MANE TONAMON DART (MIOSIT 09/25/02

Effect of Perchloroethylene,
Smoking, and Race on Oxidative
DNA Damage in Female
Drycleaners

Objective

Determine if increased oxidative damage, including the formation of oxidative DNA adducts, is elevated in female dry cleaners to PERC PERC oxidized by cytochrome P450 to trichloroacetyl chloride trichloroethanol trichloroacetic acid (TCA)

TCA is a metabolite of PERC in both man and experimental animals

TCA is also a rodent hepatocarcinogen

TCA induces lipid peroxidation and oxidative DNA damage

Vitamin E protects against PERC toxicity in mice

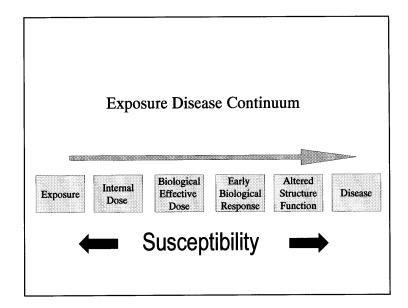
Un-repaired oxidative DNA damage can result in mutation and possibly cancer

Oxidative DNA damage used to assess exposures to chemicals that induce oxidative stress

8-Hydroxydeoxyguanosine (8-OHdG) adduct used extensively as a biomarker of effect

Rats did not exhibit elevated oxidative DNA damage following a single exposure to PERC

If PERC induces oxidative stress, biomarkers of oxidative damage could serve as biomarkers of exposure for workers in the dry cleaning industry.



Demographics

	Laundries	Dry cleaners	
Facilities	3	4	
Workers	20	18 13/5 10	
Race W/B	14/6		
Smokers	10		
Age	39 <u>+</u> 9	40 <u>+</u> 13	
BMI, kg	29 <u>+</u> 4	28 <u>+</u> 5	

Exposure Indices

Launderers	Dry cleaners
<0.02	2.4 <u>+</u> 3.4
<0.02	3.8 <u>+</u> 5.3
0.19 <u>+</u> 0.44	74.8 <u>+</u> 104.3
	<0.02

Potential Confounders

	Launderers	Drycleaners
Cotinine (smokers) mg/g creatinine	1504 <u>+</u> 879	1642 <u>+</u> 1216
Beta-carotene ug/Dl blood	8.7 <u>+</u> 5.6	19.9 <u>+</u> 24.0
Vitamin E alpha	8.4 <u>+</u> 3.1	9.0 <u>+</u> 2.2
Vitamin E beta mg/l blood	2.4 <u>+</u> 1.5	1.9 <u>+</u> 0.9

Oxidative Damage in PERC Exposed and Control Workers

	Launderers	Drycleaners
Leukocyte 8-OHdG	16.0 ± 7.3	8.1 ± 3.6 *
Pre-shift 8-epiPGF	535 ± 258	604 ± 314
Post-shift 8-epiPGFf	433 + 141	417 ± 174
Pre-shift 8-OHdG	2.8 ± 1.7	2.8 ± 1.2
Post-shift 8-OHdG	3.5 ± 1.3	3.1 ± 1.2

	PERC TWA	Blood PERC Launderers	Blood PERC Dry Cleaners	Blood PERC Dry Cleaners + Launders
Pre-shift				
Urine 8-epi-PGF ng/g creatinine	0.3587 * 0.157 b 17 °	-0.3199 0.211 17	0.0932 0.741 15	-0.0854 0.642 32
Urine 8-OHdG, μg/g creatinine	-0.1199 0.635 18	0.4661 0.044 19	0.3079 0.246 16	0.1782 0.306 35
Leukocyte 8-OHdG ng/mg dG	0.3296 0.196 17	-0.3032 0.237 17	0.2016 0.471 15	-0.5812 0.001 32
Post-shift				
Urine 8-epi-PGF ng/g creatinine	0.4688 0.058 17	0.0797 0.761 17	0.1843 0.511 15	-0.0954 0.604 32
Urine 8-OHdG µg/g creatinine	-0.3577 0.145 18	0.4296 0.075 18	0.0878 0.747 16	-0.0549 0.758 34

Oxidative Damage in PERC Exposed and Control Workers Stratified by Smoking Status

PERC exposure (n)	Urinary 8-epi-POF ng/gcreatinine	Urinary 8-OHdG µg/gcreatinine	Leukocyte 8-OHdG/dG ng/mg dG
Control nonsmoker (10)	476 ± 199	3.1 ± 1.5	17.1 ± 7.4^{a}
Control smoker (10)	479 ± 141	3.1 ± 1.3	14.8 ± 7.5^a
Exposed nonsmoker (8)	439 ± 177	28 ± 1.3	7.8 ± 26^{b}
Exposed smoker (10)	539 ± 256	3.1 ± 1.1	8.4 ± 4.4^{b}

Oxidative Damage in PERC Exposed and Control Workers Stratified by Race

PERC exposure (n)	Urinary 8-epi-PGF ng/g creatinine	Urinary 8-OHdG µg/g creatinine	Leukocyte 8-OHdG ng/mg dG
Control, black (6)	411 ± 90	2.8 ± 1.1	11.8 ± 5.9 a
Control, white (14)	506 ± 188	3.2 ± 1.5	17.8 ± 7.4^{b}
Exposed, black (5)	457 ± 171	3.1 ± 1.4	5.9 ± 1.2^{-a}
Exposed, white (13)	509 ± 246	2.9 ± 1.1	9.0 ± 3.9^{a}

PERC exposure was not associated with oxidative stress or increased oxidative DNA damage

Results indicate a reduction in oxidative DNA damage in PERC exposed dry cleaners relative to launderers,

but PERC could not clearly be defined as the source of the effect

The Guilty

- Mary Ann Butler
- Avima Ruder
- Christy Forrester
- Lauralynn Taylor
- David L. Ashley
- Patty Mathias
- Kate L. Marlow
- Kenneth L. Cheever
- Edward Krieg
- Howard Wey