
11 112	total THM	(70 change)
Baseline	1.0012	
Bladder infection prior to 12 months	1.0012	(0)
Kidney infection prior to 12 months	1.0013	(0)
Family history of bladder cancer	1.0012	(o) (<1)
Ever worked with dyes	1.0011	(0)
Worked with dyes in the last year	1.0012	(o) (<1)
Ever worked with chemicals	1.0013	(0)
Ever worked with chemicals  Ever worked making rubber or electrical cables	1.0012	(o) (<1)
Ever worked with leather	1.0011	(0)
Ever worked with paint	1.0012	(0)
Worked with paint worked with paint worked with paint in the last year	1.0012	(o) (<1)
Worked with paint in the last fortnight	1.0010	(<1) (<1)
Ever worked as a truck driver	1.0010	(<1) (<1)
Worked as a truck driver in the last year	1.0011	(0)
<u> </u>	1.0012	
Worked as a truck driver in the last fortnight Ever worked in an aluminium smelter	1.0012	(0)
Ever used dyes in hobbies	1.0013	(0) (0)
Used dyes in hobbies in the last year	1.0013	(o) (<1)
Ever worked with leather as part of a hobby	1.0013	(<1) (<1)
Worked with leather as part of a hobby in the last	1.0010	(<1) (<1)
Ever used paint in a hobby	1.0013	(<1) (<1)
Used paint in a hobby in the last year	1.0013	(0)
Used paint in a hobby in the last year Used paint in a hobby in the last fortnight	1.0013	(0)
Ever used permanent hair dyes	1.0012	(0)
Used permanent hair dyes in the last year	1.0012	(o) (<1)
Ever used semi-permanent hair dyes	1.0013	(<1) (<1)
Used semi-permanent hair dyes in the last year	1.0011	(<1) (<1)
Ever used hair rinses	1.0011	(<1) (<1)
Ever smoked tobacco	1.0011	(0)
Passive smoking home	1.0012	(0)
Passive smoking workplace	1.0012	(0)
Used dyes at workplace	1.0012	(<1)
Used chemicals in workplace	1.0013	(<1)
Used paint in the workplace	1.0019	(<1)
Worked as a truck driver	1.0010	(0)
Used paint in a hobby	1.0012	(0)
Swimming in chlorinated water	1.0012	(<1)
Smoke tobacco	1.0013	(<1)
Passive smoking - home	1.0013	(0)
Passive smoking - workplace	1.0012	(0)
Vitamin B12	1.0012	(0)
Folate	1.0012	(<1)
Age	1.0012	(<1)
1150	1.0011	( ^1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of total THM (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome

Appendix 23: Relative risks in the assessment of confounding in the association between intake dose of total THM as estimated by questionnaire, and DNA index from flow cytometry

n=112	Relative risk total	% change
	THM	C
Baseline	1.0001	
Bladder infection prior to 12 months	1.0001	(<1)
Kidney infection prior to 12 months	0.9999	(<1)
Family history of bladder cancer	1.0001	(<1)
Ever worked with dyes	0.9999	(<1)
Worked with dyes in the last year	1.0001	(0)
Ever worked with chemicals	1.0001	(0)
Ever worked making rubber or electrical cables	0.9999	(<1)
Ever worked with leather	1.0001	(0)
Ever worked with paint	1.0001	(0)
Worked with paint in the last year	1.0000	(<1)
Worked with paint in the last fortnight	1.0000	(0)
Ever worked as a truck driver	0.9998	(<1)
Worked as a truck driver in the last year	1.0001	(0)
Worked as a truck driver in the last fortnight	1.0001	(0)
Ever worked in an aluminium smelter	1.0001	(0)
Ever used dyes in hobbies	0.9999	(<1)
Used dyes in hobbies in the last year	1.0001	(0)
Ever worked with leather as part of a hobby	0.9999	(<1)
Worked with leather as part of a hobby in the last year	0.9999	(<1)
Ever used paint in a hobby	1.0000	(0)
Used paint in a hobby in the last year	1.0001	(<1)
Used paint in a hobby in the last fortnight	1.0000	(<1)
Ever used permanent hair dyes	1.0001	(0)
Used permanent hair dyes in the last year	1.0000	(0)
Ever used semi-permanent hair dyes	1.0000	(<1)
Used semi-permanent hair dyes in the last year	1.0000	(<1)
Ever used hair rinses	1.0001	(<1)
Ever smoked tobacco	1.0003	(<1)
Passive smoking home	1.0001	(0)
Passive smoking workplace	1.0001	(0)
During study period:		
Used dyes at workplace	1.0000	(<1)
Used chemicals in workplace	1.0000	(0)
Used paint in the workplace	0.9999	(<1)
Worked as a truck driver	1.0001	(0)
Used paint in a hobby	1.0001	(<1)
Swimming in chlorinated water	1.0002	(<1)
Smoke tobacco	1.0002	(<1) (<1)
Passive smoking - home	1.0002	(0)
Passive smoking - nome Passive smoking - workplace	1.0001	(0)
Vitamin B12	1.0001	(o) (<1)
Folate	1.0000	(0)
Age	1.0000	(<1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of total THM (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model i.e. baseline model + listed variable + outcome

Appendix 24: Relative risks in the assessment of confounding in the association between internal dose of total THM and DNA index from flow cytometry

(n=112)	Relative risk	(% change)
	Total THM	
Baseline	1.03	
Interaction term 1 (table 5-1)	1.02	(<1)
Bladder infection prior to 12 months	1.03	(<1)
Kidney infection prior to 12 months	1.03	(<1)
Family history of bladder cancer	1.01	(1.68)
Ever worked with dyes	1.03	(<1)
Worked with dyes in the last year	1.03	(<1)
Ever worked with chemicals	1.03	(<1)
Ever worked making rubber or electrical cables	1.02	(<1)
Ever worked with leather	1.03	(<1)
Ever worked with paint	1.03	(<1)
Worked with paint in the last year	1.03	(<1)
Worked with paint in the last fortnight	1.03	(<1)
Ever worked as a truck driver	1.02	(<1)
Worked as a truck driver in the last year	1.03	(<1)
Worked as a truck driver in the last fortnight	1.03	(<1)
Ever used dyes in hobbies	1.06	(<1)
Ever worked with leather as part of a hobby	1.03	(<1)
Ever used paint in a hobby	1.03	(<1)
Used paint in a hobby in the last year	1.03	(<1)
Used paint in a hobby in the last fortnight	1.02	(<1)
Ever used permanent hair dyes	1.02	(<1)
Ever used semipermanent hiar dyes	1.02	(<1)
Ever used hair rinses	1.02	(<1)
Ever smoked tobacco	1.05	(1.94)
Passive smoking home	1.03	(<1)
Passive smoking workplace	1.02	(<1)
During study period:		
Used dyes at workplace	1.02	(<1)
Used chemicals in workplace	1.03	(<1)
Used paint in the workplace	1.03	(<1)
Worked as a truck driver	1.03	(<1)
Used paint in a hobby	1.03	(<1)
Swimming in chlorinated water	1.05	(1.95)
Smoke tobacco	1.03	(<1)
Passive smoking home	1.03	(<1)
Passive smoking workplace	1.02	(<1)
Vitamin B12	1.04	(1.30)
Folate	1.02	(<1)
Age	1.03	(<1)

- Variables with no events or less than five events reported are not listed here. These variables were not tested because the numbers were too small to detect a contribution towards the model
- Baseline model = intake dose of total THM (exposure), and DNA index from flow cytometry (outcome). Each subsequent line represents a multivariate model – i.e. baseline model + listed variable + outcome

## Appendix 25: NCEPH - Environmental Survey 1997

Study Director Dr Geetha Ranmuthugala - 6249 5603 or 019 443 879

# Environmental Health Questionnaire 1 EHQ1

ID N	umber _					
Name	e					Telephone
The I	Home Vi	sitor was _				on
Urine	e 1 Yes 1	No	Urine	1 collectio	n	
Speci	ial Notes					
Date	EHQ1 S	cheduled _				
Cont	act Log	- To be co	mpleted	for each a	ittempt	at the number
		Start	Stop			
#	Date	Time	Time	Result	Int.	Comments
1						
2						
3						
4						
5						
6						
7						
8						
9						
10						
11						
12						
13						
14						

#### Introduction

15

Good evening, its ...(int name)... here from the Australian National University could I speak to ...(participant)... please.

Hello, ...(participant)... its ...(int name)... here from the Australian National University, how are you today. I'm ringing today to conduct the 1st telephone interview with you. As my colleague ...(HV name).. would have told you when she came to see you, this telephone interview is the 1st thing that we do in the study. Now this interview will take about 18 minutes to go through, may be a bit longer and maybe a bit shorter, it depends. Some of the questions will need you to remember back a bit so its best if we can make sure that you are comfortable and not likely to be interrupted for the next few minutes. Is now a good time to do this interview or would you like me to call back later tonight?

**If necessary,** It is important that we do this interview tonight as we want to move on to the first urine collection and to starting the fluid consumption diary tomorrow.

Now, during this interview I will read out each question to you exactly as it is written and then I'll read out the answer categories for you to choose from. You then choose the answer category which best answers the question and I will record that answer exactly. Take as much time as you like for each question and ask me to repeat things if I am not being clear. It's important that we get exact details on these questions, even ones that may seem unimportant to you so we want you to think carefully about each question.

#### **PART 1 Medical problems**

To begin with, I want to ask you about certain medical conditions and treatment that you may have had in the past.

Q1. First, have you had a bladder infection, also known as cystitis, that was diagnosed by a doctor in the last 12 months that is, any time from October last year through to now?

Yes	l
No	2
Cannot remember	3
Cannot remember	3

**Q2.** How many times did this occur in the past 12 months, that is since October 1996?

No. of times

Q3. Can you tell me when you had these bladder infections during the last 12 months, that is which months did they occur? Tick appropriate boxes. More than one response allowed

Oct '96 🖵	Feb '97 📮	Jun '97 🗖
Nov '96 <b>□</b>	Mar '97 📮	Jul '97 📮
Dec '96 □	April '97 🗖	Aug '97 <b>□</b>
Jan '97 🗖	May '97 📮	Sept '97□
		Oct '97 🗖

**Q4**. Prior to the last 12 months, have you ever had a bladder infection that was diagnosed by a doctor?

Q5. How many times did this occur?

No. of times

Q6. Can you tell me the year(s) in which you had bladder infections diagnosed by a doctor?

\_\_\_\_list years

Q7. Have you had a kidney infection diagnosed by a doctor in the last 12 months? (that is, from October '96 to October '97)

Yes 1	
No2	Go to Q10
Cannot remember 3	Go to Q10

Q8. How many times did this occur in the past 12 months, that is since October 1996?

No. of times
Q9. Can you tell me when you had these kidney infections *during the last 12 months* (that is in which months did they occur)? (Tick appropriate boxes. More than one response allowed) Tick appropriate boxes. More than one

response allowed

**Q10**. Prior to the last 12 months, have you *ever* had a kidney infection that was diagnosed by a doctor?

Q11. How many times did this occur?

\_\_\_\_No. of times

Q12. Can you tell me the year(s) in which you had a kidney infection diagnosed by a doctor?

list years

Q13. Have you had a bladder stone diagnosed by a doctor in the last 12 months? that is, since October last year, and here we are interested in bladder stones and not kidney stones?

Q14. How many times did this occur in the past 12 months, that is since October 1996?

No. of times

Q15. Can you tell me when you had these	//19
bladder stones during the last 12 months, that is which months did they occur? Tick appropriate boxes. More than one response allowed	(dd/mm/yy)  Q23. What was the reason or the diagnosis that led to the radiation treatment?
Oct '96  Feb '97  Jun '97    Nov '96  Mar '97  Jul '97    Dec '96  April '97  Aug '97    Jan '97  May '97  Sept '97    Oct '97    Oct '97     **Q16. Prior to the last 12 months, have you ever had a bladder stone diagnosed by a doctor?  Yes	Q24. Prior to the last 12 months, have you ever had radiation treatment?  Yes
Q18. Can you tell me the year(s) in which you had a bladder stone confirmed by a doctor?	No. of times  Q26. On those occasions did you receive radiation treatment? Time frame is all time prior to last 12 months.  to your head and or neck Yes No
Q19. Have you had radiation treatment in the last 12 months, that is, since October last year?  Yes	to your chest
radiation treatment in the past 12 months, that is since October 1996? No. of times	Q27. What was the approximate date of the last treatment? Time frame is all time prior to last 12 months.
Q21. On these occasions did you receive radiation treatment?  to your head and or neck	(dd/mm/yy)  Q28. What was the reason or the diagnosis that led to the radiation treatment? Time frame is all time prior to last 12 months

the last treatment?

<b>Q30a</b> . Have either of your parents ever	2
been diagnosed as having bladder cancer?	2
Yes1	3
No2	
30b. What about any of your brothers or	5
sisters?	6
Yes1	7 8.
No2	Q302. In how many separate years would
30c. Any of your children, have they ever	you have worked with dyes throughout
been diagnosed as having bladder cancer?	your life?
Yes1	your me.
No2	No. of years
30d. Any of your grandparents?	
Yes1	Q303. During that time did you make or
No2	use them, every day, every week, every month or less often?
30e. What about your aunts or uncles that	Daily 1 Goto 304a
is blood aunts or uncles, brothers or sisters	Weekly2 Goto 304b
to your parents? Yes1	Monthly 3 Goto 304c
No2	Less often 4 Goto 304d
	304a. On average how many hrs per day
30f. What about their children, your first	did you make or use them?
cousins, have any of them ever been	hrs per day
diagnosed as having bladder cancer?	304b. On average how many hrs per
Yes1 No2	week did you make or use them?
	hrs per week
30g. Any other close family members,	304c. On average how many hrs per
have any of them ever been diagnosed as	month did you make or use them?
having bladder cancer?	hrs per month
Yes1	304d. On average how many hrs per year
No2	did you make or use them?
30h. Who was that?	hrs per year
	<b>Q305</b> . You have told me about your
PART 2 Job History	lifetime contact with dyes. Now I would
Next I'd like to discuss jobs, either full-	like to ask you specifically about the last
time or part-time occupations that you	12 months. Have you worked with dyes,
may have worked in at any time during	that is making or using dyes in the past 12
your life from when you first started	months?
working right up to and including now.	Yes
	No
<b>Q300</b> . Firstly, have you ever had a job	Don't know not aware3 <b>Go to Q310</b>
where you used or made dyes?	Cannot remember4 Go to Q310
Yes1	
No2 Go to Q313	
Don't know not aware3 Go to Q313	
Cannot remember4 Go to Q313	
Q301. What dyes were you or are you	
working with?	
<del>-</del>	•

Q306. In the last 12 months did you make or use dyes, every day, every week, every month or less often?  Daily1 Goto 307a  Weekly2 Goto 307b  Monthly3 Goto 307c  Less often.4 Goto 307d  307a. On average how many hrs per day did you make or use dyes?  hrs per day  307b. On average how many hrs per week did you make or use dyes?  hrs per week	a3. One average, have you been using this dye: daily weekly, monthly or less often?  Daily
307c. On average how many hrs per	
month did you make or use dyes?	a3. One average, have you been using this
hrs per month	dye: daily weekly, monthly or less often?
307d. On average how many hrs per year	Daily 1
did you make or use dyes?	Weekly2
hrs per year	Monthly 3
Q308. Now I would like to know if you	Less often 4
have had contact with dyes specifically in	c. Fuchsin?
the last two week period. Have you made	Yes1
or used dyes in the last 2 weeks?	No
Yes1	Don't know not aware .3 Go to Q312d
No2 Go to Q310	·
Q309. How many hours in total did you	c2. In how many separate years have you
spend using or making dyes in the last two	been making or using this dye?
weeks?	No. of years
total no. of hours	No. of years
Q310. When was the last time you made	a3. One average, have you been using this
or used dyes, please tell me the month and	dye: daily weekly, monthly or less often?
the year?	Daily 1
/19 (mm/yy)	Weekly2
``	Monthly 3
Q311. Can you tell me the year in which you first used or made dyes?	Less often4
-	d. Safranin T
19	Yes1
Q312. Have you ever worked with any of	No
the following dyes used in dye industries?	Don't know not aware .3 Go to Q313
a. Benzedine?	
Yes1	d2. In how many separate years have you
No	been making or using this dye?
Don't know not aware3 Go to Q312b	
a2. In how many separate years have you	No. of years
been making or using this dye?	
-	
No. of years	

a3. One average, have you been using this dye: daily weekly, monthly or less often?  Daily	No
Q313. Have you ever worked as a	often?
fisherman?	Daily 1 Goto 320a
Yes1	Weekly2 Goto 320b
No	Monthly 3 Goto 320c
Q314. Did you use bronze dyes to stain	Less often4 Goto 320d
bait? Yes1 No	Q320a. Again, in the last 12 months, on average how many hrs per day did you work with bronze dyes?
Q315. In how many years did you work	hrs per day
with bronze dyes to stain bait, throughout your lifetime?	Q320b. Again, in the last 12 months, on average how many hrs per week did you work with bronze dyes?
No. of years	hrs per week
Q316. During that time, did you work with bronze dyes daily weekly, monthly or less often?	Q320c. Again, in the last 12 months, on average how many hrs per month did you work with bronze dyes? hrs per month
Daily1 Goto 317a	Q320d. Again, in the last 12 months, on
Weekly2 Goto 317b	average how many hrs per year did you
Monthly3 Goto 317c	work with bronze dyes?
Less often.4 Goto 317d	hrs per year
Q317a. On average how many hrs per day did you work with bronze dyes? hrs per day	Q321. Now I would like to know if you have been working with bronze dyes specifically in the last two week period. Have you been working with bronze dyes in the last 2 weeks?
Q317b. On average how many hrs per	Yes1
week did you work with bronze dyes?	No2 Go to Q323
hrs per week	Q322. How many hours in total did you
Q317c. On average how many hrs per	spend working with bronze dyes in the
month did you work with bronze dyes?	last two weeks?
hrs per month	total no. of hours
Q317d. On average how many hrs per	Q323. When was the last time you worked
year did you work with bronze dyes?hrs per year	with bronze dyes? Please tell me the month and year.
Q318. You have told me about your	/19 (mm/yy)
lifetime contact with bronze dyes used to	Q324. Can you tell me the year in which
stain bait. Now I would like to ask you	you first worked with bronze dyes?
specifically about the last 12 months.	19
Have you worked with bronze dyes to	
stain bait in the past 12 months ie since November '96?	

Yes .....1

Q325. Have you ever worked making or	Q331b. On average how many hrs per
using chemicals?	week did you work making rubber or
Yes1	making electrical cables?
No	hrs per week
Don't know	Q331c. On average how many hrs per
	month did you work making rubber or
Q326. Were you making chemicals,	making electrical cables?
handling chemicals or involved in some other way	hrs per month
Making chemicals 1	Q331d. On average how many hrs per year did you work making rubber or
Handling chemicals2	making electrical cables?
Involved in some other way3	hrs per year
	Q332. You have told me about your
Q327. Can you list these chemicals in the	lifetime history of making rubber or
blue sheet titled "List of Chemicals" that	making electrical cables. Now I would
was given to you by our Explanatory	like to ask you specifically about the last
Visitor. Do you recall getting this sheet?	12 months. Have you been making rubber
Yes	or making electrical cables in the past 12
No2	months ie since November '96?
Can you please check you package and let	Yes
me know. We will send you one in the	No
mail. Please return the completed sheet with the diary and the residential sheet	Q333. In the last 12 months, that is since
when we collect Urine 2. We may need to	November 1996, did you work making rubber or making electrical cables: daily;
speak to you about these chemicals in	weekly; monthly; or less often?
more detail at a later date.	Daily1 Goto 334a
Q328. Have you ever worked making	Weekly2 Goto 334b
rubber or making electrical cables?	Monthly 3 Goto 334c
This does not include making goods out	Less often 4 Goto 334d
of already made rubber?	Q334a. Again, in the last 12 months, on
Yes	average how many hrs per day did you
	work making rubber or making electrical
Q329. In how many separate years were	cables?
you working making rubber or making	hrs per day
electrical cables, throughout your lifetime?	Q334b. Again, in the last 12 months, on
	average how many hrs per week did you
No. of years	work making rubber or making electrical cables?
Q330. During that time, did you make	hrs per week
rubber or make electrical cables: daily;	Q334c. Again, in the last 12 months, on
weekly; monthly; or less often? Daily1 Goto 331a	average how many hrs per month did you
Weekly2 Goto 331b	work making rubber or making electrical
Monthly3 Goto 331c	cables?
Less often.4 Goto 331d	hrs per month
Q331a. On average how many hrs per day	Q334d. Again, in the last 12 months, on
did you work making rubber or making	average how many hrs per year did you
electrical cables?	work making rubber or making electrical
hrs per day	cables? hrs per year
	Q335. Now I would like to know if you
	have been making rubber or making

electrical cables specifically in the last	Q342b. On average how many hrs per
two week period. Have you been making	week did you work with leather?
rubber or making electrical cables in the	hrs per week
last 2 weeks?	Q342c. On average how many hrs per
Yes1	month did you work with leather?
No	hrs per month
Q336. How many hours in total did you spend making rubber or making electrical	Q342d. On average how many hrs per year did you work with leather?
cables in the last two weeks?	hrs per year
total no. of hours	Q343. You have told me about your
Q337. When was the last time you worked	lifetime history of leather work. Now I
making rubber or making electrical	would like to ask you specifically about
cables? Please tell me the month and year.	the last 12 months. Have you worked with
/19 <b>(mm/yy)</b>	leather in the past 12 months ie since
Q324. Can you tell me the year in which	November '96?
you first started making rubber or making	Yes1
electrical cables?	No
19	Q344. In the last 12 months, that is since
Q339. Have you ever worked as a	November 1996, did you work with
leather worker? This includes making,	leather: daily; weekly; monthly; or less
cutting, finishing leather goods, shoe	often?
making or shoe repairing?	Daily 1 Goto 345a
Yes1	Weekly2 Goto 345b
No	Monthly 3 Goto 345c
Q339a. Were you (read out the following	Less often 4 Goto 345d
options)	Q345a. Again, in the last 12 months, on
making leather 1	average how many hrs per day did you
cutting leather	work with leather?
finishing leather goods	hrs per day
a shoe maker	Q345b. Again, in the last 12 months, on
a shoe repairer 5 involved in another activity of leather	average how many hrs per week did you work with leather?
work (specify ) 6	hrs per week
\ <b>1</b>	Q345c. Again, in the last 12 months, on
Q340. In how many separate years did	average how many hrs per month did you
you work as a leather worker, throughout	work with leather?
your lifetime?	hrs per month
No. of years	Q345d. Again, in the last 12 months, on
Q341. During that time, did you work	average how many hrs per year did you
with leather: daily; weekly; monthly; or	work with leather?
less often?	hrs per year
Daily1 Goto 342a	
Weekly2 Goto 342b	
Monthly3 Goto 342c	
Less often.4 Goto 342d	
Designation of Gold 5720	
Q342a. On average how many hrs per day	
did you work with leather?	
hrs per day	

Q346. Now I would like to know if you	Q354b. On average how many hrs per
have been working with leather	week did you work with paint?
specifically in the last two week period.	hrs per week
Have you been working with leather in the last 2 weeks?	Q354c. On average how many hrs per
Yes1	month did you work with paint?
No	hrs per month
Q347. How many hours in total did you	Q354d. On average how many hrs per year did you work with paint?
spend working with leather in the last two weeks?	hrs per year
total no. of hours	Q355. You have told me about your
Q348. When was the last time you worked	lifetime contact with paint. Now I would
with leather? Please tell me the month and	like to ask you specifically about the last
year.	12 months. Have you worked with paint in the past 12 months ie since November
/19 (mm/yy)	'96?
Q349. Can you tell me the year in which	Yes1
you first worked with leather?	No
19	Q356. In the last 12 months, that is since
Q350. Have you ever worked as a	November 1996, did you work with paint:
painter or in a job that regularly	daily; weekly; monthly; or less often?
involved painting?	Daily 1 Goto 357a
Yes1	Weekly2 Goto 357b
No	Monthly 3 Goto 357c
Q351. What types of paint did you use?	Less often 4 Goto 357d
	Q357a. Again, in the last 12 months, on
	average how many hrs per day did you work with paint?
	hrs per day
	Q357b. Again, in the last 12 months, on
	average how many hrs per week did you
	work with paint?
	hrs per week
	Q357c. Again, in the last 12 months, on
Q352. In how many separate years did	average how many hrs per month did you work with paint?
you work as a painter, throughout your	hrs per month
lifetime?	Q357d. Again, in the last 12 months, on
No of ware	average how many hrs per year did you
No. of years	work with paint?
Q353. During that time, did you work as	hrs per year
a painter: daily; weekly; monthly;or less	Q358. Now I would like to ask you
often?	specifically about the last two week
Daily1 Goto 354a	period. Have you been working with paint
Weekly2 Goto 354b	in the last 2 weeks?
Monthly3 Goto 354c	Yes
Less often.4 Goto 354d	No
Q354a. On average how many hrs per day did you work with paint?	
hrs per day	
o poi day	

Q359. How many nours in total did you	Q367. In the last 12 months, that is since
spend working with paint in the last two	November 1996, did you drive trucks:
weeks?	daily; weekly; monthly; or less often?
total no. of hours	Daily 1 Goto 368a
Q360. When was the last time you worked	Weekly2 Goto 368b
with paint? Please tell me the month and	
year.	Monthly 3 Goto 368c
•	Less often 4 Goto 368d
/19 (mm/yy)	Q368a. Again, in the last 12 months, on
Q361. Can you tell me the year in which	average how many hrs per day did you
you first worked with paint?	drive trucks?
19	hrs per day
Q362. Have you ever worked as a truck	Q368b. Again, in the last 12 months, on
driver. This includes long distance or	average how many hrs per week did you
around town truck driving?	drive trucks?
Yes1	
No	hrs per week
2 30 10 4373	Q368c. Again, in the last 12 months, on
Q363. In how many separate years did	average how many hrs per month did you
you work as a truck driver, throughout	drive trucks?
your lifetime?	hrs per month
	Q368d. Again, in the last 12 months, on
No. of years	average how many hrs per year did you
Q364. During that time, did you drive	work as a truck driver?
trucks: daily; weekly; monthly; or less	hrs per year
often?	Q369. Now I would like to ask you
	specifically about the last two week
Daily1 Goto 365a	period. Have you been working as a truck
Weekly2 Goto 365b	driver in the last 2 weeks?
Monthly3 Goto 365c	Yes1
Less often.4 Goto 365d	No
Q365a. On average how many hrs per day	
did you drive trucks?	Q370. How many hours in total did you
hrs per day	spend working driving trucks in the last
Q365b. On average how many hrs per	two weeks?
week did you drive trucks?	total no. of hours
-	Q371. When was the last time you worked
hrs per week	as a truck driver? Please tell me the month
Q365c. On average how many hrs per	and year.
month did you drive trucks?	/19 (mm/yy)
hrs per month	Q372. Can you tell me the year in which
Q365d. On average how many hrs per	you first as a truck driver?
year did you drive trucks?	19
hrs per year	
Q366. You have told me about your	Q373. Have you ever worked in an
lifetime history of your work as a truck	aluminium smelter? This does include
driver. Now I would like to ask you	working in the offices of an aluminium
specifically about the last 12 months.	smelter.
Have you worked as a truck driver in the	Yes1
past 12 months ie since November '96?	No
Yes 1	
1 (.5	

Q374. In how many separate years did average how many hrs per week did you you work in an aluminium smelter, work in an aluminium smelter? throughout your lifetime? hrs per week Q379c. Again, in the last 12 months, on No. of years average how many hrs per month did you Q375. During that time, did you work in work in an aluminium smelter? an aluminium smelter: daily; weekly; hrs per month monthly; or less often? Q379d. Again, in the last 12 months, on Daily ......1 Goto 376a average how many hrs per year did you Weekly.....2 Goto 376b work in an aluminium smelter? Monthly ...3 Goto 376c hrs per year Less often.4 Goto 376d Q380. Now I would like to know if you Q376a. On average how many hrs per day have been working in an aluminium did you work in an aluminium smelter? smelter specifically in the last two week hrs per day period. Have you been working in an aluminium smelter in the last 2 weeks? Q376b. On average how many hrs per week did you work in an aluminium Yes ......1 smelter? hrs per week Q381. How many hours in total did you Q376c. On average how many hrs per spend working in an aluminium smelter in month did you work in an aluminium the last two weeks? smelter? total no. of hours hrs per month Q382. When was the last time you worked in an aluminium smelter? Please tell me Q376d. On average how many hrs per year did you work in an aluminium the month and year. smelter?  $\__/19$  $\___$  (mm/yy) \_\_\_\_ hrs per year Q383. Can you tell me the year in which O377. You have told me about your you first started working in an aluminium lifetime work in an aluminium smelter. smelter? Now I would like to ask you specifically 19 about the last 12 months. Have you **Hobbies** worked in an aluminium smelter in the Q384. Next I would like to ask you past 12 months ie since November '96? about certain hobbies, past-times, work Yes .....1 around the house, or any other non-work related activities that you may Q378. In the last 12 months, that is since have been involved in. Have you ever November 1996, did you work in an made or used dyes, including the use of aluminium smelter: daily; weekly; dyes in hairdressing, in any of you monthly; or less often? hobbies? Daily ......1 Goto 379a Yes.....1 Weekly.....2 Goto 379b Monthly ...3 Goto 379c Q384a. What were these dyes? Less often.4 Goto 379d 1.\_\_\_\_ 2.\_\_\_\_ Q379a. Again, in the last 12 months, on 3.\_\_\_\_ average how many hrs per day did you 4.\_\_\_\_\_ work in an aluminium smelter? 5.\_\_\_\_ hrs per day

Q379b. Again, in the last 12 months, on

Q385. In how many separate years did	hrs per week
you use dyes in this hobby?	390c. On average how many hrs per
	month did you make or use dyes as part
No. of years	of your hobby?
Q386. During that time did you use dyes	hrs per month
as part of this hobby: every day; every	390d. On average how many hrs per year
week; every month; or less often?	did you make or use dyes as part of your
Daily1 Goto 387a	hobby?
Weekly2 Goto 387b	hrs per year
Monthly3 Goto 387c	Q391. Now I would like to ask you specifically about the last two weeks.
Less often.4 Goto 387d	Have you been using dyes as part of your
387a. On average how many hrs per day	hobby in the last two weeks.
did you use dyes as part of your hobby?	Yes1
hrs per day	No
387b. On average how many hrs per	Q392. How many hours in total did you
week did you use dyes as part of your	spend using dyes as part of your hobby in
hobby?  hrs per week	the last two weeks?
387c. On average how many hrs per	total no. of hours
month did you use dyes as part of your	Q393. When was the last time that you
hobby?	used dyes as part of your hobby? Please
hrs per month	tell me the month and year.
387d. On average how many hrs per year	/19(mm/yy)
did you use dyes as part of your hobby?	Q394. Can you tell me the year in which
hrs per year	you first used dyes as part of your hobby?  19
<b>Q388</b> . You have told me about your	Q394a. Have you used any of the
lifetime contact with dyes as part of your	following dyes in any of your hobbies?
hobby. Now I would like to ask you	a. Benzedine?
specifically about the last 12 months.	Yes1
Have you been using dyes as part of your	No2
hobby in the past 12 months ie since	Don't know not aware .3
November '96?	b. 2-napthylamine?
Yes 1 No 2 Go to Q393	Yes1
	No2
Q389. In the last 12 months, that is since November 1996, did you make or use	Don't know not aware .3
dyes as part of your hobby: every day;	c. Fuchsin?
every week; every month; or less often?	Yes1
Daily1 Goto 390a	No2 Don't know not aware .3
Weekly2 Goto 390b	
Monthly3 Goto 390c	d. Safranin T Yes1
Less often.4 Goto 390d	No2
390a. On average how many hrs per day	Don't know not aware .3
did you make or use dyes as part of your	Zon time, not arraid .
hobby?	
hrs per day	

390b. On average how many hrs per week did you make or use dyes as part of

your hobby?

Q100. Have you ever been fishing, as a hobby, using bronze dyes to stain bait?	bronze dyes to stain bait as part of your hobby?
Yes1	hrs per day
No	Q401b. Again, in the last 12 months, on
Q396. In how many separate years have	average how many hrs per week did you
you been fishing using bronze dyess to	use bronze dyes to stain bait as part of
stain bait?	your hobby?
	hrs per week
No. of years	Q401c. Again, in the last 12 months, on
Q397. During that time, did you use	average how many hrs per month did you
bronze dyes to stain bait: daily; weekly;	use bronze dyes to stain bait as part of
monthly; or less often?	your hobby?
Daily1 Goto 398a	hrs per month
Weekly2 Goto 398b	Q401d. Again, in the last 12 months, on
Monthly3 Goto 398c	average how many hrs per year did you
2	use bronze dyes to stain bait as part of
Less often.4 Goto 398d	your hobby?
Q398a. On average how many hrs per day	hrs per year
did you use bronze dyes to stain bait?	Q402. Now I would like to ask you
hrs per day	specifically about the last two weeks.
Q398b. On average how many hrs per	Have you been using bronze dyes to stain
week did you use bronze dyes to stain	bait as part of your hobby, in the last two
bait?	weeks.?
hrs per week	Yes1
Q398c. On average how many hrs per	No
month did you use bronze dyes to stain	Q403. How many hours in total did you
bait?	spend using bronze dyes to stain bait as
hrs per month	part of your hobby in the last two weeks? total no. of hours
Q398d. On average how many hrs per	
year did you use bronze dyes to stain bait?	Q404. When was the last time that you
hrs per year	used bronze dyes to stain bait as part of your hobby? Please tell me the month and
Q399. You have told me about your	year.
lifetime use of bronze dyes as part of your hobby. Now I would like to ask you	
specifically about the last 12 months.	/19(mm/yy)
Have you been using bronze dyes to stain	Q405. Can you tell me the year in which
bait as part of your hobby in the past 12	you first used bronze dyes to stain bait as
months ie since November '96?	part of your hobby?
Yes1	19
No	Q406. Have you ever made or handled
Q400. In the last 12 months, that is since	chemicals in any of your hobbies?
November 1996, did you use bronze dyes	Yes 1 No 2 Go to Q407
to stain bait as part of your hobby: daily;	
weekly; monthly; or less often?	Q406a. What were these chemicals?
Daily1 Goto 401a	1
Weekly2 Goto 401b	2
Monthly3 Goto 401c	3
Less often.4 Goto 401d	4Q406b. Were you making chemicals,
	handling chemicals or involved in some
Q401a. Again, in the last 12 months, on	nananng chemicais of myorved in some

average how many hrs per day did you use other way

Making chemicals 1	Have you been making rubber or making
Handling chemicals2	electrical cables as part of your hobby in
Involved in another	the past 12 months ie since November
process with chemicals3	'96?
Can you list these chemicals in the blue	Yes1
sheet titled "List of Chemicals". We may	No
need to speak to you about these	Q412. In the last 12 months, that is since
chemicals in more detail at a later date.	November 1996, did you make rubber or
Q407. Have you ever made rubber or	make electrical cables as part of your
made electrical cables as a hobby? This	hobby: daily; weekly; monthly; or less
does not include making things using	often?
rubber.	Daily 1 Goto 413a
Yes1	Weekly2 Goto 413b
No	Monthly 3 Goto 413c
Q408. In how many separate years did	Less often 4 Goto 413d
you make rubber or make electrical cables	Q413a. Again, in the last 12 months, on
as part of your hobby?	average how many hrs per day did you
	spend making rubber or making electrical
No. of years	cables as part of your hobby?
Q409. During that time, did you make	hrs per day
rubber or make electrical cables: daily;	Q413b. Again, in the last 12 months, on
weekly; monthly; or less often?	average how many hrs per week did you
Daily1 Goto 410a	spend making rubber or making electrical
Weekly2 Goto 410b	cables as part of your hobby?
-	hrs per week
Monthly 3 Goto 410C	
Monthly3 Goto 410c Less often 4 Goto 410d	Q413c. Again, in the last 12 months, on
Less often.4 Goto 410d	average how many hrs per month did you
Less often.4 Goto 410d Q410a. On average how many hrs per day	average how many hrs per month did you spend making rubber or making electrical
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby?
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?hrs per day	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q413d. Again, in the last 12 months, on
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q413d. Again, in the last 12 months, on average how many hrs per year did you
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?hrs per day	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?hrs per day Q410b. On average how many hrs per week did you spend making rubber or	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables?
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?hrs per week	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year Q414. Now I would like to ask you specifically about the last two weeks.
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?  hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?  hrs per week Q410c. On average how many hrs per	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks.  Have you been making rubber or making
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?  hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?  hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby?	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year Q414. Now I would like to ask you specifically about the last two weeks.
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?  hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?  hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby?  hrs per month	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?  hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?  hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby?  hrs per month Q410d. On average how many hrs per	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby? hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q410d. On average how many hrs per year did you spend making rubber or	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?  Yes
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby? hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q410d. On average how many hrs per year did you spend making rubber or making electrical cables as part of your	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?  Yes
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?  hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?  hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby?  hrs per month Q410d. On average how many hrs per year did you spend making rubber or making electrical cables as part of your hobby?	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?  Yes
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby? hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q410d. On average how many hrs per year did you spend making rubber or making electrical cables as part of your hobby? hrs per weekhrs per year	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?  Yes
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby?  hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby?  hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby?  hrs per month Q410d. On average how many hrs per year did you spend making rubber or making electrical cables as part of your hobby?  hrs per wear Q411. You have told me about your	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?  Yes
Less often.4 Goto 410d Q410a. On average how many hrs per day did you spend making rubber or making electrical cables as part of your hobby? hrs per day Q410b. On average how many hrs per week did you spend making rubber or making electrical cables as part of your hobby? hrs per week Q410c. On average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month Q410d. On average how many hrs per year did you spend making rubber or making electrical cables as part of your hobby? hrs per weekhrs per year	average how many hrs per month did you spend making rubber or making electrical cables as part of your hobby? hrs per month  Q413d. Again, in the last 12 months, on average how many hrs per year did you spend making rubber or making electrical cables? hrs per year  Q414. Now I would like to ask you specifically about the last two weeks. Have you been making rubber or making electrical cables as part of your hobby in the last two weeks?  Yes

you specifically about the last 12 months.

Q415. How many nours in total aid you	Q421c. On average now many hrs per
spend making rubber or making electrical	month did you work with leather as part of
cables as part of your hobby in the last	your hobby?
two weeks?	hrs per month
total no. of hours	
	Q421d. On average how many hrs per
Q416. When was the last time that you	year did you work with leather as part of
made rubber or made electrical cables as	your hobby?
part of your hobby? Please tell me the	hrs per year
month and year.	Q422. You have told me about your
/19 (mm/yy)	- ·
	lifetime work with leather as part of your
Q417. Can you tell me the year in which	hobby. Now I would like to ask about the
you first made rubber or made electrical	last 12 months specifically. Have you
cables as part of your hobby?	been working with leather as part of your
19	hobby in the past 12 months ie since
Q418. Have you ever worked with	November '96?
<u> </u>	Yes1
leather in any of your hobbies? This	No
includes making leather, cutting,	
finishing leather goods, shoe making or	Q423. In the last 12 months, that is since
shoe repairing.	November 1996, did you work with
Yes1	leather as part of your hobby: daily;
No2 Go to Q429	weekly; monthly; or less often?
	Daily 1 Goto 424a
Q418a. Were you (read out the following	Weekly2 Goto 424b
options)	
making leather 1	Monthly 3 Goto 424c
cutting leather	Less often 4 Goto 424d
finishing leather goods	Q424a. Again, in the last 12 months, on
a shoe maker 4	average how many hrs per day did you
a shoe repairer 5	work with leather as part of your hobby?
involved in another process of leather	
work (specify) 6	hrs per day
work (specify) 0	Q424b. Again, in the last 12 months, on
Q419. In how many separate years did	average how many hrs per week did you
you work with leather as a hobby?	work with leather as part of your hobby?
	hrs per week
No. of years	Q424c. Again, in the last 12 months, on
0.420 Di 4144 1:1	average how many hrs per month did you
Q420. During that time, did you work	
with leather: daily; weekly; monthly; or	work with leather as part of your hobby?
less often?	hrs per month
Daily1 Goto 421a	Q424d. Again, in the last 12 months, on
Weekly2 Goto 421b	average how many hrs per year did you
Monthly3 Goto 421c	work with leather as part of your hobby?
Less often.4 Goto 421d	hrs per year
	Q425. Now I would like to ask you
Q421a. On average how many hrs per day	specifically about the last two weeks.
did you work with leather as part of your	Have you been working with leather as
hobby?	part of your hobby in the last 2 weeks?
hrs per day	
Q421b. On average how many hrs per	Yes1
week did you work with leather as part of	No
your hobby?	Q426. How many hours in total did you
-	spend working with leather, as part of
hrs per week	your hobby, in the last two weeks?

total no. of hours	Q432d. On average how many hrs per
Q427. When was the last time that you	year did you spend use paints as part of
worked with leather as part of your	your hobby?
hobby? Please tell me the month and year.	hrs per year
/19 <b>(mm/yy)</b>	Q433. You have told me about your
``	lifetime contact with paint as part of your
Q428. Can you tell me the year in which	hobby. Have you been using paints as part
you first worked with leather as part of	
your hobby?	of your hobby, in the past 12 months ie
19	since November '96?
<del></del>	Yes1
Q429. Have you ever used paint in any	No 2 Go to Q438
of your hobbies (Past-times, work	Q434. In the last 12 months, that is since
around the house etc)?	November 1996, did you use paints as part
Yes1	of your hobby: daily; weekly; monthly; or
No	less often?
Q429a. What were these paints?	
Q 125 m. What were those pulled.	Daily 1 Goto 435a
	Weekly2 Goto 435b
<del></del>	Monthly 3 Goto 435c
<del></del>	Less often4 Goto 435d
	Q435a. Again, in the last 12 months, on
	average how many hrs per day did you use
	paint as part of your hobby?
	hrs per day
	Q435b. Again, in the last 12 months, on
	average how many hrs per week did you
O420 In hory many concrete years did	use paint as part of your hobby?
Q430. In how many separate years did	hrs per week
you use paints in this hobby?	Q435c. Again, in the last 12 months, on
No. of years	average how many hrs per month did you
Q431. During that time, did you use	use paint as part of your hobby?
paints, as part of your hobby: daily;	hrs per month
weekly; monthly; or less often?	Q435d. Again, in the last 12 months, on
Daily1 Goto 432a	average how many hrs per year did you
-	use paint as part of your hobby?
Weekly2 Goto 432b	hrs per year
Monthly3 Goto 432c	Q436. Now I would like to ask you
Less often.4 Goto 432d	specifically about the last two weeks.
Q432a. On average how many hrs per day	Have you been using paint as part of your
did you spend using paint as part of your	hobby in the last 2 weeks?
hobby?	Yes1
hrs per day	No
Q432b. On average how many hrs per	Q437. How many hours in total did you
week did you use paints as part of your	spend using paints as part of your hobby
hobby?	in the last two weeks?
hrs per week	total no. of hours
Q432c. On average how many hrs per	Q438. When was the last time that you
month did you use paints as part of your	used paint as part of your hobby? Please
hobby?	tell me the month and year.
hrs per month	/10 (mm/m)

Q439. Can you tell me the year in which	No		
you first used with paint as part of your	Q126. During the last 12 months, how		
hobby?	many times in total did you use permanent		
19	hair dyes? (that is since November '96)		
Q440. Do you have any other hobbies	Total number of times		
on which you have spent more than 10	Q127. Have you used permanent hair dyes		
hours per week, for at least 1 year at	in the last two weeks?		
any stage in your life?	Yes1		
Yes	No		
·	Q128. How many times have you used		
Q441. Please name these hobbies and tell	permanent hair dyes in the last two		
me how many hours per week you spent	weeks?Total number of times		
on each of these hobbies	Q129. When was the last time that you		
Hobby Hrs / wk	used permanent hair dyes in the last two		
1	weeks?		
2	//19 (dd/mm/yy) Go to Q131		
3	Q130. When was the last time that you		
4	used permanent hair dyes? Please give the		
5,	year, if possible. If you have used		
6	permanent hair dyes in the last 12 months		
7 8.	(ie. since October '96), can you tell me		
	the month that you last used it?		
Hair tints and dyes	/19( <b>mm/yy</b> )		
I would now like to ask you about hair			
dyes or colour rinses that may have been	<b>Q131.</b> Have you ever used <b>semi- permanent hair dyes</b> , that is, hair dyes		
applied to your hair. This does not include	that wash out after about six washes?		
assisting others to apply hair dyes on themselves.	Yes1		
	No		
Q123. Have you <i>ever</i> used <b>permanent</b>			
hair dyes, that is, hair dyes that do not	Q132. In how many separate years did		
wash out but only grow out?	you use, or have you been using, semi-		
Yes1	permanent hair dyes?		
No	No. of years		
Q124. In how many separate years did	Q132a. In those years, on average how		
you use, or have you been using,	many months per year did you use semi-		
permanent hair dyes?	permanent hair dyes?		
No. of years	months per year		
Q124a. In those years, on average how	Q133. Have you used semi-permanent		
many months per year did you use	hair dyes in the past 12 months (ie. since		
permanent hair dyes?	October '96)?		
months per year	Yes1		
monaio poi you	No		
O125 Hove were and a second of the control of the c			
Q125. Have you used permanent hair			
dyes in the past 12 months (ie. since October '96)?			
Yes1			
1 651	l		

Q143. Have you used colour rinses in the
last two weeks?
Yes1
No2 Go to Q146
Q144. How many times have you used
colour rinses in the last two weeks?
Total number of times
Q145. When was the last time that you
used colour rinses in the last two weeks?
//19 (dd/mm/yy) Go to Water
Q146. When was the last time that you
used colour rinses? Please give the year, if
possible. If you have used colour rinses in
the last 12 months (ie. since October '96),
can you tell me the month that you last
used it?
/19 (mm/yy)

## Water History Code for portion siz

code for portion size: (for column /)	
Cup	. 1
Mug	. 2
Glass - small/pony	
Glass - medium/middy	
Glass - large/schooner	
Small bottle/can (285 ml)	. 6
Half bottle/375 ml stubbie/normal can	. 7
Large bottle (750 ml bottle of wine)	. 8
Half nip	9
Nip	0
Double nip	1
Other – specify	

Now I would like to ask you about what you have had to drink, including water, in the last 12 months (Interviewer – For each of the drinks listed in column 1 ask the following: Q1) Have you drunk ... in the last 12 months? If no, write no and skip to next drink. If yes, Q2) Would you have drunk ... every day, every week, every months or less often?

(1)	(2)	(3)	(4)	(5)	(6)	(7)
Type of beverage	NO -	YES-(daily) On	YES-(every week) On	YES (every month) On		What portion size do you
	did not		average, how many	average, how many	many drinks would you	usually have?
	drink	drinks would you	drinks would you have	drinks would you have	have had in the last 12	(use code above for portion
		have had per day?	had per week?	had per month?	months?	sizes)
Tap Water, as just water, from the community						1 2 3 4 5 6 7 8 9 10 11
water supply where you live						
Tap Water, as just water, from the community						1 2 3 4 5 6 7 8 9 10 11
water supply of your work place.						
Where do you work?						
Bottled Water						1 2 3 4 5 6 7 8 9 10 11
Tank Water						1 2 3 4 5 6 7 8 9 10 11
Well Water						1 2 3 4 5 6 7 8 9 10 11
Other Water 1 (specify)						1 2 3 4 5 6 7 8 9 10 11
Other Water 2 (specify)						1 2 3 4 5 6 7 8 9 10 11
Instant coffee						1 2 3 4 5 6 7 8 9 10 11
Brewed, expresso, filtered coffee						1 2 3 4 5 6 7 8 9 10 11
Decaf coffee						1 2 3 4 5 6 7 8 9 10 11
Hot chocolate, Milo, made with water not milk						1 2 3 4 5 6 7 8 9 10 11
Tea						1 2 3 4 5 6 7 8 9 10 11
Carbonated, fizzy drinks (non alcoholic)						1 2 3 4 5 6 7 8 9 10 11
Fruit juice – bought						1 2 3 4 5 6 7 8 9 10 11
Fruit juice - home-made						1 2 3 4 5 6 7 8 9 10 11
Cordial						1 2 3 4 5 6 7 8 9 10 11
Home Brews						1 2 3 4 5 6 7 8 9 10 11
Beer						1 2 3 4 5 6 7 8 9 10 11
Wine						1 2 3 4 5 6 7 8 9 10 11
Fortified wines (sherry, port)						1 2 3 4 5 6 7 8 9 10 11
Spirits specify if ice or water added						1 2 3 4 5 6 7 8 9 10 11
Is there any other type of drink that you have that I haven't mentioned (specify)						1 2 3 4 5 6 7 8 9 10 11

For ACT and SA only. (Bungendore – Go to smoking questions)	Q4. Have you ever stopped smoking tobacco for a more than 6 months at a
Q8. If you drink water from the tap, do	time?
you currently use a chlorine filter for your	Yes1
drinking water?	No2 Go to Q7
Yes	Q5. When was the last time that you stopped smoking tobacco for at least 6 months? Please tell me the month and year.
Do not use tap water	/19 (mm/yy)
for drinking purposes4 Go to Q10 Q9. How many years have you used a chlorine filter for your drinking water?	Q5a. When you stopped smoking that time, when did you start smoking again? Please tell me the month and year.
Number of years	/19 <b>(mm/yy)</b>
Number of months if less than one year	Q6. Not counting the times that you stopped smoking for a at least 6 months,
Q10. What is the <i>main</i> source of water that you use for showering and bathing?	how many years in total have you been smoking tobacco?
Tap water from the community supply 1	Number of years
Household tank water 2 Go to smoking A household well 3 Go to smoking	Number of months if less than one year
Other (specify)	Q7. Over the last few years, what type of
4 Go to smoking	tobacco did you mainly smoke? (one
Q11. Do you use a chlorine filter for	option only)
water that you use for showering and	Cigarettes, filtered
bathing?	Cigarettes, unfiltered2
Yes 1	Cigars
No, never used filter . 2 Go to smoking	
Used filter in past,	Q8. At present, what type of tobacco are
but not at present 3	you <i>mainly</i> smoking? (one option only)
Q12. How many years have you used a	Cigarettes, filtered
chlorine filter for your domestic water	Cigarettes, unfiltered2
source?	Cigars3
source:	Pipes4
Number of years	Q9. At present, on average, how many of
	the following do you smoke a day? (give
Number of months if less than one year	a number for each type of tobacco
Smoking	smoked)
Q1. Have you ever smoked tobacco?	No. / day
Yes1	Cigarettes, filtered
No	Cigarettes, unfiltered
Q2. Are you currently a tobacco smoker?	Cigars
Yes1	Pipes
No	Q10. Has this always been your pattern of
	smoking?
Q3. How old were you when you first	Yes
started smoking tobacco?	No2

Age in years

Q11. In the past, on average, how many of the following did you smoke a day? (give a number for each type of tobacco smoked)  No. / day  Cigarettes, filtered  Cigarettes, unfiltered  Pipes  Pipes  Q12. How old were you when you first stared smoking tobacco?	Q16. During the time that you were smoking tobacco, what type of tobacco did you <i>mainly</i> smoke?  Cigarettes, filtered
Age in years  Q14. How old were you when you last stopped smoking tobacco? Age in years  Q15. Not counting the times that you stopped smoking for periods greater than 6 months at a time, how many years in total were you smoking tobacco? Number of years	No. / day  Cigarettes, filtered  Cigarettes, unfiltered  Cigars  Pipes  Q18. Have you been smoking tobacco in the last 4 weeks?  Yes
Number of months if less than one year  Q20. In the last 4 weeks, what type of tobacco were you mainly smoking? (One option only)  Cigarettes, filtered	Number of days

## **Passive Smoking**

Q23. How many people in your household, apart from yourself, smoke inside the house? \_\_\_\_\_\_persons. (If zero go to next page)

Now I would like to ask you some questions about each of these inside smokers separately. If one of these inside smokers is your spouse or partner, please tell us about that person first.

	Person 1	Person 2	Person 3
Q24. Is this your spouse or partner? Ask for person 1 only)	Yes		
Q25. Does this person smoke while in the same room that you are in: every day; every week; every month; or less often?	Every day	Every day	Every day
Q26. In the last 4 weeks, how many smokes in total would he/she have smoked while in the same room as you?	smokes	smokes	smokes
Q27. In the last 4 weeks, has he/she been mainly smoking: filtered cigarettes; unfiltered cigarettes; cigars or a pipe?	Filtered cigarettes	Filtered cigarettes 1 Unfiltered cigarettes 2 Cigars 3 Pipes 4	Filtered cigarettes 1 Unfiltered cigarettes 2 Cigars 3 Pipes 4

Q28. In the last 4 weeks, have you been sharing an office, work room or other enclosed space, including vehicles, with a person who smoked tobacco inside that space?

Now I would like to ask you some questions about the persons apart from yourself who smoke tobacco inside that space.

	Person 1	Person 2	Person 3
Q29. Does this person smoke while in the same room that you are in: every day; every week; every month; or less often?	Every day	Every day	Every day
Q30. In the last 4 weeks, how many smokes in total would he/she have smoked while in the same room as you?	smokes	smokes	smokes
Q31. In the last 4 weeks, has he/she been mainly smoking: filtered cigarettes; unfiltered cigarettes; cigars or a pipe?	Filtered cigarettes1 Unfiltered cigarettes2 Cigars3 Pipes4	Filtered cigarettes 1 Unfiltered cigarettes 2 Cigars 3 Pipes 4	Filtered cigarettes 1 Unfiltered cigarettes 2 Cigars 3 Pipes 4

Q32. Next, I want to ask you about how many years in total of primary schooling and secondary schooling you completed. How many years of both primary and secondary schooling did you complete?

#### Years

Q33. And did you complete any further education or training after leaving secondary school?

Q34. What was the highest level of education that you have completed was it, a vocational certificate, a trade certificate, an associate diploma, a full diploma, a bachelor degree, a graduate diploma or a master or higher degree?

 Voc C.
 1

 Trad C.
 2

 Ass D.
 3

 Dipl.
 4

 Bach.
 5

 Grad D.
 6

 Mast.
 7

Q35. Thankyou, that is the end of the questions for now, before I go I want to remind you:

**If urine 1 is indicated.** To remember to give us a urine sample in the small bottle in the bag labelled 1. Now from my notes I see that you are going to leave it for us c

Also I want to remind you to start recording your fluid intake on the yellow fluid intake diary that ...(HV)... showed you. That starts when you first get up tomorrow morning. We also want you to record your showering and bathing outside ...(Respondent Location)... as per the yellow diary. We want you to keep the diary until we ring you again in exactly 14 days from now on .......

**If blood is to be done**, Also, some time in the next 14 days the Macquarie pathology lab will call you to arrange for your blood sample to be taken.

If you have any questions about procedure during the next 14 days please give us a

call on the number that is on top of your yellow appointment sheet list.

Thank you very much once again for assisting us in this medical research and we will speak to you again in 14 days.

#### Interviewer's comments

(to be completed at the end of each of interviews)

interviews)
1. Interview outcome
Completed1
Refusal2
Language problem3
Too ill4
Unable to complete5
Give reason
2. Was the participant responsive?
Not at all - uninterested, reticent 1
Fairly cooperative and responsive 2
Very cooperative, responsive 3
Very cooperative, responsive,
interested
3. Did the participant seem to remember
his/her medical history?
No1
Fairly well, some problems2
Very well3
Declined these questions4
4. Did the participant seem to remember
his/her occupation/hobbies history?
No1
Fairly well, some problems2
Very well 3
Declined these questions4
5. Did the participant seem to remember
his/her fluid consumption?
No1
Fairly well, some problems2
Very well3
Declined these questions4
6. Did the participant seem to remember
his/her smoking history?
No1
Fairly well, some problems2
Very well 3

Declined these questions......4

## Appendix 25a: NCEPH - Environmental Survey 1997

Study Director Dr Geetha Ranmuthugala - 6249 5603 or 019 443 879

## Environmental Health Questionnaire 2 EHQ2

ID N	ID Number					
Name						Telephone
Special Notes						
Date	EHQ2 S	cheduled _				
Cont	act Log			for each a	ıttempt	at the number
		Start	Stop			
#	Date	Time	Time	Result	Int.	Comments
1						
2						
3						
4						
5						
6						
7						
8						
9						
10						
11						
12						
13						
14						
1.5						

#### Introduction

Good evening, its ...(int name)... here from the Australian National University could I speak to ...(participant)... please.

Hello, ...(participant)... its ...(int name)... here from the Australian National University, how are you today. I'm ringing today to conduct the 2nd telephone interview with you.

Now this interview is shorter than the 1<sup>st</sup> one you did and will take about 12 minutes to go through. Can we do that now.?

Now, during this interview I will read out each question to you exactly as it is written and then I'll read out the answer categories for you to choose from. You then choose the answer category which best answers the question and I will record that answer exactly. Take as much time as you like for each question and ask me to repeat things if I am not being clear. It's important that we get exact details on these questions, even ones that may seem unimportant to you so we want you to think carefully about each question.

#### **PART 1 Medical problems**

I would like to ask you about some medical conditions and treatment that you may have had in the last 2 weeks (ie since your last telephone interview a fortnight ago).

Q1. Have you had a bladder infection	on, or
cystitis, diagnosed by a doctor during	ng the
last 2 weeks (ie since your last telep	hone
interview a fortnight ago)?	
Yes1	
No2 Go to	o Q4

Q2. Can you give me an approximate date that this diagnosis was made? (if unable to give a date, identify the week that the diagnoses was made)

	/19	_ (dd/mm/yy)
Q3. Are the	symptoms still pre	sent?
Yes	1	
No	2	

**Q4.** Have you had a kidney infection diagnosed by a doctor during the last 2 weeks (ie since your last telephone interview a fortnight ago)?

Yes	1	
No	2	Go to Q7

Q5. Can you give me an approximate date that this diagnosis was made? (if unable to give a date, identify the week that the diagnoses was made)

	//19(	(dd/mm/yy)
Q6. Are the	symptoms still prese	ent?
Yes	1	
No	2	

Q7. Have you had bladder stones (not kidney stones) diagnosed by a doctor during the last 2 weeks (ie since your last telephone interview a fortnight ago)?

Yes1	
No2	0 1 010

Q8. Can you give me an approximate date that this diagnosis was made? (if unable to give a date, identify the wee that the diagnoses was made)

	_//19_	(dd/mm/yy)
Q9. Are the sym	ptoms still	present?
Yes	1	
No	2	Go to Q4

Q10. Have you received radiation
treatment during the last 2 weeks (ie since
your last telephone interview a fortnight
ago) This does not include diagnostic x-
rays?

Yes	1	
No	2 Go to Q1	0
Q11. How many	times did you receive	
radiation treatm	ent during the last 2	
weeks?		

Q12. On those	occasions	did you	receive

No. of times

No

radiation treatment? to your head and or neck ...... Yes No to your chest ......Yes No to your abdomen ......Yes No

to your pelvis ......Yes No to your upper limbs ......Yes No to your lower limbs ...... Yes No to some other part of your

body ......Yes What part was that?

Q13. When was the last treatment date / /19 (dd/mm/yy)

### **PART 2 Job History**

Now I'd like to discuss jobs, either fulltime or part-time that you may have worked in during the last 2 weeks. (ie since your last telephone interview a fortnight ago)

Q1. Have you worked making or using dyes in the last 2 weeks?

Yes1	
No2	Go to Q3

Q2. How many hours in total did you spend using or making dyes in the last two weeks?

total no. of hours

Q3. Have you worked as a fisherman (as a job, not just for pleasure) in the last 2 weeks?

Yes1	
No2	Goto Q6

Q4. Did you use bronze or other dyes to	Q12. Were you (read out the following
stain bait?	options)
Yes1	making leather1
No	cutting leather2
Don't know	finishing leather goods
Cannot remember4 Goto Q6	a shoe maker4
Q5. How many hours in total did you	a shoe repairer5
spend working with bronze or other dyes	or another activity
used a bait in the last two weeks?	(specify)6
total no. of hours	Q13. In the last two weeks, how many
Q6. Have you worked making or	hours in total did you spend working with
handling chemicals in the last 2 weeks?	leather total no. of hours
Yes1	
No	Q14. Have you worked as a painter in
Q7. Please name these chemicals?	the last two weeks?
	Yes
	Q15. What were the types of paint that
	you used?
	you used?
Q8. How many hours in total did you	
spend making or handling these chemicals	
in the last two weeks?	
total no. of hours	
Q9. Have you worked making rubber	Q16. In the last two weeks, how many
or making electrical cables in the lagst	hours in total did you spend working with
2 weeks?	paint?
Yes1	total no. of hours
No 2 Go to Q11	Q17. Have you ever worked as a truck
Q10. How many hours in total did you	driver in the last two weeks. This
spend making rubber or making electrical	includes long distance or around town
cables in the last two weeks?	truck driving?
total no. of hours	Yes1
Q11. Have you worked as a leather	No
worker in the last 2 weeks? This	Q18. In the last two weeks, how many
includes leather cutting, finishing	hours in total did you work as a truck
leather goods, shoe making or shoe	driver?
repairing?	total no. of hours
Yes1	Q19. Have you ever worked in an
No2 Go to Q14	aluminium smelter in the last two
	weeks?.
	Yes1
	No

Q20. In the last two weeks, how many hours in total did you spend working in	
that job? total no. of hours	Q29. In the last two weeks, how many
Hobbies	hours in total would you have spent
I will now go through some of these	painting as a hobby?
activities again that you may have done	total no. of hours
as a hobby, past-time, as part of work	Q30. Are there any other hobbies that
around the house, or any other non-	you did for more than 10 hours per
· · · · · · · · · · · · · · · · · · ·	
work related activities, since your last	week, in the last two weeks?
telephone interview a fortnight ago.	Yes1
Q21. Have you made or used dyes as a	No
hobby in the last 2 weeks?	Q31. Please tell me what these hobbies
Yes1	were how many hours in total you spent
No2 Go to Q23	on each hobby during the last two weeks
Q22. How many hours in total would you	Hobby Hrs/wk
have spent working on this hobby during	1
the last 2 weeks?	2
total no. of hours	3
Q23. Have you fished, as a hobby, using	4
bronze dyes to stain bait, in the last two	5
weeks?	6
Yes1	7
No	8.
Q24. How many hours in total would you	
have spent using bronze dyes to stain bait	Hair tints and dyes
<del>-</del>	I would now like to ask you about any
during the last 2 weeks? total no. of hours	hair dyes you may have used in the last
<del></del>	two weeks.
Q25. Have worked with leather as a	Q32. In the last 2 weeks, have you used
hobby, this includes making leather,	permanent hair dyes, that is, hair dyes that
cutting, finishing leather goods, shoe	do not wash out but only grow out?
making or shoe repairing in the last 2	Yes1
weeks?	No
Yes1	Q33. How many times have you used
No	permanent hair dyes in the last 2 weeks?
Q26. How many hours in total would you	
have spent working on this hobby during	No. of times
the last 2 weeks?	Q34. When was the last time that you
total no. of hours	used permanent hair dyes in the last 2
Q27. Have been painting as a hobby in	weeks, please give me a date?
the last two weeks?	/ /19 (dd/mm/yy)
Yes1	Q35. Have you used semi-permanent hair
No2 Go to Q30	dyes in the last 2 weeks. Semi-permanent
Q28. What types of paint did you use?	hair dyes are those that wash out after
71 - r - r	about six washes?
	Yes1
	No
	Q36. How many times have you used
	semi-permanent hair dyes in the last 2
	weeks?

No. of times
Q37. When was the last time that you used semi-permanent hair dyes in the last 2 weeks, please give me a date?//19(dd/mm/yy)
Q38. Have you used colour rinses in the
last 2 weeks. Colour rinses are hair dyes that wash out the next time you wash your hair?
Yes1
No
Q36. How many times have you used colour rinses in the last 2 weeks?
No. of times
Q37. When was the last time that you used colour rinses in the last 2 weeks, please give me a date?
//19 (dd/mm/yy)

Water History
Code for portion size: (for column 6 of Q 38)
Cup1
Mug2
Glass - small/pony3
Glass - medium/middy4
Glass - large/schooner5
Small bottle/can (285 ml)6
Half bottle/375 ml stubbie/normal can7
Large bottle (750 ml bottle of wine)8
Half nip9
Nip10
Double nip11
Other – specify12

Q38. Now I would like to ask you about what you have had to drink, including water, in the last 2 weeks (Interviewer – For each of the drinks listed in column 1 ask the following: i) Have you drunk ... in the last 2 weeks? If no, write no and skip to next drink. If yes, ii) Would you have drunk ... every day, every week, or less often?

(1)	(2)	(3)	(4)	(5)	(6)
Type of beverage	NO -	YES-(daily) On			What portion size do you
	did not	average, how many	average, how many	many drinks would you	usually have?
	drink	drinks would you	drinks would you have		(use code above for portion
		have had per day?	had per week?	weeks?	sizes)
Tap Water, as just water, from the community					1 2 3 4 5 6 7 8 9 10 11
water supply where you live					
Tap Water, as just water, from the community					1 2 3 4 5 6 7 8 9 10 11
water supply of your work place.					
Where do you work?					
Bottled Water					1 2 3 4 5 6 7 8 9 10 11
Tank Water					1 2 3 4 5 6 7 8 9 10 11
Well Water					1 2 3 4 5 6 7 8 9 10 11
Other Water 1 (specify)					1 2 3 4 5 6 7 8 9 10 11
Other Water 2 (specify)					1 2 3 4 5 6 7 8 9 10 11
Instant coffee					1 2 3 4 5 6 7 8 9 10 11
Brewed, expresso, filtered coffee					1 2 3 4 5 6 7 8 9 10 11
Decaf coffee					1 2 3 4 5 6 7 8 9 10 11
Hot chocolate, Milo, made with water not milk					1 2 3 4 5 6 7 8 9 10 11
Tea					1 2 3 4 5 6 7 8 9 10 11
Carbonated, fizzy drinks (non alcoholic)					1 2 3 4 5 6 7 8 9 10 11
Fruit juice – bought					1 2 3 4 5 6 7 8 9 10 11
Fruit juice - home-made					1 2 3 4 5 6 7 8 9 10 11
Cordial					1 2 3 4 5 6 7 8 9 10 11
Home Brews					1 2 3 4 5 6 7 8 9 10 11
Beer					1 2 3 4 5 6 7 8 9 10 11
Wine					1 2 3 4 5 6 7 8 9 10 11
Fortified wines (sherry, port)					1 2 3 4 5 6 7 8 9 10 11
Spirits specify if ice or water added					1 2 3 4 5 6 7 8 9 10 11
Is there any other type of drink that you have that I haven't mentioned (specify)					1 2 3 4 5 6 7 8 9 10 11

Q39. Have you been swimming in the las
2 weeks ie since your first telephone
interview a fortnight ago?
Yes1
No
Q40. Did you swim in: chlorinated pools
only; un-chlorinated pools only; or both
cholrinated and un-chlorinated pools?
Chorinated pools only1
Un-chorinated pools only .2 Go to Q44
Both3
Uncertain4 Go to Q44
I will now ask you a few questions about
the times you swam in chlorinated pools
only. Do not include the times that you
swam in un-chlorinated pools, rivers,
dams, or lakes when responding to these
questions. If you are not sure whether the
pools were chlorinated or not, do not
include it as a swim in a chlorinated pool
•
Q41. How many times did you swim in a
chlorinated pool over the last 2 weeks.?
Number of years
Number of years
Q42. On average, how much time, in
minutes, did you sepnd in the water each
time?
Number of minutes on average
Q43. When was the last time that you
swam, please give me a date?
//19 (dd/mm/yy)
Smoking
Q44. Have you been smoking tobacco in
the last two weeks?
Yes1
No
Q45. Did you smoke tobacco: daily;
weekly less often than that?
Daily1 Go to Q45a
Weekly 2 Go to Q45b
Less often than that2 Go to Q45c
Q45a During the last two weeks, on
average, how many of the following did
you smoke a day? No. / day
Cigarettes, filtered
Cigarettes, unfiltered
Cigars
Pipes
-
Q45b. During the last two weeks, on
average, how many of the following did
you have per week?

No. / wk
Cigarettes, filtered
Cigarettes, unfiltered
Cigars
Pipes
Q45c. During the last two weeks, on
average, how many of the following
would you have had in the two week time
period?
No. in 2 wks
Cigarettes, filtered
Cigarettes, unfiltered
Cigars
Pipes
Q46 When was the last time you smoked
tobacco in the last two weeks, please give
me a date
//19 (dd/mm/yy)

Passiv	re Smoking
$\Omega$ 47 H	as anyone inside your home been smoking tobacco in the last 2 weeks?

Q47. Thas anyone inside your nome been smoking tobacco in the last 2 weeks?
Yes1
No
Now I would like to ask you about any persons who have been smoking inside the house in the last 2 weeks. Please tell me about each one of them
separately. If one of those inside smokers is your spouse or partner, please tell me about that person first.

	Person 1	Person 2	Person 3
Q48. Is this your spouse or partner? Ask for person 1 only)	Yes		
Q49. In the last 2 weeks, would the person have smoked in the same room that you were in: every day; every week; or less often?	Every day	Every day	Every day
Q50. In the last 2 weeks, how many smokes in total would he/she have smoked while in the same room as you?	smokes	smokes	smokes
Q51. In the last 2 weeks, has he/she been mainly smoking: filtered cigarettes; unfiltered cigarettes; cigars or a pipe?	Filtered cigarettes	Filtered cigarettes 1 Unfiltered cigarettes 2 Cigars	Filtered cigarettes

Q52. In the last 2 weeks, have you been sharing an office, work room or other enclosed space, including vehicles, with a person who smoked tobacco inside that space?

I will ask you some details about the persons apart from yourself who smoke tobacco inside that space.

	Person 1	Person 2	Person 3
Q53. In the last two weeks has			
the person been smoking while	Every day1	Every day 1	Every day 1
in the same room that you are in:	Every week2	Every week2	Every week2
every day; every week; or less	Less often3	Less often 3	Less often 3
often?			
Q54. In the last 2 weeks, how			
many smokes in total would			
he/she have smoked a day while	smokes	smokes	smokes
he/she have smoked a day while in the same room as you?	smokes	smokes	smokes
_	smokes	smokes	smokes
in the same room as you?	smokes Filtered cigarettes1	smokes Filtered cigarettes 1	smokes Filtered cigarettes 1
in the same room as you?  Q55. In the last 2 weeks, has			
in the same room as you?  Q55. In the last 2 weeks, has he/she been mainly smoking:	Filtered cigarettes1	Filtered cigarettes 1	Filtered cigarettes 1
in the same room as you?  Q55. In the last 2 weeks, has he/she been mainly smoking: filtered cigarettes; unfiltered	Filtered cigarettes1 Unfiltered cigarettes2	Filtered cigarettes 1 Unfiltered cigarettes 2	Filtered cigarettes 1 Unfiltered cigarettes 2

Q56. Do you regularly spend time in an enclosed space where there are one or more other inside smokers? For example, do you spend time, on a regular basis, at a	
pub where there are other inside smokers, or do you travel in a vehicle with a person who regularly smokes inside the vehicle?  Yes	Thank you very much for being part of this study and helping us with this community health research. We are very grateful to you.
Q57. Where does this take place?	For Bungendore respondents only.  We have one more interview to be done in another 14 days from today and that will be about the same length as today's and will have similar questions.  So we will talk to you next on
	For all respondents
Q58. How often are you in this enclosed space at the same time that at least one other person is smoking within the enclosed space. Is it daily; weekly; or less often?  Every day	Now tomorrow we want you to give us two urine samples, one from the second time you empty your bladder in the day and the other from the third time you empty your bladder. Those samples go in the bottles marked 2 and 3 on the caps. The second urine of the day goes in the bottle marked 2 and the third in the bottle marked 3. Write the time of the sample and the date on the labels then put the bottles in the plastic bags and then into the brown paper bag marked with the number 2.  Now we will collect that from you tomorrow, I have here that we should collect that from:
average hours per week Q61. In the last two weeks, how many	
hours in total did you spend in this enclosed area while at least one other person was smoking in the enclosed area? total hours	and that you expect to have both samples ready by (Time). Is that still OK? If not change details  Please remember to put your residential
Q62. Finally can I have you date of birth please? //19(dd/mm/yy)	history sheet and your fluid intake diary in the gold coloured envelope and put that inside the bag together with the sample
Q63. Would you like to make any comments on any aspect of the study?	bottles for us to collect. Have you finished the residential history sheet? ( <b>If not</b> , please do so before tomorrow).
	Urine 3 Yes1

## No......2-> Skip this section Now after you have done all that you should have another two sample bottles in a another brown paper bag marked 3. Now the bottles in this bag have got 2 and 3 on the caps just like the ones that you will use tomorrow but these are for the next day, that is day)... \_\_\_...(name of day)... We Again on want you to give us two urine samples one from the second time you empty your bladder in the day and the other from the third time you empty your bladder in the day. Again they go in the bottles marked 2 and 3 on the caps. Write the time of the sample and date on the labels then put the bottles into the plastic bags and then into the brown paper bag marked with the number 3. We will collect these from you ...(name of day)... I have here that we should collect that from: and that you expect to have both samples ready by **(Time)**. Is that still OK? Blood Yes.....1 No.....2-> Skip this section Have you had your blood test done? Yes .....1 No.....2-Have you got an ..... appointment made ..... for the blood test? Yes.....1 No .....2 ->We will get someone to call ..... you in the next 2 ..... days to arrange that.

If you have any questions about the procedure during the next 14 days please give us a call on the number that is on top of your appointment sheet.

The results of your blood test will be mailed out to you before Christmas.

## **Final Statement**

Once again, thank you very much for assisting us in this medical research. (and we will speak to you again in 14 days.)

#### Interviewer's comments

(to be completed at the end of each of interviews)

interviews)
1. Interview outcome
Completed1
Refusal2
Language problem3
Too ill4
Unable to complete5
Give reason
2. Was the participant responsive?
Not at all - uninterested, reticent 1
Fairly cooperative and responsive 2
Very cooperative, responsive 3
Very cooperative, responsive,
interested 4
3. Did the participant seem to remember
his/her medical history?
No1
Fairly well, some problems2
Very well3
Declined these questions4
4. Did the participant seem to remember
his/her occupation/hobbies history?
No1
Fairly well, some problems2
Very well3
Declined these questions4
5. Did the participant seem to remember
his/her fluid consumption?
No1
Fairly well, some problems2
Very well
Declined these questions4
6. Did the participant seem to remember
his/her smoking history?
No1
Fairly well, some problems2
Very well3
Declined these questions 4

ID number	

## **Appendix 26:**

# Diary of fluid intake and swimming & Diary of showering / bathing away from your place of residence

ID Number	
Start date	
End date	

Start this record the morning after telephone interview 1.

#### Record:

- All that you have to drink until you go to bed on the day of telephone interview 2.
- If you go swimming.
- If you have a shower, bath, sauna or spa away from your place of residence.

Please leave the completed diary and residential history for collection with 'urine 2'.

## Guidelines for keeping fluid intake record

Thank you for taking time to keep this diary. Following are guidelines to help you record your beverage consumption.

- 1. Start your beverage / fluid recording the morning after your first telephone interview. Starting that morning, record all that you have to drink for two weeks. You can stop recording when you go to bed the day you have your second telephone interview. Drink as you normally would during the two weeks.
- 2. Record everything that you have to drink in column 2. This includes water from the town water supply drunk as just water, bottled water, water from any other source (eg. well or tank), coffee (instant, brewed, decaf), tea, fruit juice, cordials, carbonated (fizzy) drinks, wine, beer, spirits, or anything else that you may have to drink. When ever possible, record the beverage soon after drinking. If this is not possible, enter each days record at the end of the day. It is important that you record all that you have had to drink at least on a daily basis.
- 3. Record the amount that you have had to drink in column 3. Use the guide given below to record the amount consumed. Record the number of serves in column 4. For example, if you have had *two medium glasses of orange juice*, record it as follows:

(1) Time	(2) TYPE of drink (eg town water, tank water, coffee, tea etc.)	(3) Amount	(4) Number of serves	(5) Place (town AND state)	Please use the following codes when filling in the amount of beverage consumed in column 2, and specify the number of serves.
10.30 am	orange juice	4)—	2 serves	Adelaide city, SA	Cup
					Glass - small/pony
					Small bottle/can (285 ml)
					Large bottle (750 ml bottle of wine)
					Nip         10           Double nip         11           Other (specify volume)         12

- 4. Please write down the place where you had this drink in column 5. For example, if you had a cup of coffee in Adelaide city, please write 'Adelaide city, SA' in column 5.
- 5. List only one beverage per line. Include a time for each drink in column 1.
- 6. Start a new page for each day. Circle the day (Sunday, Monday, Tuesday etc when you start each day), and enter the date.

Please do not hesitate to contact Geetha Ranmuthugala on 1 800 812 954 if you have problems. Your help is greatly appreciated.

Thank you.

ID number			

## Instructions for Swimming / Sauna / Spa record

If you go swimming (in Adelaide or elsewhere) in the time period between your first and second telephone interviews, please write down the place and duration of the swim. If you know whether the water was chlorinated or not, please write in this information. For example, if you went swimming in a private chlorinated pool on the 15th of October for 20 minutes, you would write the following (on the page that you record fluid intake for 15 October):

	Swimming (anywhere	•	Showering / bathing/ Sauna / Spa (outside your place of residence)			
Where (pool, sea, beach etc.)	Duration of swim (minutes)	Chlorinated / Unchlorinated/ Do not know	Where (town AND state)	Duration of shower/bath/spa /sauna (minutes)	Chlorinated/ Unchlorinated/ Do not know	
Private pool Adelaide	20 minutes	Chlorinated				

## Instructions for Showering / Bathing / Spa / Sauna

If you have a shower, bath, spa or sauna **away from your place of residence** in the time period between your first and second telephone interviews, please write down the place and duration of shower, bath, spa or sauna. Please indicate whether it was a shower, bath, spa or sauna. For example, if you had a shower in Port Adelaide, or if you used a spa in Sydney, you would write the following:

Swimming	Showeri	ng / bathing/ Sa	una / Spa	
(anywhere)		(outside	your place of r	esidence)
(pool, sea, swim Ur	nlorinated / nchlorinated/ o not know	Where (town AND state)	Duration of shower/bath/spa/ sauna (minutes)	Chlorinated/ Unchlorinated/ Do not know
		Port Adelaide, SA Sydney NSW	20 minutes shower 20 minutes spa	chlorinated chlorinated

See over for diary

Date							
Day (circle one):	Sun	Mon	Tues	Wed	Thurs	Fri	Sat

	Fluid intake						
(1)	(2)	(3)	(4)	(5)			
Time	TYPE of drink (eg	Amount	Number	Place (town			
	town water, tank water,		of serves	AND state)			
	coffee, tea, etc.)			ŕ			

## Codes for 'amount' in fluid intake diary (column 3)

Cup	
Mug	
Glass - small/pony	
Glass - medium/middy	
Glass - large/schooner	
Small bottle/can (285 ml)	
Half bottle/375 ml stubbie/normal can	
Large bottle (750 ml bottle of wine)	
Half nip.	
Nip	
Double nip	
Other (specify volume)	
(-r)	

## Special instructions for fluid type (column 2)

Please record <u>all</u> fluids consumed

If type = water - indicate
if 'boiled' or
straight off 'tap.'

If type = spirits, indicate if
water or ice added

Swimming (anywhere)			0	-
Duration of swim (minutes)	Chlorinated / Unchlorinated/ Do not know	Where (town AND state)	Duration of shower/bath/spa/ sauna (minutes)	Chlorinated/ Unchlorinated/ Do not know
	(anywhere Duration of swim	(anywhere)  Duration of Swim Chlorinated / Unchlorinated/	(anywhere)(outside)Duration of swimChlorinated / Unchlorinated / Unchlorinated / AND state)	(anywhere)(outside your place of red)Duration of swimChlorinated / SwimWhere (town AND state)Duration of Shower/bath/spa/

	ID number
Comments on fluid consumption	

## **Appendix 27: Introductory letter (tailored for ACT)**



20 October 1997

«Name» «Name» «Address» «Suburb», «State» «POST\_\_CODE»

Dear Mr. «NAME»

I am writing to you to ask for your help in a medical research project conducted by the National Centre for Epidemiology and Population Health (NCEPH), at the Australian National University.

We are studying the level of community exposure to substances in the environment that are believed to increase the risk of developing urinary bladder cancer. Exposure to certain chemicals and other substances in the work place and home environment may increase the risk of developing urinary bladder cancer. Tobacco smoking, alcohol consumption, and certain medical conditions are also thought to increase the risk of developing bladder cancer. This study will help examine a link between exposure to these potentially harmfully substances and the extent of bladder cell damage in the community. The degree of bladder cell damage is used as a biological marker of the communities potential to develop cancer. Health regulatory organisations depend on such research to set safety values for human exposure to potentially harmful substances. This study will therefore benefit the health of the community.

We are seeking your help as a resident of the ACT, and asking you, and all other eligible member of your household, to take part in this study. Your name was randomly selected from the white pages. Any person in your household who meets the following eligibility criteria can participate in the study:

- Male aged between 30 and 65 years;
- Resident in the ACT for a minimum period of six months;
- Not a regular swimmer in chlorinated swimming pools; and
- Never been diagnosed as having cancer. This does not include skin cancer.

NATIONAL CENTRE FOR EPIDEMIOLOGY AND POPULATION HEALTH
THE AUSTRALIAN NATIONAL UNIVERSITY
CANBERRA ACT 0200 AUSTRALIA

Telephone: (02) 6249 2378 Facsimile: (02) 6249 0740

Participation in this study involves the following.

- A visit by one of our researchers to explain the study further, and to go through the process with you.
- Participation in two telephone interviews, a fortnight a part. During these interviews, we will ask you about certain substances that you may have come in to contact with as part of your daily activities.
- Maintain a diary of fluid consumption for two weeks.
- Provide urine samples on two days, a fortnight apart
- Provide a blood sample for vitamin levels. This blood test can be done at your home, or at any one of the many Macquarie Pathology Services collection rooms in the ACT, at no cost to you.

I would be most grateful to you if you could find the time to help us with this study. Your participation is invaluable for the success of this project. Participation is voluntary and all information provided by you will be kept strictly confidential. You will not be identifiable from the reporting of the findings. Please take some time to read the enclosed brochure with answers to frequently asked questions.

An ANU researcher will call you in the next few days to speak to you about this project. If you are able to assist us, we would like to make an appointment to meet you at a convenient time. In the mean time, if you would like to discuss any aspect of the study, please do not hesitate to call me on 1 800 812 954.

Your assistance is greatly appreciated.

Thank you.

Yours sincerely,

Dr Geetha Ranmuthugala Project Coordinator

## **Appendix 28: Study brochure**

## **Environmental Study**

## **Answers to Frequently Asked Questions**

NATIONAL CENTRE FOR EPIDEMIOLOGY AND POPULATION HEALTH

THE AUSTRALIAN NATIONAL UNIVERSITY

## What is the purpose of this study?

There is evidence to indicate that exposure to certain substances in the environment can increase the risk of developing urinary bladder cancer. Many of these substances are often found in our work and home environment. Although research to date indicates an increased risk of developing bladder cancer following contact with these substances over many years, the evidence is still inconclusive and therefore merits further investigation.

People working in certain jobs are more likely to come in contact with certain substances that have been implicated in the development of bladder cancer. For others, exposure to these potentially harmful substances may be through water, tobacco smoking or alcohol consumption. We are asking the general population to tell us whether they have been exposed to these substances that are believed to be harmful to the bladder. This information will be supplemented with analysis of urine samples. The results of this research will help establish a link between the extent of contact at the community level to possible cancer causing agents and the risk of developing bladder cancer. Analysis will be undertaken at the community level rather than at the individual participant level. This study therefore will act as a first but very important step for undertaking further research on individuals considered to be at high risk of developing bladder cancer.

Furthermore, by measuring the degree of cell damage that results from contact with potential cancer causing substances, it is possible to identify the level of exposure beyond which contact is considered hazardous. Such information is invaluable to official regulatory authorities for reviewing existing guidelines and determining safe cut-off levels.

The results of this study will help us all - the community, the researchers and the officials - in developing a better understanding of the causes and risk of developing bladder cancer. The ultimate goal would be to improve the health of the community not only at the local and state level, but also nationally.

## Why am I getting a letter?

Your name was randomly selected from the Adelaide white pages. We are seeking the help of 150 individuals in Adelaide.

### Why should I participate?

We are studying the risk to the community as a whole. There is a lot of variation from person to person, and studying the community as a whole will help us to overcome variation between persons. For a community-based study to be successful, it is important that all people selected complete the study. Your participation is therefore essential in ensuring the results of the study are of value to you and to your community.

## **Should I have cancer to participate?**

No, we are not looking for people with any particular kind of health problem(s).

## How do you assess the risk of developing bladder cancer?

Cell damage, cell death and replacement of dead cells with healthy cells is a normal process for every living being. Even in a perfectly healthy human being, there is a baseline level of cell damage and cell death. However, when healthy individuals are exposed to potentially harmful compounds in the environment, the number of cells that undergo damage and death can increase considerably.

Scientists now agree that the assessment of the extent of cell damage (to a particular body organ) in a community as a whole, can be used to estimate a community's risk of developing cancer (for that particular organ). Since we are studying urinary bladder cancer, we will be studying cells in the urine. Cells lining the bladder are continuously being shed into the urine and we will examine these cells for cell damage. It must be emphasised that this test cannot be used to predict the chances of an individual developing bladder cancer as there are vet no agreed upon normal levels of cell damage for an individual. However, the amount of damage seen in the community as a whole provides a crude estimate of the risk faced by that community. Comparisons across communities will help us to determine which communities are at higher risk than therefore. others. and. merit further investigation.

#### How do I benefit?

Your participation will help provide a better understanding of the risk of developing bladder cancer for your community. Health regulatory authorities depend on such research findings to set safety values for exposure to potentially harmful substances. This type of research is essential to improve the health of the community as a whole.

#### Who can participate in the study?

We are seeking the assistance of people who meet the following eligibility criteria:

- Males aged 30-65 years;
- Living in Adelaide for at least six months prior to the survey;
- Do not swim regularly (ie, once or more a week) in chlorinated swimming pools; and
- Never been diagnosed as having cancer.

All members of your household who meet the above criteria are eligible to participate.

We would be calling you in the near future to set up appointments. However, if you have any questions or concerns prior to our call, please do not hesitate to contact the project coordinator - Dr Geetha Ranmuthugala. The coordinator can be reached at:

Tel 1 800 812 954 Fax (02) 6249 0740

E-mail: gpr868@nceph.anu.edu.au

Thank you for your assistance

## Appendix 29:

## National Centre for Epidemiology and Population Health The Australian National University

## **Environmental Study Consent Form**

(To be signed by the participant in the presence of the ANU interviewer during the home visit. The participant's signature is to be witnessed by the ANU interviewer.)

	Ž
I, hereby of environment study of my free will.	consent to participate in the
I will assist by responding to the telephone interviews a urine samples as has been explained to me. I also agree trained laboratory person.	1 0
I am free to withdraw from the study at any time.	
Signed	Date
Witness	Date
Name of witnessing person	

a

## Appendix 30:



## **Environmental study**

## **Interviewers Declaration**

information pro Identifiable info	vided to me during this study will be ormation will only be accessible to the No person will be identifiable from	e kept completely confidential. nose directly involved with this
Signed		Date

NATIONAL CENTRE FOR EPIDEMIOLOGY AND POPULATION HEALTH
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e-mail:

## **Appendix 31:**



**Environmental study** 

## **Instruction for collection of urine sample 1 (one)**

Thank you for assisting us with this study. We need you to provide us with a sample of urine that you pass for the first time in the morning.

- 1. Use the glass bottle that we have provided to collect this sample. As it is glass, please be careful when handling the bottle.
- 2. It is important to fill the container right to the top, almost to overflow level. It is important that there is no space for an air bubble when you put the lid on.
- 3. Screw the cap on firmly.
- 4. Rinse the outside of the bottle thoroughly and wipe it dry with a clean paper towel or cloth.
- 5. Write the date and time on the label.
- 6. Place the bottle in the plastic bag that we have provided.
- 7. Place this in the brown paper bag and fold the top of the bag down.
- 8. Keep the sample out of direct light or heat at all times.
- 9. Place the sample in the agreed place.
- 10.If it is raining, or it is likely to rain, and we have arranged to collect the sample from a previously identified spot at your home, place the brown paper bag in a plastic bag to prevent it from getting wet in the rain.

We will be using this urine sample to measure organic matter in the urine picked up from the environment. These results cannot be interpreted on an individual basis and we will be looking at the results of the community as a whole. Therefore, the results of individual tests on these urine samples will not be available. Thank you once again for your support of this study.

## Appendix 32.



**Environmental study** 

## Instruction for collection of urine sample 2/3

Thank you for assisting us with this study. We need you to provide us with the entire amount of urine you pass for the *second and third* times for the day. Please do not collect any urine from the first time you pass urine for the day.

- 1. Use the bottle marked '2' to collect the second void (ie. the second time you pass urine for the day)
- 2. Use the bottle marked '3' to collect the third void (ie. the third time you pass urine for the day)
- 3. For each collection, collect the ENTIRE VOLUME in the appropriate bottle
- 4. Screw the cap on firmly
- 5. Rinse the outside of the bottles thoroughly and wipe it dry with a clean paper towel or cloth
- 6. For each of the bottles, write the collection date and time on the labels
- 7. Place each bottle in the plastic bags that we have provided
- 8. Place the two plastic bags with the bottles in the one brown paper bag and fold the top of the bag down
- 9. Keep the samples out of direct light or heat at all times
- 10.Place the samples in the agreed place
- 11. Place the envelope containing the completed residential history sheet and the diary with the samples (inside the paper bag) for collection by our collectors.
- 12.If it is raining or it is likely to rain, and we have arranged to collect the sample from a previously identified spot at your home, place the brown paper bag in a plastic bag to prevent it from getting wet in the rain

We will be using these urine samples to count a particular type of cell in urine that may be affected by exposure to certain agents in the environment. As with the previous urine test, these results cannot be interpreted on an individual basis and we will be looking at the results of the community as a whole. The results of individual tests will not be available.

Thank you for your support of this study.

## **Appendix 33: Randomised controlled trial**

Since the discovery of its disinfection properties, chlorine has been used widely to disinfect community water supplies in industrially developed and developing countries. The discovery of disinfection by-products (DBP) in the early 1970's caused much concern over the potential public health risks associated with these by-products.

Although many epidemiological studies have been undertaken in an attempt to establish the causal link between DBP in drinking water and cancer, the evidence has not been convincing for several reasons. The long latency period associated with cancer development, the small relative risk over and above those drinking unchlorinated water, (as is with most environmental carcinogens), difficulty in assessing confounders, and the difficulty in accurately measuring and quantifying exposure, are all contributory factors.

This is a prospective study that measures and quantifies exposure to DBPs at an individual level. It overcomes the chronicity issue by using a biomarker of genotoxic damage.

## 1.1 Objectives

This study will determine if exposure to DBP in chlorinated drinking water will alter the frequency of micronucleated bladder epithelial cells in humans, thereby providing evidence for, or against, the carcinogenic potential of DBP. Micronuclei in bladder epithelial cells are a biomarker of genotoxic damage. Micronuclei are DNA fragments or whole chromosomes that are not incorporated into daughter cells during mitosis: Chromosomal aberrations are thought to be an initial step in carcinogenesis. It is therefore proposed that the effect of DBP on the frequency of micronucleated bladder epithelial cells will serve as an indicator of the potential carcinogenicity of DBP in drinking water.

## 1.2 Setting

To establish a baseline for micronucleated bladder epithelial cells, it was necessary to identify an unchlorinated water supply. Bungendore is a community in NSW, 40 km north of Canberra, with a population of about 2500 people. A single water bore supplies the community with the bulk of its water for most of the year, demand being augmented by a second bore, especially during summer. The town supply was unchlorinated.

In response to the occasional finding of coliforms in routine monitoring, the Yarrowlumla Shire Council was considering the need to introduce chlorination in 1998. This situation presented an opportunity to conduct a pre- and post- chlorination RCT.

## 1.3 Sample size and power

Sample size calculation and justification of eligibility criteria are only summarised here as they have been provided in more detail in the methods chapter of this thesis.

Sample size was established from risk ratios estimated by case-control studies. Case control studies have a higher degree of measurement error resulting in bias of the risk assessment towards the null. It was therefore expected that the true relative risk would be higher than the estimated 1.2 - 1.8 collectively found in these studies. Accordingly, the sample size calculation was adjusted in order to detect a relative risk as low as 1.4. A total of 140 participants, (70 in each group) would be sufficient to demonstrate a relative risk of 1.4 with 80% power (table 33.1). These calculations used a confidence level of 0.05.

Table 33.1: Summary of sample size and power calculation

Number of participants	140			150		
Relative risk	1.35	1.40	1.45	1.40	1.45	1.50
Power	73.5	80.8	86.4	75.7	81.8	86.6

## 1.4 Sampling frame and study population

The Bungendore electoral role by age group will be used to select all males aged between 45 and 65 years of age, resident in Bungendore. The rationale of using this age group, and a male only group, is provided in the methods chapter. Due to the relatively small population in Bungendore of approximately 2,500 residents, sampling is not thought necessary to find 70 males, aged 30 and 65 of years, meeting all eligibility criteria. All Bungendore households will therefore be sent letters to identify eligible persons. If more than 70 eligible persons (willing to participate) are identified, a sample of 70 persons will be randomly selected.

## 1.5 Study procedure

All selected persons will be sent study information letters. Follow up telephone calls will be made to offer more detailed information and to determine eligibility and willingness to participate. Consenting persons will be randomly allocated to one of two groups by allocating each participant a number, and generating a list of random numbers using SAS software. The total number of random numbers generated will be half the total number of participants. Those participants whose number was drawn will be allocated to be in the initial experimental group.

On recruitment to the study, participants will be invited to provide informed consent and demographic information on themselves. An initial urine sample will be collected to determine the baseline rate of micronuclei in bladder epithelial cells.

Study participants will then be delivered bottled water on a weekly basis for five weeks. Study participants will be blinded to the chlorination status of the water. Each participant will be asked to drink at least two glasses of bottled water every day for the duration of the study. At the end of the fifth week, they will be invited to complete an environmental and health questionnaire, which will identify and quantify exposure to other known risk factors for bladder cancer. Detailed information on water consumption and exposure to chlorinated water supplies (e.g. chlorinated swimming pools, drinking patterns at work place, visiting people in chlorinated areas) will be collected in this questionnaire. A semi-quantitative food frequency questionnaire (FFQ) will be used to assess dietary intake of micronutrients which affect cell integrity, over

the same time period. At the end of the fifth week, second and third samples of urine will be collected on two consecutive days.

The next stage of the study will be a four-week wash out period where no bottled water will be delivered. Since micronuclei are measures of acute exposure, a four-week period is sufficient for levels to return to baseline level. It has been demonstrated by Rosin and Stich in 1983 that the frequency of micronucleated mucosal cells (of head and neck region) increase within seven days of commencing radiotherapy, and return to baseline levels within a month of cessation of treatment [Rosin, 1992 #489]).

The washout period will be followed by a crossover phase of the study. Individuals who were initially receiving chlorinated water will now receive unchlorinated water, and those initially receiving unchlorinated water will now receive chlorinated water. Participants will once again be blinded to the chlorination status of the bottled water. A fourth urine sample will be collected when water delivery commences, to ensure the return of micronuclei levels to baseline levels. Once again, participants will be requested to drink at least two glasses of bottled water a day. At the end of the fifth week, the environment and health questionnaire and the FFQ will be administered, and a fifth and sixth urine sample will be collected.

## 1.6 Source of water, bottling of water, and quality control

Canberra water will be used as the source for bottling chlorinated water, while Bungendore water will be used as the source for unchlorinated water. Bottling of water and delivery will be done in accordance with established guidelines, and the water will be tested to ensure that NHMRC guidelines for drinking water are adhered to. (See pilot study 1). Water will be tested by ACTEW Corporation for DBP levels and microbiological quality prior to distribution to participants.

## 1.7 Measuring exposures

Exposure will be quantified for each participant using the volume of water intake and the concentration of DBP in the water supply. This will be the external dose of

exposure. The concentration of DBP in urine will be used as a surrogate for the internal dose of exposure.

## 1.8 Measuring outcomes

All collected urine samples will be examined for the frequency of micronucleated bladder epithelial cells, using the Fluorescence in-situ Hybridisation (FISH) technique. This will be done by a person trained in the method, and who will be blind to the exposure status of the study subject.

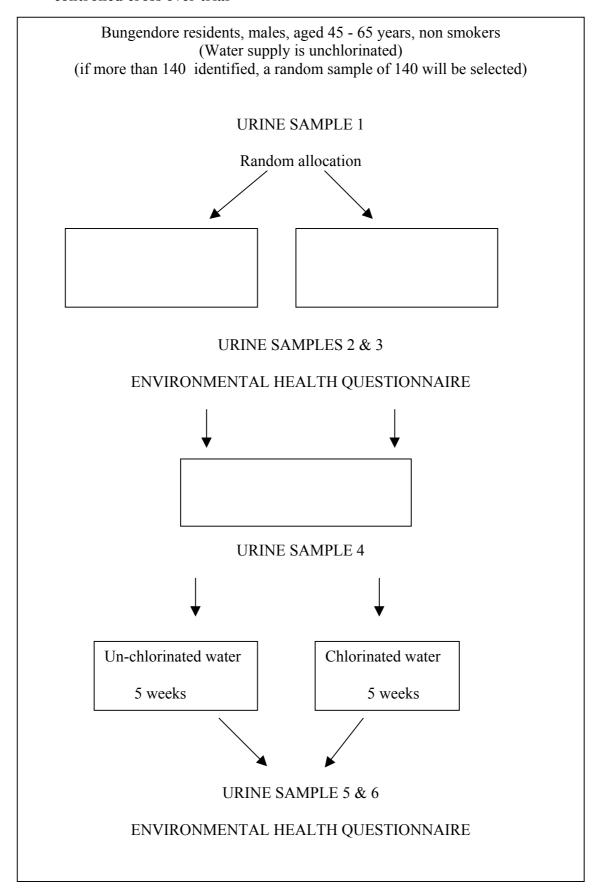
## 1.9 Summary of respondent burden

A total of 140 participants will be invited to participate in the study. Each participant will be enrolled in the study for a period of 14 weeks, during which time they will be expected to drink at least two glasses of the provided bottled water per day, during the first and last five weeks of the study. They will also be required to provide six samples of urine, and respond to the environment and health questionnaire and FFQ twice. This has been diagrammatically represented in figure 33-1

## 1.10 Statistical Analysis

Frequencies of micronucleated bladder epithelial cells will be examined for differences in the two groups using t-tests and analysis of variance. The frequency of micronucleated cells will also be examined in relation to the various DBP levels to determine if a dose response relationship exists using generalised linear modeling. The analysis will adjust for potential confounders using linear models including multiple regression, and ANOVA techniques.

Figure 33-1. Diagrammatic representation of the double-blinded randomised controlled cross over trial



## Appendix 34: Abbreviations used

ACT Australian Capital Territory

AOX Absorbable Organic Halogen

ASR Age standardised incidence rate

DBP Disinfection by-product

CHBrCl<sub>2</sub> Bromodichloromethane

CHBr<sub>2</sub>Cl Dibromochloromethane

CHBr<sub>3</sub> Bromoform CHCl<sub>3</sub> Chloroform

FFQ Food frequency questionnaire

FISH Fluorescence in Situ Hybridisation

MAC Maximum admissible concentration

ml millilitre

NHMRC National Health & Medical Research Council (Australia)

OR Odds ratio

RR Relative risk

RCT Randomised Controlled Trial

THM Trihalomethane

SE Standard Error

RR Relative risk

US EPA United States Environmental Protection Agency

WFR Weighted food record

## **Appendix 35: Glossary**

**Adduct** – A complex that forms when a chemical binds to a biological molecule, such as DNA or a protein.

Apoptosis – Programmed cell death as signaled by the nuclei in normally functioning human and animal cells when age or state of cell health and condition dictates. It is an active process requiring metabolic activity by the dying cell, often characterised by cleavage of the DNA into fragments that give a so-called laddering pattern on gels. Cells that die by apoptosis do not usually elicit the inflammatory responses that are associated with necrosis, though the reasons are not clear.

**Carcinogen** – A substance that causes cancer.

**Carcinogenesis** - The generation of cancer from normal cells, correctly the formation of a carcinoma from epithelial cells, but often used synonymously with transformation,

tumourigenesis.

**Carcinogenicity** – The ability or tendency to produce cancer.

**Diploid** – Having a pair of each chromosome characteristics of a species (in man - 2n or 46 chromosomes).

Genotoxic – Describes a substance which harms an organism by damaging its DNA

**Guideline** – Directing principle, not enforceable by law (as opposed to standard).

**Hypoploid** – Having less than the full complement of DNA.

Lignin - Lignins are derived from an abundant and renewable resource: trees, plants, and agricultural crops. Commercial lignin is currently produced as a co-product of the paper industry, separated from trees by a chemical pulping process.

Lignosulfonates (also called lignin sulfonates and sulfite lignins) are products of sulfite pulping. Kraft lignins (also called sulfate lignins) are obtained from the kraft pulping process. Other delignification technologies use an organic solvent or a high pressure steam treatment to remove lignins from plants. Because lignins are very complex natural polymers with many random couplings, the exact chemical structure is not known. commercial lignosulfonates products comes from their dispersing, binding, complexing and emulsifying properties.

Industry first began to use lignins in the 1880s when lignosulfonates were used in leather tanning and dye baths. Since then, lignosulfonates have even found applications in food products, serving as emulsifiers in animal feed and as raw material in the production of vanillin. (Vanillin is widely used as an ingredient in food flavors, in pharmaceuticals and as a fragrance in perfumes and odormasking products.) Lignin uses have expanded into literally hundreds of applications - impacting on many facets of our daily lives. (Source: Lignine Institute Home Page at http://www.assnhq.com/li/whatis.htm)

**Micronuclei** - Nuclei, separate from and additional to the main nucleus of a cell, produced during the telophase of mitosis or meiosis by lagging chromosomes or chromosome fragments derived from spontaneous or experimentally induced chromosomal structural changes.

**Mutation** – A permanent transmissible change in the genetic material. Origin: L. Mutatio from mutare = to change.

Mutagenic – Inducing genetic mutation.

**Necrosis** - The sum of the morphological changes indicative of cell death and caused by the progressive degradative action of enzymes, it may affect groups of cells or part of a structure or an organ. Origin: Gr. Nekrosis = deadness.

**Standard** – Measure or specification to which others conform. It is enforceable by law.

Acknowledgment: Definitions for medical terms were obtained from the On-line Medical Dictionary (Cancer Web) http://www.graylab.ac.uk/omd/index.html