For ONLINE APPENDIX for Thorax/2006/072199

CONSTITUENTS AND CONTAMINANTS WITH THE POTENTIAL TO CAUSE OR EXACERBATE RESPIRATORY DISEASE IN THE AEROSOL MISTS AT THE FACTORY

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CONSTITUENTS OR CONTAMINANTS, WHICH MAY HAVE PLAYED A PART IN CAUSING OR EXACERBATING SYMPTOMS OF BOTH EXTRINSIC ALLERGIC ALVEOLITIS(EAA) AND OCCUPATIONAL ASTHMA(OA)

Constituent or Contaminant and links to respiratory disease	Quantities found or identified in metalworking or wash fluids/mists	Comment
Bacteria - linked to both occupational asthma (OA) and extrinsic allergic alveolitis (EAA).	In 2004 high bacteria levels were counted in fluid samples from a number of washing machines and small sumps belonging to stand – alone metalworking machines. Two of these washing machines 38969 and 700315 were capable of expelling large quantities of contaminated mist in the area where most of those diagnosed with disease were working when they first became breathless	Some workers diagnosed with EAA demonstrated a serological response both to a crude metalworking fluid extract, taken in April 2004, from the large Mayfram sump, and to predominant bacteria, Ochrobactum anthropi and Acinetobacter sp², present as DNA fragments in the sump. In 2005 Two workers diagnosed with EAA have demonstrated adverse respiratory effects when challenged with samples of fluid from the Mayfram sump taken by HSE in September 2004 Ochrobactum anthropi has been linked to an outbreak of EAA in a car component factory in the USA³.

¹ Sera were taken from those diagnosed with EAA.
² HSL Report dated 9 September 2004 "Results of bacterial identifications from environmental sampling" 12 August 2004.
³ Project Sensor News Volume 16, No 1, winter 2004 – 5, Michigan State University, College of Human Medicine, 117 West Fee Hall, East Lansing, MI 48824 - 1316

Constituent or	Quantities found or	Comment
Contaminant and links to respiratory disease	identified in metalworking or wash fluids/mists	Comment
Endotoxin - linked to short term health effects and exacerbation of pre – existing asthma	In 2004 high endotoxin levels were recorded in samples taken by HSL from a number of machines and locations ⁴ . These were frequently the same locations where high levels of bacteria had been recorded e.g. in and around 38969 (camshaft wash)	May have had a role in some of the disease diagnosed. As they were found in wash 39869 they may have been spread over considerable area.
Fungi – linked to extrinsic allergic alveolitis and other respiratory conditions, but limited evidence of any link to occupational asthma	At all times, generally low levels of common fungi were found in all sumps and washing machines, although in August 2004 moderate levels were found in air in various parts of the factory (for example, on 12 August 2004 1.4 x 10³ were found in air near washer 700315, and 1.2 x 10³ in air in CPK 24 at the other end of the factory)	May have had a role in some respiratory disease.

⁴ See HSL Report dated 19/8/04

Constituent or Contaminant and links to respiratory disease	Quantities found or identified in metalworking or wash fluids/mists	Comment
Cobalt - linked to hardmetal disease, a severe lung disease, and asthma, both normally from exposure to cobalt dust. There was no exposure to cobalt dust at the factory, but it is possible	A number of samples ⁵ of fluids were taken from metalworking and washing machine sumps; generally levels were found < 0.1mg/l (or below the detection limit)	Respiratory disease from exposure to cobalt is normally associated with exposure to cobalt dust in the hard metal and diamond polishing industries
there was some cobalt in metalworking and wash fluids from the metals being machined or metalworking tools used	Concentrations of cobalt in air sampled ⁶ were below 0.01mg/m ³ Levels of cobalt in urine from those diagnosed with respiratory disease was not consistent with occupational exposure ⁷	The Occupational Exposure Standard (OES) ⁸ was 0.1mg/m ³ (8 hr time weighted average)
Other metals — including other hard metals, Iron, Chrome, Manganese and Nickel - either as constituents of metals being machined or metalworking tools used — linked to a variety of health effects on the skin and lungs, but not EAA or OA	Concentrations of these metals in air were sampled ⁹ at very low exposures, generally <0.01	Such exposures are not generally associated with any health effect, and were always below any Maximum Exposure Limit (MEL) ¹⁰ or OES

⁵ See HSL Environmental Monitoring Requests - samples collected on 12/8/04, 16/8/04, 23/8/04, and 14/9/04

See HSL Environmental Monitoring Request – samples collected on 25/1/05

⁷ See letter from University of Surrey, dated 26 July 2005

⁸ Occupational Exposure Standards (OESs) were in force at the time of the outbreak and represent concentrations of hazardous substances in air at which there is no indication of risk to the health of workers exposed by inhalation day by day. OESs have now been replaced by Workplace Exposure Limits (WELs). The actual levels at which all relevant OESs and WELs were set remained unchanged over the period of the outbreak.

⁹ See Environmental Monitoring Request – samples collected on 25/1/05

¹⁰ Maximum Exposure Limits (MELs) were in force at the time of the outbreak for substances which may cause the most serious health effects, such as occupational asthma, and for which "safe levels of exposure" cannot be determined. A MEL is also set for substances, for which although safe levels may exist, it is not reasonably practicable to reduce to those levels. MELs have now been replaced by WELs.

Constituent	Oughtities formal an	Commont
Constituent or	Quantities found or	Comment
Contaminant and links	identified in	
to respiratory disease	metalworking or wash	
	fluids/mists	
Mycobacteria – these	No Mycobacteria were	
have been linked to	detected in HSL	
outbreaks of EAA in the	samples ¹¹ taken from	
United States have	sumps	
been found in	_	
metalworking fluids after	HSL cultured	No significant reactions
large outbreaks of EAA	Mycobacteria for	were recorded between
in the USA	precipitin tests involving	sera and Mycobacteria
	sera from those	
	diagnosed	
Tramp oil – consists of	Between October 2003	It may be expected that
oils, such as machine	and February 2004	levels of exposure to
hydraulic and lubricating	concentrations of	mineral oil over the
oils, which leak into	mineral oil mist ¹² in air	period of the outbreak
metalworking fluid. At	of between 1 and 4	were somewhat higher
the factory over the	mg/m³ and an average	than those after steps
period of the outbreak	concentration of just	had been taken to
hydraulic and lubricating	above 1 mg/m³ were	reduce the amount of
oils were highly refined	recorded. Personal	tramp oil in the
mineral oils, with an	samples taken over the	metalworking fluid. No
exposure guidance	same period indicated	direct link between such
value of 3mg/m ³ .	exposures of between 1	exposure and either
Exposure to mineral oil	– 1.7 mg/m ³ , with an	occupational asthma or
mist alone has been	average exposure of 1.3	extrinsic allergic
linked to respiratory	mg/m ³ . These levels	alveolitis has been
irritation, shortness of	were recorded after the	demonstrated either in
breath, coughing and	outbreak had peaked	this outbreak or others.
tachypnea, but not	and action taken to	
occupational asthma or	reduce tramp oil.	
extrinsic allergic		
alveolitis.	The hydraulic oil in all	Onsets of
Tramp oil may form	systems was changed in	breathlessness peaked
mineral oil/water	April 2003, which	in March 2003
emulsions, which may	resulted in increased	
easily become	levels of tramp oil.	
microbiologically	·	
contaminated and		
support the proliferation		
of microorganisms.		
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¹¹ See HSL Reports dated 17/11/03 and 4/5/04
12 At the time of the outbreak, the guidance value for exposure to mineral oil, or neat oil, mist was 3mg/m³. This value is not based on any possible health effects at this level of exposure, but like the water – mix guidance value, represents good practice.

CONSTITUENTS OR CONTAMINANTS, WHICH ALONE MAY HAVE PLAYED A PART IN CAUSING OR EXACERBATING SYMPTOMS OF OCCUPATIONAL ASTHMA (OA)

Generally these are constituents, which have been linked to occupational asthma, but not extrinsic allergic alveolitis. There is no pattern in the use of particular fluids, or additives, such as biocides, or changes in them or of them which correspond to patterns of disease onset.

Constituent or Contaminant and links to respiratory disease	Quantities found or identified in metalworking or wash fluids/mists	Comment
Cases of occupational asthma have been linked to ethanolamines, including mono- and triethanolamine ¹³ .	Monoethanolamine was present in all the major metalworking and wash fluid concentrates used in the factory.	Taking into account the dilution of the concentrate the amount of Monoethanolamine in mist would have been extremely low.
		The same metalworking fluids had been in use in the factory with no major formulation changes since 1994.
		A new aluminium wash fluid was introduced in 2003, but this contained the about same proportion of Monoethanolamine as the wash fluid it replaced.
Triethanolamine - cases of asthma linked to ethanolamines, including mono- and triethanolamine ¹⁴ .	Present in wash fluid for cast iron.	Used since 1994.

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¹³ American Review of Respiratory Disease 115(5): 867-71, 1977 May, Allergy 1994; 49:877-881: 12 cases 1990 - 2005

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Constituent or Contaminant and links to respiratory disease	Quantities found or identified in metalworking or wash	Comment
Contaminant and links	identified in	All fluids in use have been unchanged (apart from the removal of diethanolamine) since 1998. Amount in mist would be extremely low. Would be present in mist at low concentrations <0.001mg/m³ at the guidance value for metalworking fluid concentrate (1mg/m³). Exposures at the factory were generally below this value.

Constituent or Contaminant and links to respiratory disease	Quantities found or identified in metalworking or wash fluids/mists	Comment
Sodium Polyacrylate – polyacrylates have been linked to occupational asthma	Present as a small proportion of a metalworking fluid.	Used unchanged since 1992 (apart from the removal of diethanolamine in August 2002).
		Polyacrylates named as suspected agents in 2 cases in nappy factory but kind of polyacrylate not named.
		About 50 cases due to monomeric acyrylates e.g. cyanoacrylates, methacrylates have been reported in the last 15 years
Tall Oil - suspected to be a sensitiser	Present in a metalworking fluid	Used unchanged since 1994 (apart from the removal of diethanolamine in August 2002).
16		Would be present in mist at low concentrations <0.1mg/m³ at the guidance value for metalworking fluid concentrate (1mg/m³). Exposures at the factory were generally below this value
Formaldehyde ¹⁵ released by biocides in use. May cause sensitisation by inhalation and skin contact	Very low concentrations of between 0.0130 and 0.1122 mg/m³ were measured¹6 at points around the factory in August 2004.	No new biocides have been introduced since 2000, with no changes in formulation since 2002. Formaldehyde had a MEL of 2ppm/2.5mg/m³ during the period of the outbreak.

¹⁵ Formaldehyde is under review by HSE.
16 HSL Sample Numbers 03500/04 to 0309/04 taken on 23/8/04