

Seminario per Medici Competenti

**Il rischio  
cancerogeno  
nelle lavorazioni  
con acciaio inox**

Venerdì 20 ottobre 2017  
ore 8.45 - 13.15

Sala della Quercia  
Padiglione A. Ziccardi  
Via Amendola, 2 - Reggio Emilia



# **Gli effetti sulla salute di Cromo, Nichel e Cadmio**

**Massimo Corradi**



**UNIVERSITÀ  
DI PARMA**

DIPARTIMENTO DI MEDICINA E CHIRURGIA

**Reggio E, 20 Ottobre 2017**

# Breath-taking jobs: a case–control study of respiratory work disability by occupation in Norway

AKM Fell,<sup>1</sup> R Abrahamsen,<sup>1,2</sup> PK Henneberger,<sup>3</sup> MV Svendsen,<sup>1</sup> E Andersson,<sup>4</sup>  
K Torén,<sup>4</sup> J Kongerud<sup>2,5</sup>

**Table 3** Risk of job change for occupations with at least 3 cases registered, and by gender

ISCO-88 code and occupation	Cases (n=125)	Controls (n=8352)	OR <sub>Crude</sub> (95% CI)	OR <sub>adj</sub> * (95% CI)	Females (n=47) OR <sub>adj</sub> * (95% CI)	Males (n=78) OR <sub>adj</sub> * (95% CI)
3230 Nurses (%)	3 (2)	303 (3.6)	0.65 (0.21 to 2.1)	1.2 (0.37 to 3.9)	1.6 (0.46 to 5.3)	–
3229 Health-associated professionals (except nursing) (%)	6 (5)	572 (6.8)	0.69 (0.30 to 1.6)	1.3 (0.53 to 3.0)	1.4 (0.5 to 3.7)	1.7 (0.23 to 13)
5122 Cooks/chefs (%)	7 (6)	196 (2.3)	<b>2.5 (1.1 to 5.4)</b>	<b>3.6 (1.6 to 8.0)</b>	<b>5.5 (2.2 to 14)</b>	1.3 (0.17 to 9.6)
5141 Hairdressers (%)	5 (4)	92 (1.1)	<b>3.7 (1.5 to 9.4)</b>	<b>6.4 (2.4 to 17)</b>	<b>8.6 (3.1 to 24)</b>	–
5220 Shop salespersons (%)	8 (6)	1127 (13.5)	<b>0.44 (0.21 to 0.90)</b>	0.75 (0.35 to 1.6)	1.0 (0.37 to 2.7)	0.58 (0.18 to 1.9)
6113 Gardeners (%)	3 (2)	47 (0.6)	<b>4.3 (1.3 to 14)</b>	<b>4.5 (1.3 to 15)</b>	–	<b>5.3 (1.5 to 18)</b>
7124 Carpenters and joiners (%)	6 (5)	176 (2.1)	<b>2.3 (1.0 to 5.4)</b>	<b>2.3 (0.98 to 5.5)</b>	–	<b>2.2 (0.92 to 5.2)</b>
7212 Welders (%)	5 (4)	53 (0.6)	<b>6.5 (2.4 to 17)</b>	<b>5.2 (2.0 to 14)</b>	–	<b>5.0 (1.9 to 13)</b>
7213 Sheet metal workers (%)	5 (4)	50 (0.6)	<b>6.9 (2.7 to 18)</b>	<b>5.4 (2.0 to 14)</b>	–	<b>5.1 (1.9 to 14)</b>
7230 Machinery mechanics and fitters (%)	5 (4)	155 (1.9)	2.2 (0.89 to 5.5)	2.0 (0.81 to 5.2)	<b>22 (2.6 to 191)</b>	1.6 (0.56 to 4.4)
7231 Motor vehicle fitters (%)	3 (2)	112 (1.3)	1.8 (0.57 to 5.9)	1.7 (0.51 to 5.4)	–	1.6 (0.50 to 5.3)
8332 Earth moving and related plant operators (%)	4 (3)	101 (1.2)	2.7 (0.89 to 7.5)	2.5 (0.88 to 7.1)	–	2.4 (0.84 to 6.8)
9131 and 9132 Cleaners (%)	7 (6)	152 (1.8)	<b>3.2 (1.5 to 7.0)</b>	<b>5.0 (2.2 to 11)</b>	<b>5.7 (2.3 to 14)</b>	3.9 (0.51 to 31)
9211 Agricultural labourers (%)	4 (3)	44 (0.5)	<b>6.2 (2.2 to 18)</b>	<b>7.4 (2.5 to 22)</b>	<b>12 (2.6 to 57)</b>	<b>5.3 (1.2 to 24)</b>

Values in bold typeface are statistically significant at  $p < 0.05$ .

\*Adjusted for each other and age, gender and smoking.

–: less than two cases.

» **Febbre da fumi metallici**

» **Polmonite**

» **Asma**

» **BPCO**

» **Fibrosi polmonare**

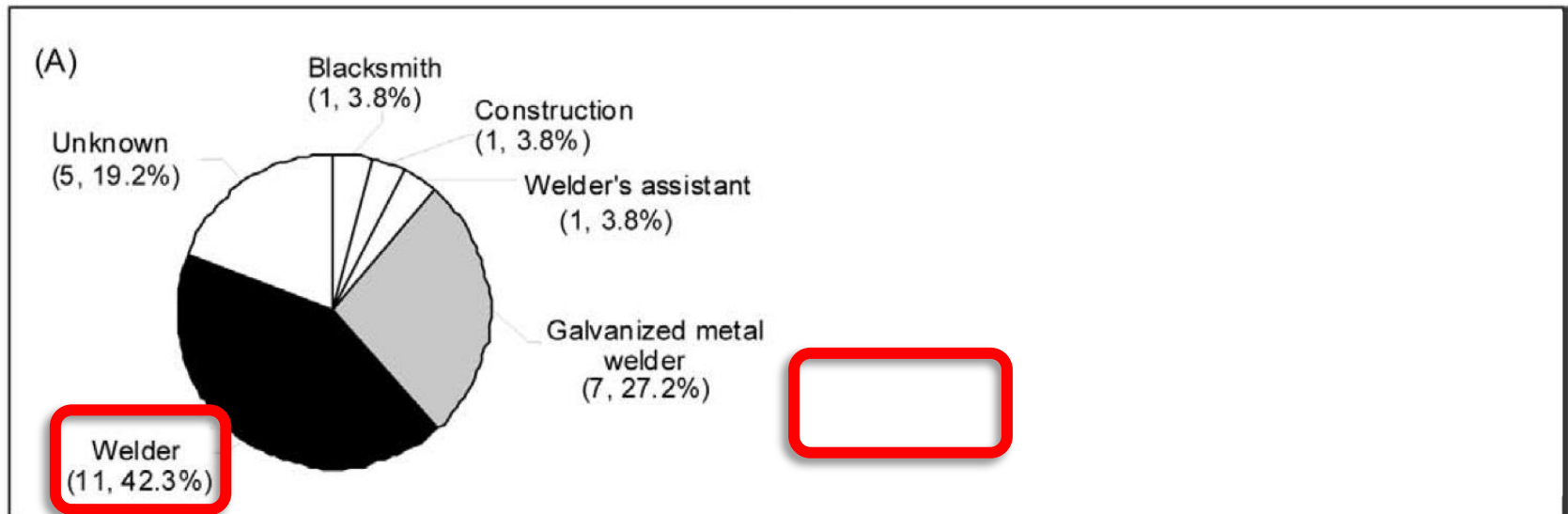
» **Tumore del polmone**



Reproduced with permission from the *JOURNAL of the Louisiana State Medical Society*  
[2009;161(6):348-351]

## Metal Fume Fever: A Review of the Literature and Cases Reported to the Louisiana Poison Control Center

Syed Atif Ahsan, MPH; Michelle Lackovic, MPH;  
Adrienne Katner, MPH; and Christine Palermo, PhD



**Figure 1.** The distribution of cases of metal fume fever reported to Louisiana Poison Control Center by occupation (A) and healthcare management (B) are shown. The number of reported cases and the percent of the total are given.

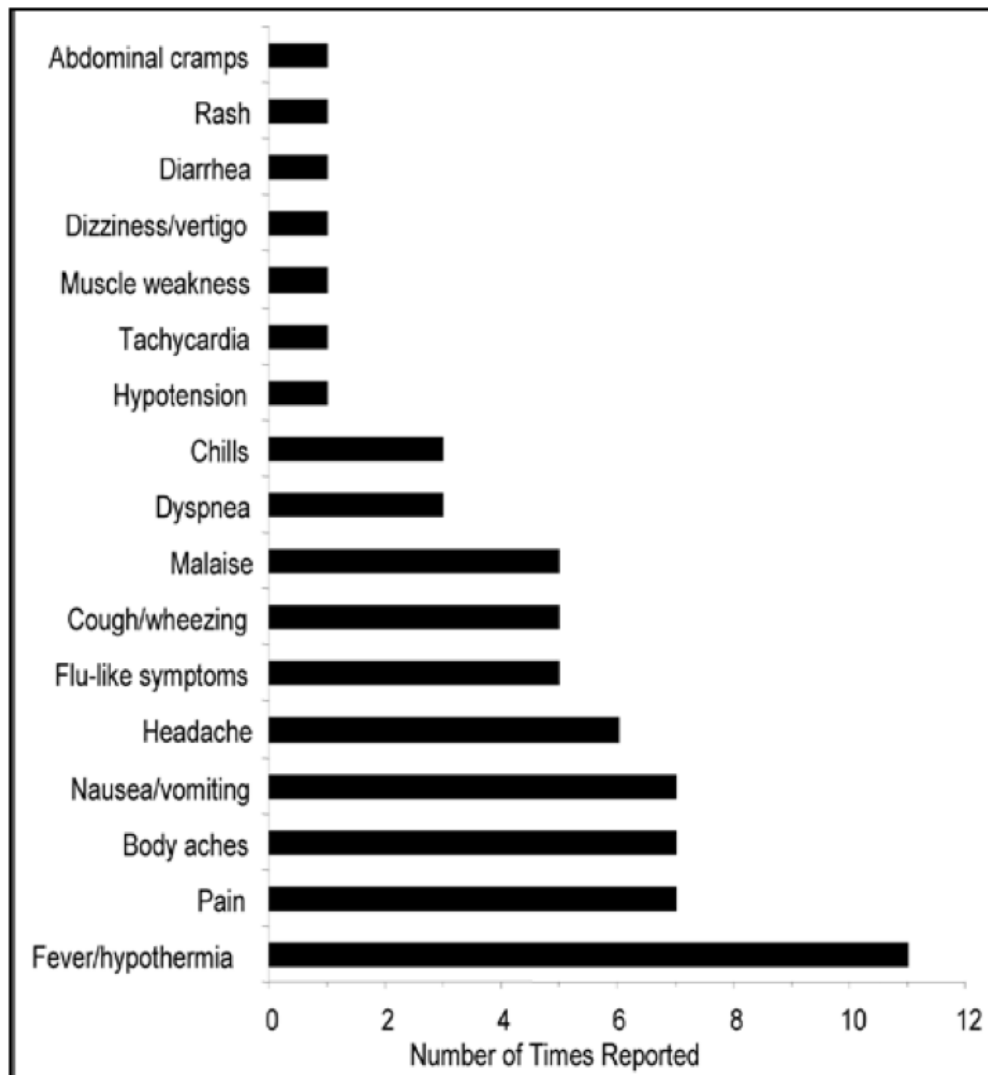
## Metal Fume Fever: A Review of the Literature and Cases Reported to the Louisiana Poison Control Center

Syed Atif Ahsan, MPH; Michelle Lackovic, MPH;  
Adrienne Katner, MPH; and Christine Palermo, PhD



**DANGER**  
**WELDING**

- Fumes and gases may cause irritation of the eyes, nose and throat
- Fumes and gases may cause chest pain/pulmonary edema
- Fumes and gases may cause chronic lung diseases/lung cancer
- Fumes and gases may cause metal fume fever/lead poisoning
- Polyester and other man-made fibers may melt and cause severe burns if struck by a welding spark
- May result in asphyxiation in confined spaces



# Metal fume fever

*Michele Malaguarnera, Filippo Drago, Giulia Malaguarnera, Giovanni Li Volti, Salvatore Salomone, Filippo Caraci, Fabio Galvano, Marco Vacante, Claudio Bucolo, Mariano Malaguarnera*

In September, 2011, a 28-year-old man with an 8-year history of occupational welding developed paraesthesiae, severe pain in both legs, weakness, myalgia, wheezing, malaise, conjunctivitis, dyspnoea, and high-grade fever. This fever developed 8 h after metal fume exposure and lasted for 24 h. Fever was recurring for 4 weeks.

In October, 2011, the possibility of heavy metal intoxication was investigated because of fever and osteoporosis in the patient. Urinary cadmium was 69·3  $\mu\text{mol/g}$  creatinine (normal values 8·8–46·2  $\mu\text{mol/g}$  creatinine). Other toxic metals did not differ from normal reference range. The patient changed jobs and started treatment with calcium and

**Lancet 2013; 381: 2298**

# Metal fume fever

*Michele Malaguarnera, Filippo Drago, Giulia Malaguarnera, Giovanni Li Volti, Salvatore Salomone, Filippo Caraci, Fabio Galvano, Marco Vacante, Claudio Bucolo, Mariano Malaguarnera*

increased osteoporosis and osteomalacia.<sup>1</sup> Diagnosis of metal fume fever is based on exposure to metal fumes within the last 48 h and development of influenza-like symptoms with resolution within 24–48 h

Although exposure to cadmium oxide fumes is the most frequent and best characterised cause of metal fume fever, other metal oxides such as arsenic, boron, zinc, chromium, copper, magnesium, manganese, nickel, and titanium are also suggested causes.<sup>2</sup>

# Is metal fume fever a determinant of welding related respiratory symptoms and/or increased bronchial responsiveness? A longitudinal study

M El-Zein, C Infante-Rivard, J-L Malo, D Gautrin

**Table 6** Association between possible MFF and welding related respiratory symptom suggestive of welding related asthma and increase in bronchial responsiveness in multiple logistic regression analysis\*

	At least one welding related respiratory symptoms (n = 32)	Persistent welding related respiratory symptoms (n = 19)	Increase in BR (n = 23)
	OR (95% CI)	OR (95% CI)	OR (95% CI)
Possible MFF (n = 91)	4.92 (2.10–11.52)	13.72 (3.63–51.84)	1.26 (0.49–3.24)

**Conclusion:** There is a strong association between welding related MFF and welding related respiratory symptoms suggestive of OA. As such, MFF could be viewed as a pre-marker of welding related OA, a hypothesis that requires further investigation.



» **Febbre da fumi metallici**

» **Polmonite**

» **Asma**

» **BPCO**

» **Fibrosi polmonare**

» **Tumore del polmone**





# Polmonite infettiva

**Table 3** Mortality risk for infectious pneumonia among male Swedish construction workers aged 20–64 years exposed to inorganic dust, chemicals, metal fumes and wood dust, adjusted for age and smoking based on Poisson regression models

Predictor (exposure)	RR (95% CI)		
	Infectious pneumonia (n = 145)	Lobar pneumonia (n = 49)	Pneumococcal pneumonia (n = 36)
Referents	1 n=26	1 n=5	1 n=3
Inorganic dust*	1.87 (1.22 to 2.87) n=108	3.37 (1.32 to 8.57) n=37	4.29 (1.28 to 13.86) n=28
Chemicals*	1.91 (1.37 to 3.22) n=31	4.53 (1.63 to 12.58) n=14	5.80 (1.62 to 20.88) n=10
Metal fumes*	2.31 (1.35 to 3.95) n=27	3.67 (1.33 to 10.11) n=9	5.77 (1.53 to 21.73) n=8
Wood dust*	0.90 (0.37 to 2.19) n=6	1.59 (0.31 to 8.17) n=2	2.61 (0.44 to 15.83) n=2
Any exposure*	1.80 (1.18 to 2.75) n=119	3.45 (1.37 to 8.70) n=44	4.21 (1.29 to 13.72) n=33

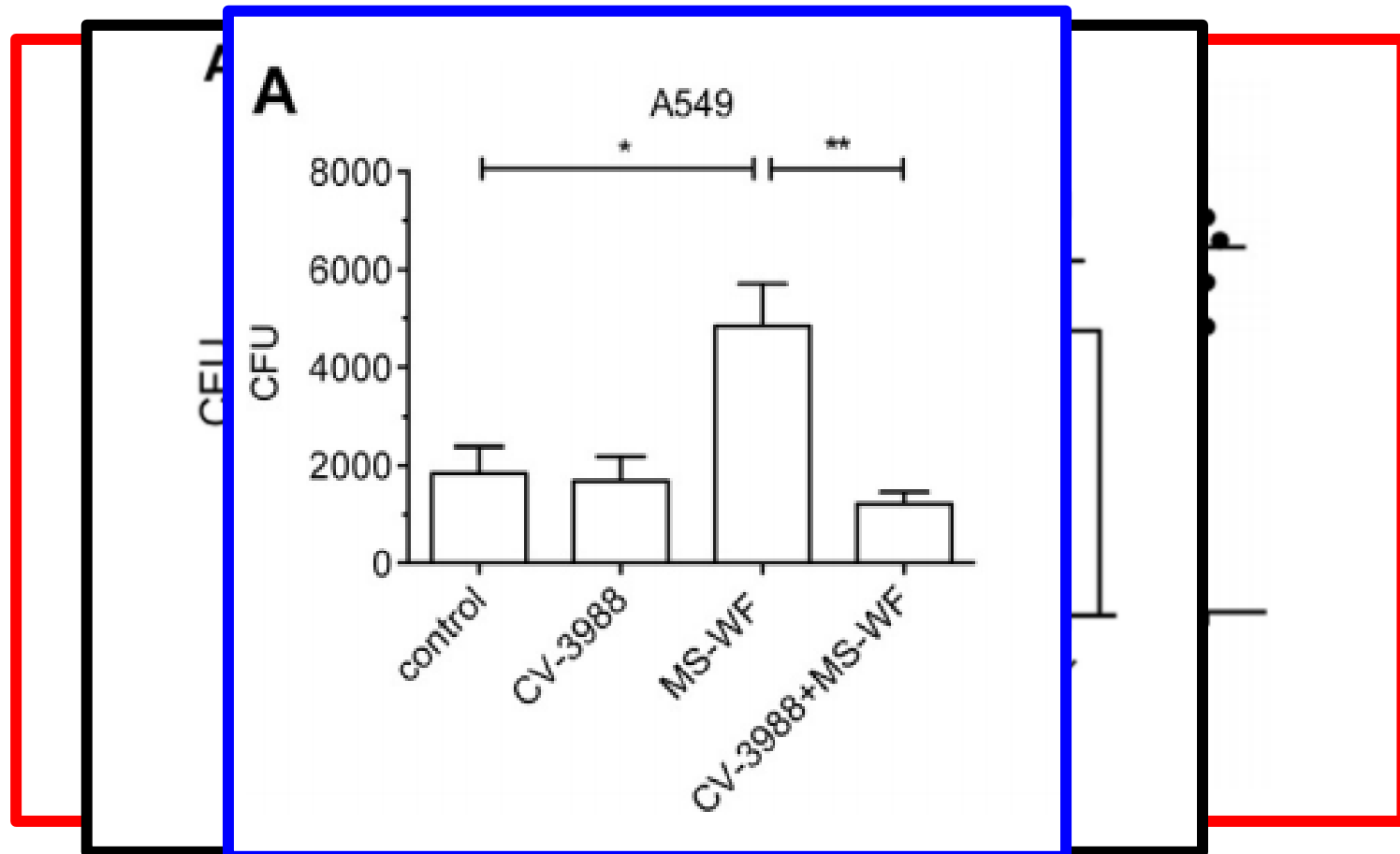
**Mortalità per polmonite lobare 3.7 volte maggiore e per polmonite da pneumococco 5.8 volte maggiore**

Increased mortality from infectious pneumonia after occupational exposure to inorganic dust, metal fumes and chemicals.

Toren K, et al. Thorax. 2011;66:992-6.

## Exposure to welding fumes and lower airway infection with *Streptococcus pneumoniae*

Reetika Suri, PhD<sup>a</sup>, Jimstan Periselneris, MB BS<sup>b</sup>, Sophie Lanone, PhD<sup>c</sup>, Patti C. Zeidler-Erdely, PhD<sup>d</sup>, Geoffrey Melton, BSc<sup>e</sup>, Keith T. Palmer, MD<sup>f</sup>, Pascal Andujar, MD<sup>c</sup>, James M. Antonini, PhD<sup>d</sup>, Vanessa Cohignac, MSc<sup>c</sup>, Aaron Erdely, PhD<sup>d</sup>, Ricardo J. Jose, MB BS<sup>b</sup>, Ian Mudway, PhD<sup>g</sup>, Jeremy Brown, MD<sup>b</sup>, and Jonathan Grigg, MD<sup>a</sup>

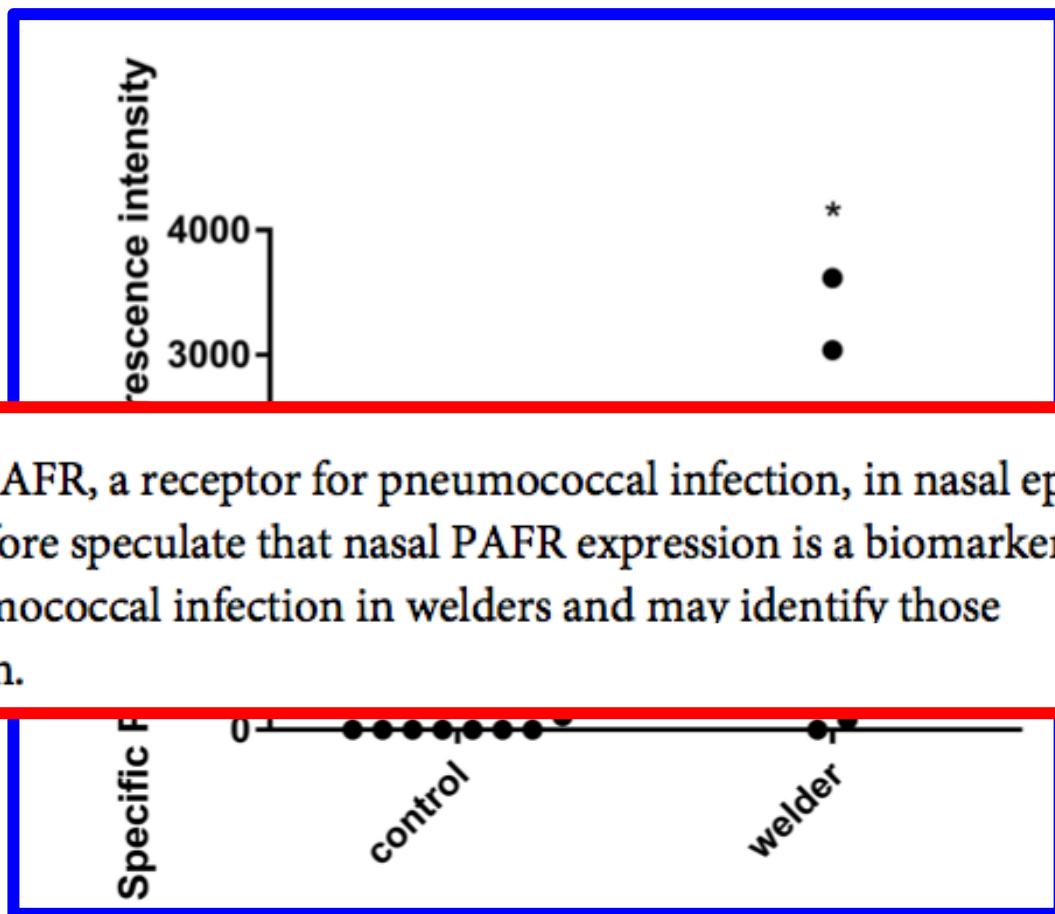


# Pneumococcal infection of respiratory cells exposed to welding fumes; Role of oxidative stress and HIF-1 alpha

Jonathan Grigg, Lisa Miyashita, Reetika Suri\*

PAFR has been proposed to

We found increased expression of PAFR, a receptor for pneumococcal infection, in nasal epithelial cells from welders. We therefore speculate that nasal PAFR expression is a biomarker for increased susceptibility to pneumococcal infection in welders and may identify those requiring pneumococcal vaccination.

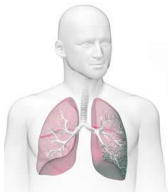




## Immunisation against infectious disease

### Individuals at occupational risk

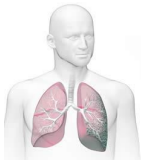
There is an association between exposure to metal fume and pneumonia and infectious pneumonia, particularly lobar pneumonia (Palmer *et al.*, 2003; Palmer *et al.*, 2009; Industrial Injuries Advisory Council, 2010; Toren *et al.*, 2011) and between welding and invasive pneumococcal disease (Wong *et al.*, 2010). PPV23 (single 0.5ml dose in those who have not received PPV previously) should be considered for those at risk of frequent or continuous occupational exposure to metal fume (e.g. welders) taking into account the exposure control measures in place. Vaccination may reduce the risk of invasive pneumococcal disease but should not replace the need for measures to prevent or reduce exposure.



# Polmonite infettiva

Nel 2011 il Dipartimento di Sanità in Inghilterra aveva **raccomandato la vaccinazione antipneumococcica per i saldatori (PPV23)**

VANTAGGI	LIMITI
Singola dose	Protegge solo dai sierotipi presenti nel vaccino
Sicuro con comparsa degli anticorpi dopo 3 settimane	Protezione per 5 anni
Riduzione dell'incidenza di Polmonite da pneumococco	Non sono raccomandate dosi successive
Riduzione della mortalità per Polmonite da Pneumococco	Età anagrafica/anni di esposizione



REPUBBLICA ITALIANA

## BOLLETTINO UFFICIALE

REGIONE DEL VENETO

Bollettino ufficiale vaccini 2015

Tab. 1 – Condizioni di rischio per malattia invasiva pneumococcica (MIP)

Condizioni	Rischio uniforme: Vaccinazione raccomandata a tutti	Rischio eterogeneo: Valutare la necessità della vaccinazione per singolo caso
<b>Condizioni con rischio moderato</b>		
- Immunosoppressione iatrogena (esclusi trapianto d'organo)		X
- Diabete mellito (in particolare in labile compenso o complicato)		X
- Malattia celiaca		X
- Malattie polmonari croniche, insufficienza respiratoria		X
- Persone che hanno avuto polmonite o malattia invasiva pneumococcica		X
- Cardiopatie croniche (in particolare patologie congenite con cianosi e insufficienza cardiaca)		X
- Alcolismo		X
- Tabagismo		X
- Individui con aumentato rischio per il tipo di lavoro (laboratoriisti, saldatori)		X

» **Febbre da fumi metallici**

» **Polmonite**

» **Asma**

» **BPCO**

» **Fibrosi polmonare**

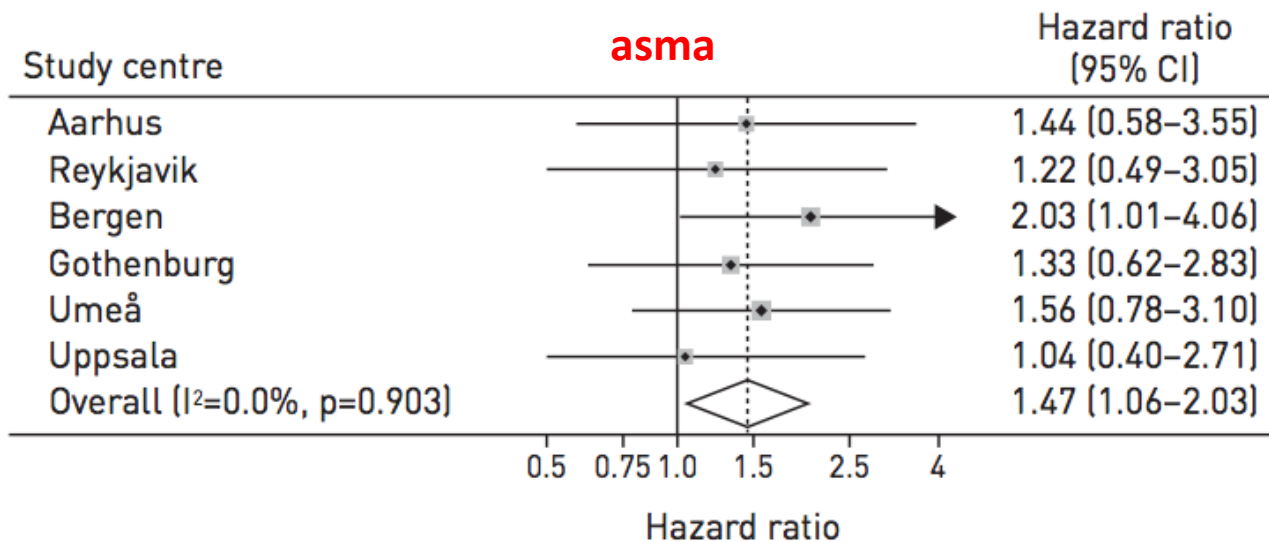
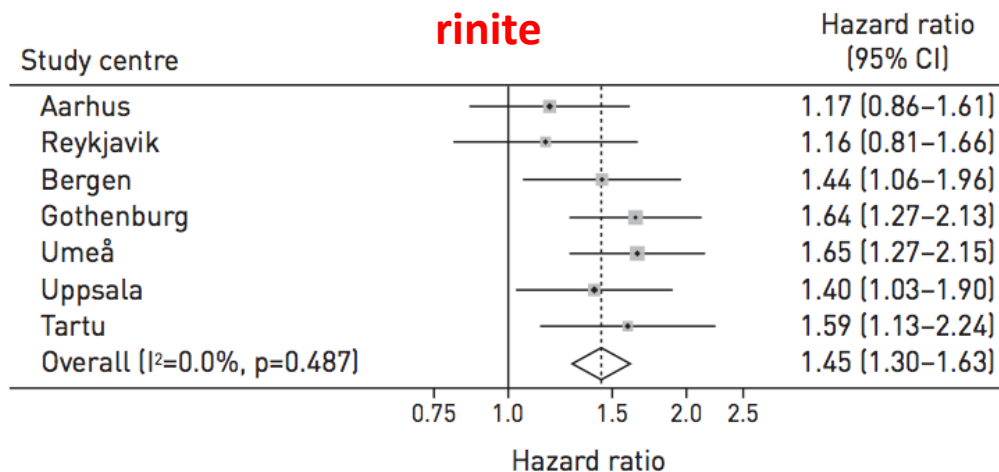
» **Tumore del polmone**





# Incidence of rhinitis and asthma related to welding in Northern Europe

Torgeir Storaas<sup>1,2</sup>, Jan-Paul Zock<sup>3</sup>, Ana Espinosa Morano<sup>3</sup>, Mathias Holm<sup>4</sup>, Eythor Björnsson<sup>5</sup>, Bertil Forsberg<sup>6</sup>, Thorarinn Gislason<sup>7,8</sup>, Christer Janson<sup>9</sup>, Dan Norback<sup>10</sup>, Ernst Omenaas<sup>11</sup>, Vivi Schlünssen<sup>12</sup>, Kjell Torén<sup>4</sup> and Cecilie Svanes<sup>1,13</sup>



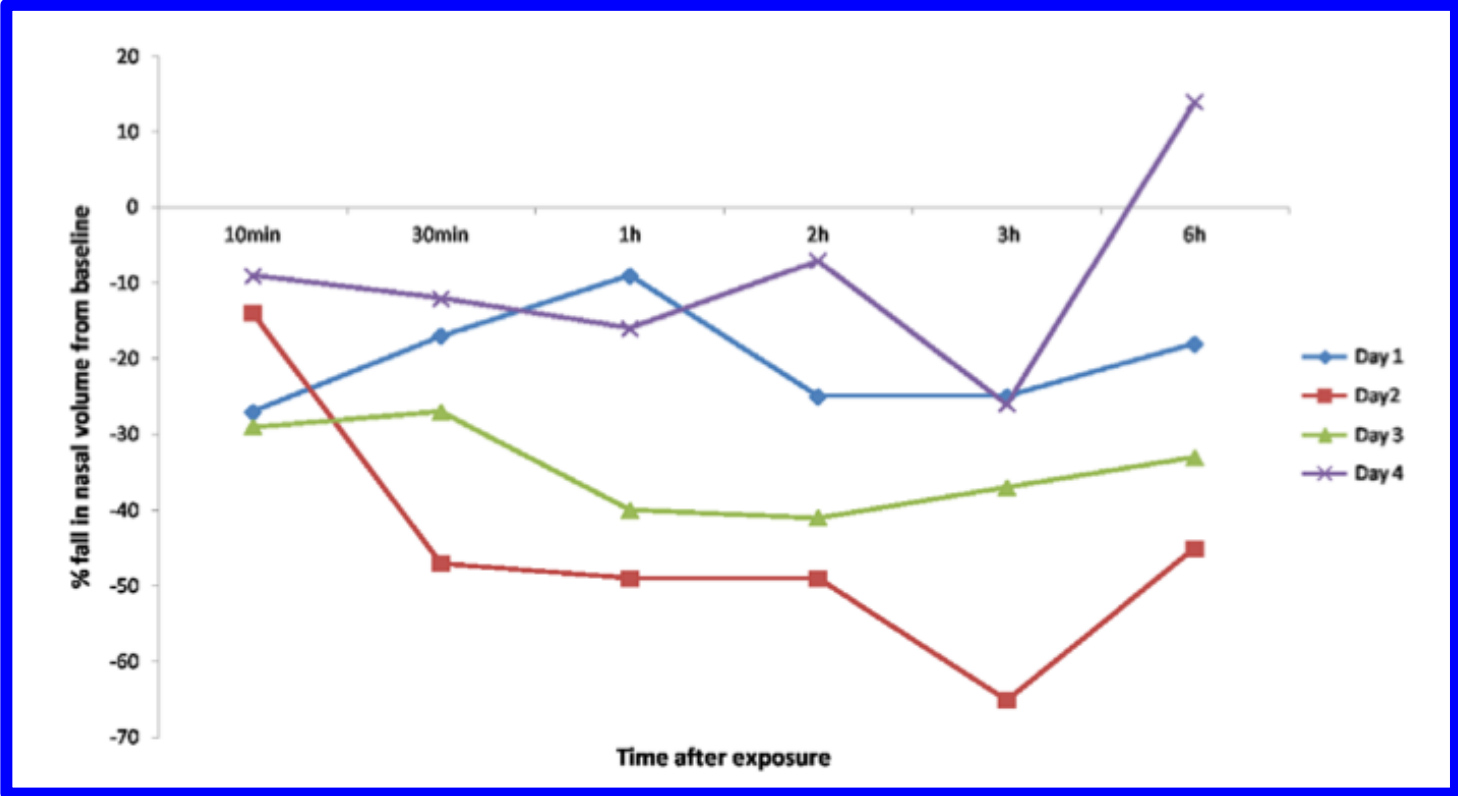
# Incidence of rhinitis and asthma related to welding in Northern Europe

Torgeir Storaas<sup>1,2</sup>, Jan-Paul Zock<sup>3</sup>, Ana Espinosa Morano<sup>3</sup>, Mathias Holm<sup>4</sup>, Eythor Björnsson<sup>5</sup>, Bertil Forsberg<sup>6</sup>, Thorarinn Gislason<sup>7,8</sup>, Christer Janson<sup>9</sup>, Dan Norback<sup>10</sup>, Ernst Omenaas<sup>11</sup>, Vivi Schlünssen<sup>12</sup>, Kjell Torén<sup>4</sup> and Cecilie Svanes<sup>1,13</sup>

	All participants		
	Number <sup>#</sup>	Incidence rate	HR
<b>Incident rhinitis subsequent to welding debut n</b>		11 203	
→ Never welded	3279	16.8	1 (Reference)
Ever welded	405	20.3	1.4 (1.3–1.6)
Welded <25% of working time	230	21.3	1.5 (1.3–1.7)
Welded >25% of working time	171	18.9	1.4 (1.2–1.6)
Welded, but not in stainless steel	364	20.0	1.4 (1.3–1.6)
Welded in stainless steel >6 months	38	23.4	1.6 (1.2–2.2)
{ Welding activity ≤8 years	162	22.0	1.5 (1.3–1.8)
{ Welding activity >8 years	222	18.8	1.3 (1.2–1.5)

# Occupational Rhinitis Due to Steel Welding Fumes

Roberto Castano, MD, PhD<sup>1,2\*</sup> and Eva Suarthana, MD, PhD<sup>2</sup>



# METAL-INDUCED ASTHMA AND CHEST X-RAY CHANGES IN WELDERS

TOMASZ WITCZAK<sup>1</sup>, WOJCIECH DUDEK<sup>2</sup>, JOLANTA WALUSIAK-SKORUPA<sup>2</sup>,  
DOMINIKA ŚWIERCZYŃSKA-MACHURA<sup>2</sup>, WOJCIECH CADER<sup>3</sup>, MONIKA KOWALCZYK<sup>1</sup>,  
and CEZARY PAŁCZYŃSKI<sup>1,2</sup>

Patient	Challenge test with:	↓FEV <sub>1</sub> ≥ 20% during test/ type of reaction	Positive SPT with:
1 (group A)	Ni	+/late	Ni
2 (group A)	Ni	+/late	Ni
3 (group A)	Ni	+/dual	V
4 (group A)	Cr	+/late	-
5 (group A)	Cr	+/late	-
6 (group A)	Cr	+/dual	Ni
7 (group A)	Cr	+/late	V
8 (group A)	Co	+/late	Co
9 (group A)	Mn	+/late	-

» **Febbre da fumi metallici**

» **Polmonite**

» **Asma**

» **BPCO**

» **Fibrosi polmonare**

» **Tumore del polmone**



# Occupational chronic obstructive pulmonary disease: a standard of care

D. Fishwick<sup>1</sup>, D. Sen<sup>2</sup>, C. Barber<sup>1</sup>, L. Bradshaw<sup>1</sup>, E. Robinson<sup>1</sup>, J. Sumner<sup>1</sup> and The COPD Standard Collaboration Group\*

**Table 2.** Evidence-based statements for occupational COPD relating to prevention, causation, behavioural issues and risk

Evidence grade	Evidence-based statement
A1 *** SIGN 2++	Occupational exposures are a risk factor for the development of COPD and account for approximately 10–15% of all COPD
A2 ** SIGN 2+	The prevalence of COPD in working populations varies and can be as high as 30% in the working age population
A3 *** SIGN 2++	Occupational agents reported to cause COPD with varying degrees of supporting evidence include coal mine dust, silica, asbestos, refractory ceramic fibres, flour, endotoxin, cadmium, carbon black, agricultural dusts (from poultry, animal and arable farming products and practices), dusts from rubber, cotton, wood, iron/steel and smelting, <u>welding fumes</u> , isocyanates and other chemicals

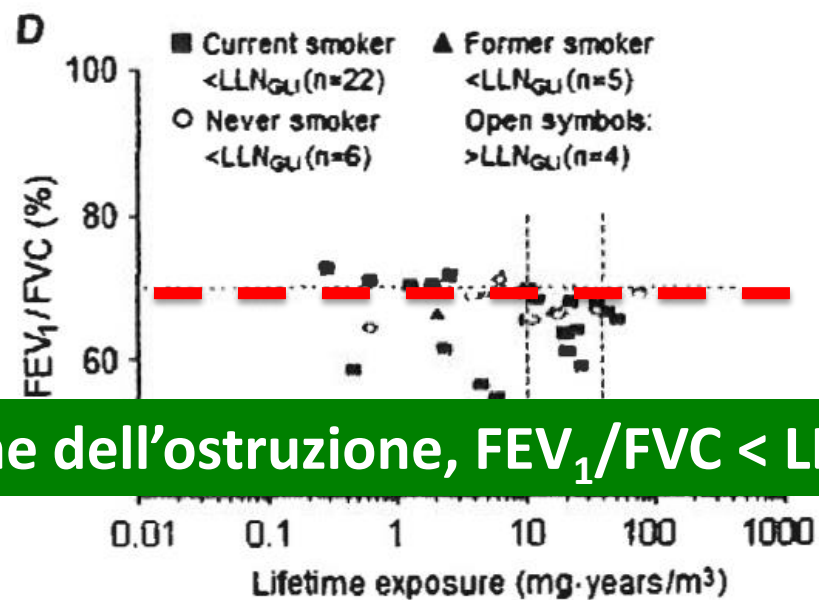
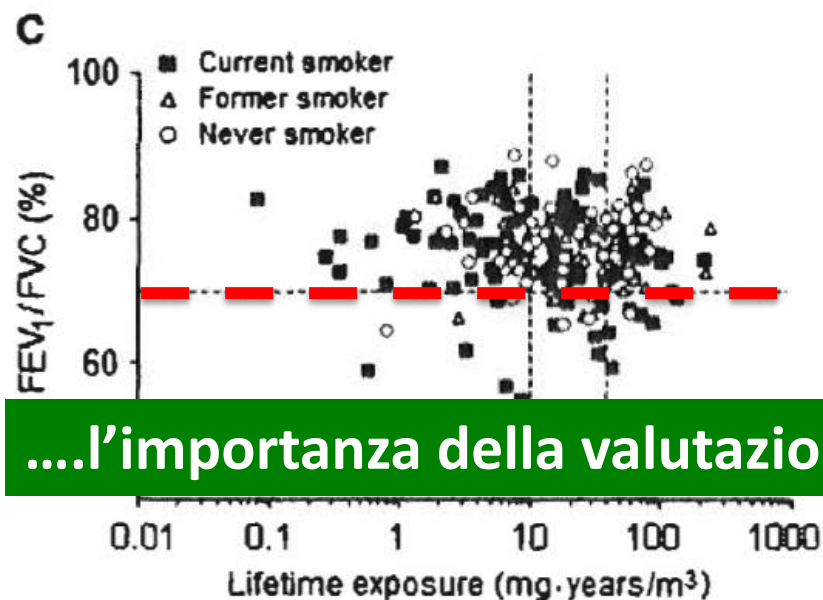
# Occupational chronic obstructive pulmonary disease: a systematic literature review

by Øyvind Omland, MD, PhD,<sup>1,2</sup> Else Toft Würtz, MHS,<sup>1</sup> Tor Brøvig Aasen, MD,<sup>3</sup> Paul Blanc, MD, PhD,<sup>4</sup> Jonas Brisman, MD, PhD,<sup>5</sup> Martin Reginald Miller, MD,<sup>6</sup> Ole Find Pedersen, MD, DMSc,<sup>2</sup> Vivi Schlünssen, MD, PhD,<sup>2</sup> Torben Sigsgaard MD, PhD,<sup>2</sup> Charlotte Suppli Ulrik, MD, DMSc,<sup>7</sup> Sven Viskum, MD<sup>1</sup>

Reference	Exposure	Study design	Population	Outcome	Measure of association
<b>Welding</b>					
Lou et al, 2006 (57)	Spot and arc welders Air sampling	Cross-sectional study	N=247 (130 referents) Age 22–56 yrs	FEV <sub>1</sub> /FVC <0.75	Non significant. Borderline linear trend (P=0.08) to FEV1 decline and spot welding.
Gennaro et al, 1993 (58)	13 job categories in shipyard workers Questionnaire-defined	Cross-sectional study	N=657 (174 referents) Mean age 45.7 yrs	Obstructive pulmonary function: Normal FVC and low FEV <sub>1</sub> /FVC Mixed pulmonary function impairment: Low FVC and low FEV <sub>1</sub> /FVC	Obstructive pulmonary function: No significant association with job title Mixed pulmonary function: No significant association with job title Significant OR 2.52 (95% CI 1.15–5.53) for >20 yrs compared with <20 yrs experience for mixed impairment
Bogadi-Sare, 1990 (59)	Dust and fumes of stainless steel welding Defined by work place.	Cross-sectional study	N=186 (80 referents) Mean age exposed 38.5 yrs referents 36.9 yrs	FEV <sub>1</sub> /FVC ratio	Significant lower FEV1/FVC ratio: Smokers: 79.2% compared with referents 84.4%, P<0.05 Non-smokers: 80.4% compared with referents 92.8%, P<0.01.
Wang et al, 1996 (67)	Dust exposure in steelworkers Record (steel corporation)-defined	Longitudinal study; 4–9 yrs	N=475 (internal referents) Age at baseline 20–61 yrs	FEV <sub>1</sub> /FVC ratio	Significant reduce in FEV <sub>1</sub> /FVC ratio of 0.03%/yr, P=0.02

# Effects of Exposure to Welding Fume on Lung Function: Results from the German WELDOX Study

M. Lehnert, F. Hoffmeyer, K. Gawrych, A. Lotz, E. Heinze, H. Berresheim, R. Merget, V. Harth, R. Van Gelder, J.-U. Hahn, A. Hartwig, T. Weiß, B. Pesch, and T. Brüning, for the WELDOX Study Group



...l'importanza della valutazione dell'ostruzione, FEV<sub>1</sub>/FVC < LLN



Smoking, cadmium, and emphysema

---

# Smoking, cadmium, and emphysema

**D J Hendrick**

---

Does cadmium contribute to the development of smoking induced emphysema?

Urinary cadmium levels predict lower lung function in current and former smokers: data from the Third National Health and Nutrition Examination Survey

**D M Mannino, F Holguin, H M Greves, A Savage-Brown, A L Stock, R L Jones**

---

*Thorax* 2004;**59**:194–198. doi: 10.1136/thorax.2003.012054

Lung function and chest radiographs of 101 men who had worked for 1 or more years manufacturing copper-cadmium alloy were compared with those of a referent group matched for age, sex, and employment status. Cigarette consumption was similar in the two groups.

The cadmium workers had an excess of abnormalities of lung function and of radiographic changes consistent with emphysema. Classification of the cadmium workers by exposure categories based on either estimated cumulative cadmium exposure or liver cadmium measured by neutron activation analysis showed that abnormalities of lung function were

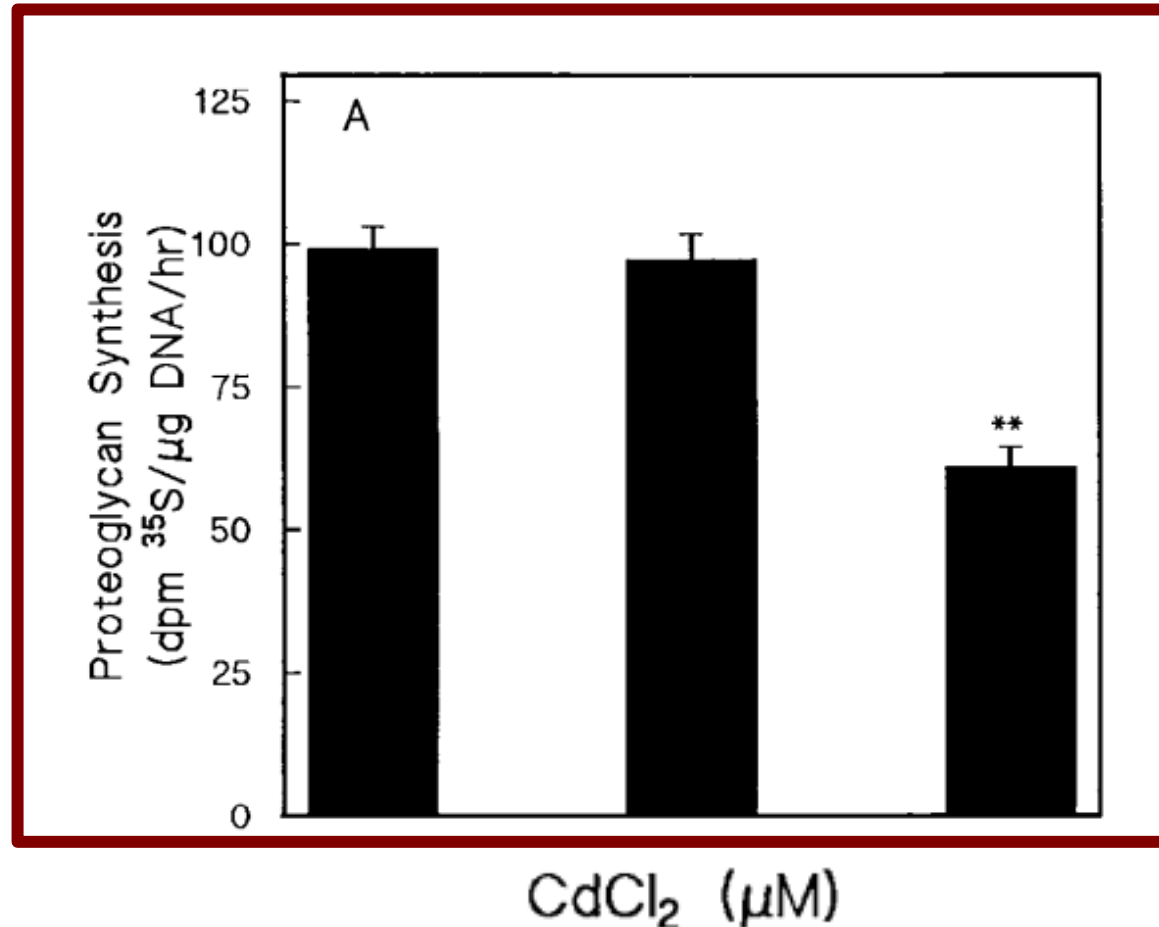
greatest in those with the highest cumulative cadmium exposure or liver cadmium. The difference in the transfer coefficient (KCO) between cadmium workers and referents increased linearly with increasing cumulative exposure without evidence for a threshold. The

estimated mean decrement in KCO for a cadmium worker employed 5 or more years with a cumulative exposure of  $2000 \text{ yr} \cdot \mu\text{g} \cdot \text{m}^{-3}$  (exposure to the current UK control limit of  $50 \mu\text{g} \cdot \text{m}^{-3}$  for a working lifetime of 40 yr) lies between  $0.05$  and  $0.3 \text{ mmol} \cdot \text{min}^{-1} \cdot \text{kPa}^{-1} \cdot \text{l}^{-1}$  (95% confidence interval). This decrement is consistent with the functional and radiological changes of emphysema observed in this group of workers.

# Cadmium Inhibits Proteoglycan and Procollagen Production by Cultured Human Lung Fibroblasts

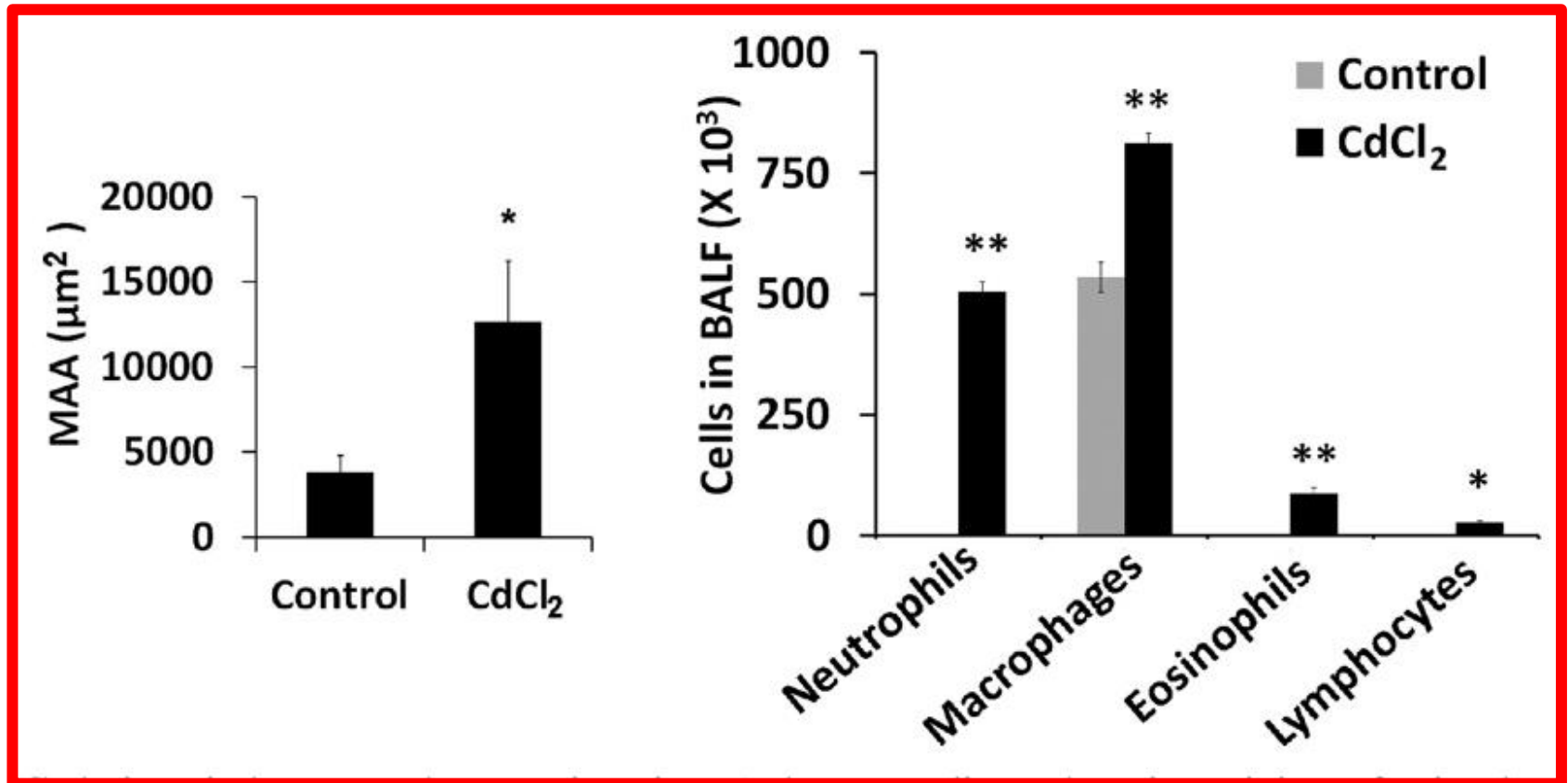
Rachel C. Chambers, Geoffrey J. Laurent, and Gunilla Westergren-Thorsson

Centre for Cardiopulmonary Biochemistry and Respiratory Medicine, University College Medical School, Rayne Institute, London, United Kingdom; and Department of Cell and Molecular Biology, Lund University, Lund, Sweden



Am. J. Respir. Cell Mol. Biol. 1998; 19:498–506.

## Heme oxygenase-1-mediated autophagy protects against pulmonary endothelial cell death and development of emphysema in cadmium-treated mice



» **Febbre da fumi metallici**

» **Polmonite**

» **Asma**

» **BPCO**

» **Fibrosi polmonare**

» **Tumore del polmone**



## Pulmonary fibrosis and exposure to steel welding fume

M. P. Cosgrove

**Table 2.** Histological findings

Paper, year of publication [reference]	Number of cases with histology	Number of cases with evidence of fibrosis	Alveolar macrophages containing welding fume	Perivascular fibrosis	Peribronchial fibrosis	Interstitial/alveolar fibrosis
Buerke, 2002 [7]	13	13	Yes	Yes	Yes	Yes
Müller, 2000 [8,33]	43	38	Yes	Yes	Yes	Yes
Radenbach, 1995 [26]	11	7	Not stated	Not stated	Not stated	Not stated
Rösler, 1995 [34]	6	6	Not stated	Yes	Yes	yes
Funahashi, 1988 [25]	10	10	Yes	Yes	Yes	Yes

# BMJ Open Effects of smoking, gender and occupational exposure on the risk of severe pulmonary fibrosis: a population-based case-control study

Magnus Ekström,<sup>1,2</sup> Torbjörn Gustafson,<sup>3</sup> Kurt Boman,<sup>3</sup> Kenneth Nilsson,<sup>3</sup> Göran Tornling,<sup>4</sup> Nicola Murgia,<sup>5</sup> Kjell Torén<sup>6</sup>

**Table 3** Dose-response effect of smoking on the risk of severe pulmonary fibrosis

Smoking, pack-years*	PF OR (95% CI)	IPF OR (95% CI)
0	1	1
1–9	1.03 (0.62 to 1.70)	0.90 (0.52 to 1.57)
10–19	2.26 (1.35 to 3.80)	2.10 (1.20 to 3.68)
≥20	2.66 (1.56 to 4.55)	2.25 (1.26 to 4.02)

ORs for levels of smoking estimated using conditional logistic regression adjusted for age and stratified for year of birth, year of diagnosis, gender and occupational exposure.

\*Pack-years of smoking up to 10 years before the year of PF diagnosis.

IPF, idiopathic pulmonary fibrosis; PF, pulmonary fibrosis.

**Table 2** Effect of smoking on the adjusted risk of pulmonary fibrosis, according to gender and occupational exposure

	PF OR (95% CI)		IPF OR (95% CI)	
	Women	Men	Women	Men
No occupational exposure	1.10 (0.50 to 2.42)	1.97 (0.64 to 6.13)	1.12 (0.49 to 2.59)	1.44 (0.43 to 4.83)
Occupational exposure	1.10 (0.52 to 2.34)	4.63 (2.08 to 10.33)	1.32 (0.58 to 3.03)	2.96 (1.34 to 6.52)

# Rischio di fibrosi e attività lavorative note per essere associate a IPF

Table 2. Job at risk for idiopathic pulmonary fibrosis, adjusted for gender, age and smoking and then stratified for gender.

	All	Men	Women
Exposure	OR (95% CI)	OR (95% CI)	OR (95% CI)
None	1	1	1
Any job at risk of IPF	<b>4.14</b> (2.27-7.53)	<b>4.40</b> (2.13-9.05)	<b>3.91</b> (1.18-12.86)
Costruction workers	1.31 (0.58-2.98)	1.37 (0.60-3.12)	-
Wood industry workers	1.36 (0.46-3.97)	1.35 (0.46-3.95)	-
<b>Metallurgical and steel industry</b>	<b>4.80</b> (1.50-15.33)	<b>4.76</b> (1.50-15.15)	-
Farmers, vets and gardeners	<b>2.73</b> (1.47-5.10)	<b>2.42</b> (1.14-5.11)	<b>3.91</b> (1.18-12.86)



# Pulmonary fibrosis and exposure to steel welding fume

M. P. Cosgrove

### Key points

- Health surveillance with spirometry is neither sensitive nor specific for detecting this condition and enquiry should also be made about dyspnoea and cough, which if present may need investigation despite normal spirometry, if the welder has been exposed to a high level of fume for a long period of time.
- Pulmonary investigation of welders in secondary care should include high-resolution computerized tomography.

» **Febbre da fumi metallici**

» **Polmonite**

» **Asma**

» **BPCO**

» **Fibrosi polmonare**

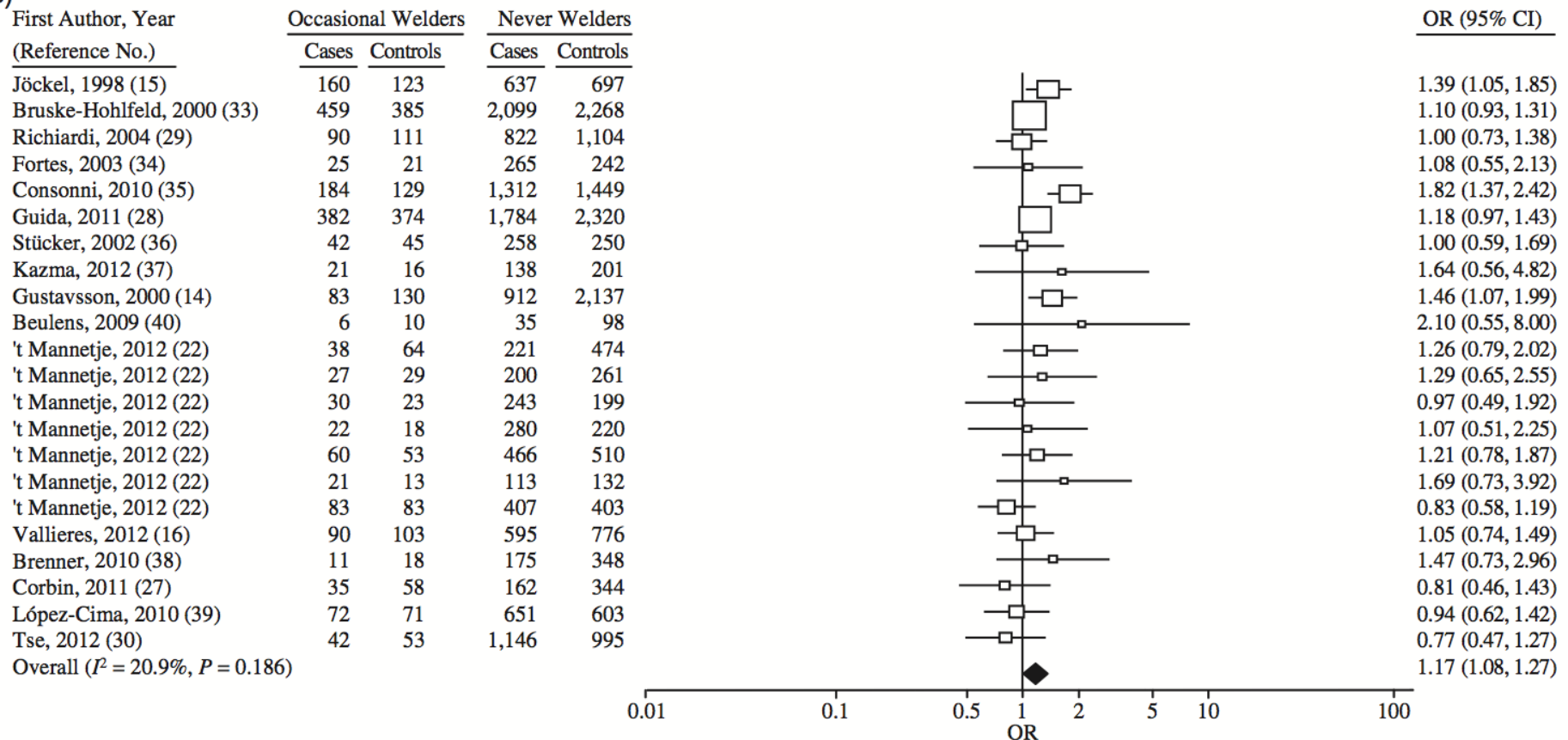
» **Tumore del polmone**



## Systematic Reviews and Meta- and Pooled Analyses

### Welding and Lung Cancer in a Pooled Analysis of Case-Control Studies

B)



# Lung cancer risk in welders and foundry workers with a history of heavy smoking in the USA: The National Lung Screening Trial

Jason Y Y Wong, Bryan A Bassig, Wei Jie Seow, Wei Hu, Bu-Tian Ji, Aaron Blair, Debra T Silverman, Qing Lan

	I) All lung cancer subtypes				
	Total cases	py	HR	95% CI Lower	95% CI Upper
Never welder or foundry worker	1824	300192.4	Ref		
Ever welder, never foundry worker	101	14056.5	1.12	0.91	1.37
Ever foundry worker, never welder	70	9715.2	1.09	0.85	1.39
Ever welder and foundry worker†	39	3870.6	1.48	1.08	2.04*



## Welding and Lung Cancer in Central and Eastern Europe and the United Kingdom

**Table 5.** Association Between Duration of Exposure to Welding Fumes and Lung Cancer, With and Without Chromium Exposure, Central and Eastern Europe and the United Kingdom, 1998–2001<sup>a</sup>

Exposures to Welding Fumes and Chromium	No. of Cases	No. of Controls	OR	95% CI
No welding fumes, no chromium	1,470	1,655	1.00	Referent
Welding fumes with chromium	190	159	1.34	1.04, 1.71
1–8 years	54	42	1.47	0.94, 2.30
9–25 years	64	52	1.28	0.85, 1.92
>25 years	71	65	1.27	0.87, 1.85
Welding fumes without chromium	393	363	1.14	0.95, 1.36
1–8 years	123	134	0.98	0.74, 1.30
9–25 years	117	120	1.00	0.75, 1.34
>25 years	153	109	1.48	1.11, 1.97
Chromium without welding fumes	144	118	1.32	1.00, 1.75

Abbreviations: CI, confidence interval; OR, odds ratio.

<sup>a</sup> Model adjusted for age, center, education, smoking, and asbestos, as well as silica exposure.

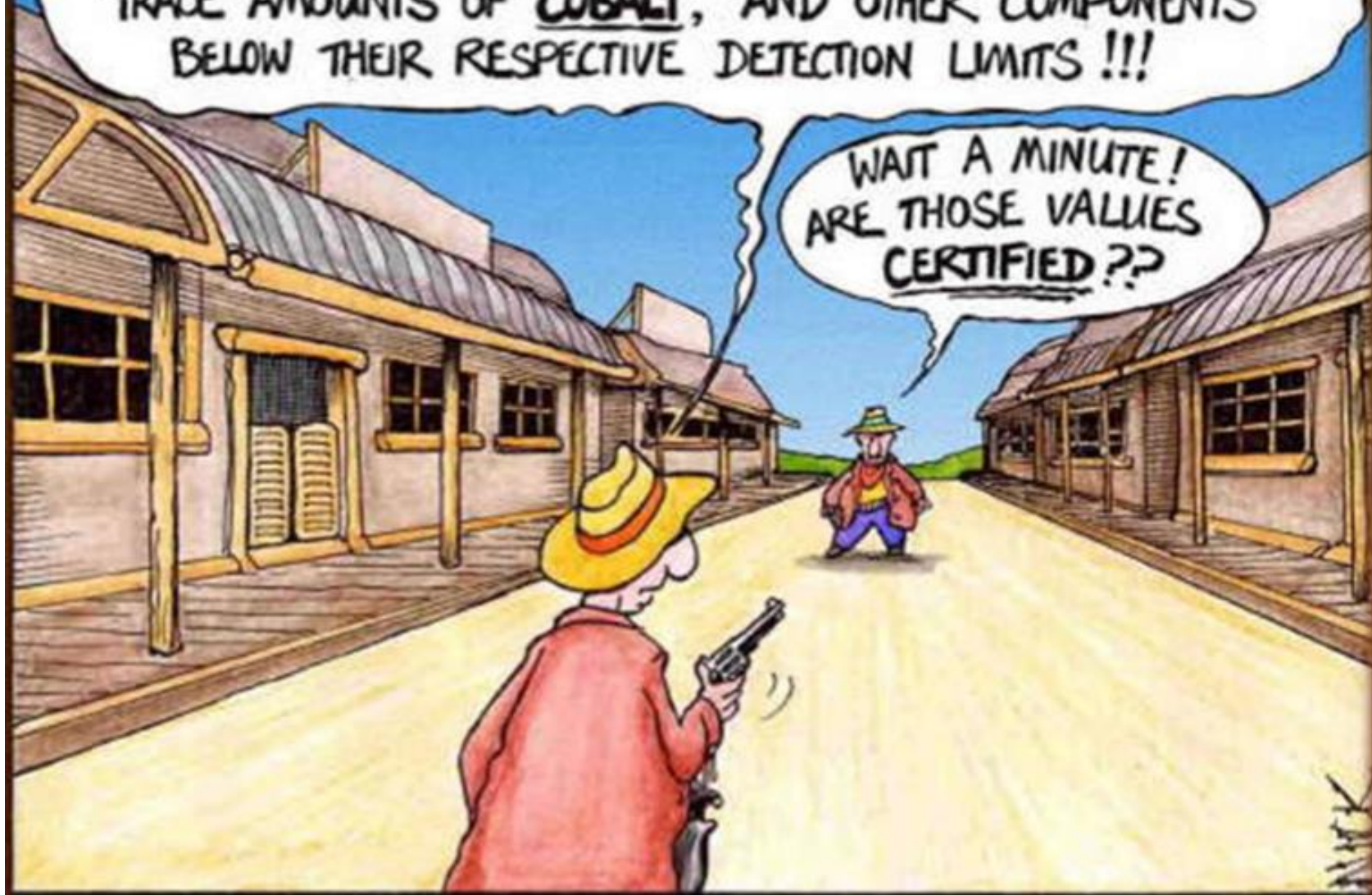
# Cancer Risks among Welders and Occasional Welders in a National Population-Based Cohort Study: Canadian Census Health and Environmental Cohort

Jill S. MacLeod<sup>1,\*</sup>, M. Anne Harris<sup>1,2,3</sup>, Michael Tjepkema<sup>4</sup>, Paul A. Peters<sup>5</sup>, Paul A. Demers<sup>1,3,6</sup>

	Welders	
	<i>n</i> = 12,845	
	Cases	HR (95% CI) <sup>†</sup>
Any cancer*	1,385	1.04 (0.99–1.10)
Lung	265	1.16 (1.03–1.31)
Mesothelioma	15	1.78 (1.01–3.18)
Stomach	45	1.25 (0.93–1.67)
Bladder	100	1.40 (1.15–1.70)
Kidney	60	1.30 (1.01–1.67)
Brain	35	1.16 (0.83–1.63)
Nasal	< 5	–
Ocular melanoma	5	1.55 (0.64–3.76)

DON'T MOVE OR I'LL FILL YOU FULL OF 98% LEAD,  
1% ANTIMONY, 0.75% SILVER, 200 PPM NICKEL, WITH  
TRACE AMOUNTS OF COBALT, AND OTHER COMPONENTS  
BELOW THEIR RESPECTIVE DETECTION LIMITS !!!

WAIT A MINUTE!  
ARE THOSE VALUES  
CERTIFIED??



**ANALYTICAL CHEMISTS IN THE WILD WEST**

Seminario per Medici Competenti

## Il rischio cancerogeno nelle lavorazioni con acciaio inox

Venerdì 20 ottobre 2017  
ore 8.45 - 13.15

Sala della Quercia  
Padiglione A. Ziccardi  
Via Amendola, 2 - Reggio Emilia



**Gli indicatori biologici di esposizione più comunemente  
impiegati e confronto con i valori limite e di riferimento**

**Massimo Corradi**



**UNIVERSITÀ  
DI PARMA**

DIPARTIMENTO DI MEDICINA E CHIRURGIA

**Reggio E, 20 Ottobre 2017**

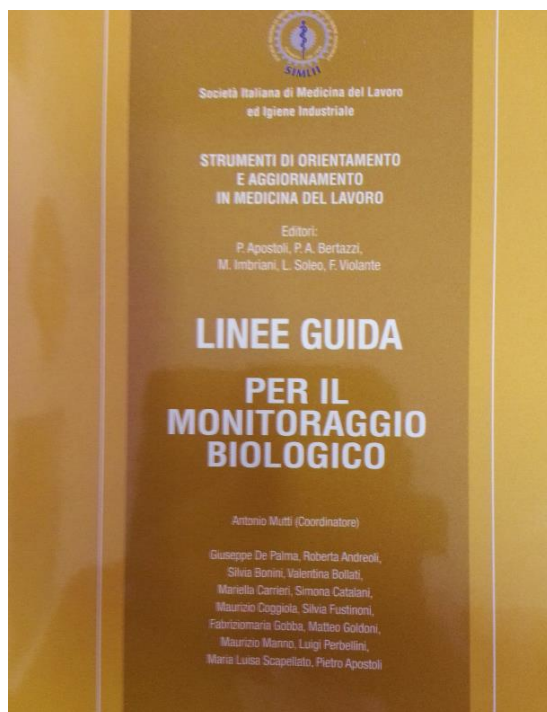


# Orientamento SIMLII

“La valutazione periodica dell’esposizione mediante *indicatori di dose interna e di effetto* è un compito di esclusiva pertinenza del medico competente, sia come pratica integrativa alla sorveglianza sanitaria, che come complemento alla valutazione del rischio ”

# Articolo 229 - Sorveglianza sanitaria

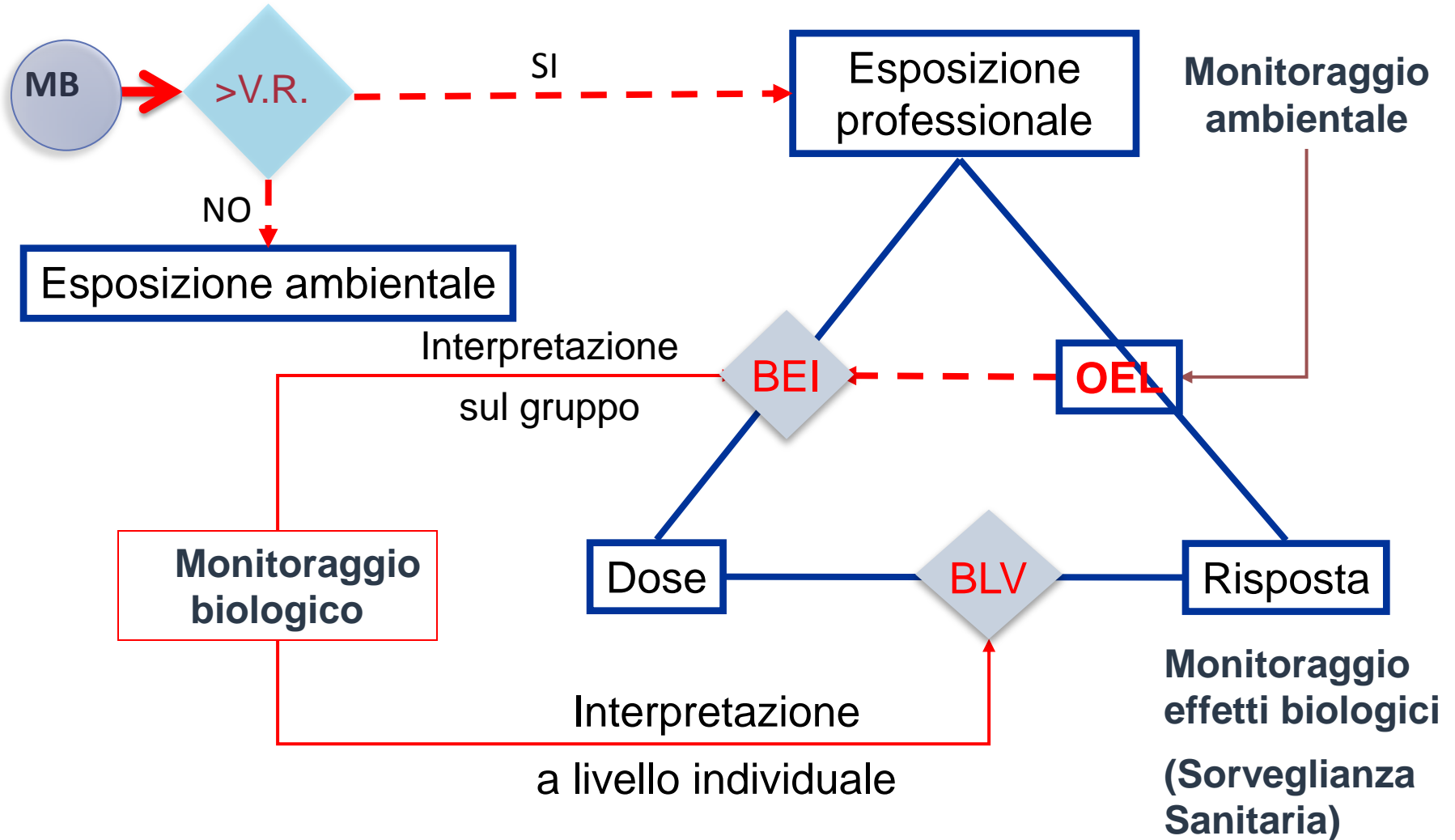
Il monitoraggio biologico è obbligatorio per i lavoratori esposti agli agenti per i quali è stato fissato **un valore limite biologico**.



Il valore limite biologico (VLB) è definito come "il limite di concentrazione del relativo agente, di un suo metabolita o di un indicatore di effetto nell'appropriato mezzo biologico". L'unico agente chimico per il quale esiste un VLB stabilito dalla legge è il piombo (allegato XXXIX D.lgs 81/2008), (vedi capitolo 10.1, sugli elementi metallici).

Nel processo di valutazione del rischio si afferma tuttavia il principio secondo il quale il monitoraggio biologico si debba estendere anche a tutti gli agenti chimici per i quali enti internazionali riconosciuti abbiano fissato dei valori limite (es i BEI dell'ACGIH, i BAT del DFG ecc, vedi in seguito) e non solo nei casi di esposizione ad agenti per cui esista un VLB (Tabella 1).

# Monitoraggio biologico





## **Position Paper**

**Criteria metodologici per la valutazione e proposta di valori-guida (VG) nazionali per il controllo dell'esposizione professionale a fattori di rischio chimici (valori-limite, livelli d'azione, valori di riferimento)**

A cura del  
**Sottogruppo di lavoro per la valutazione del rischio chimico (Sottogruppo-VRC)**

*M. Manno (Referente), M.C. Aprea, R. Bonfiglioli, A. Moretto, M.L. Scapellato*

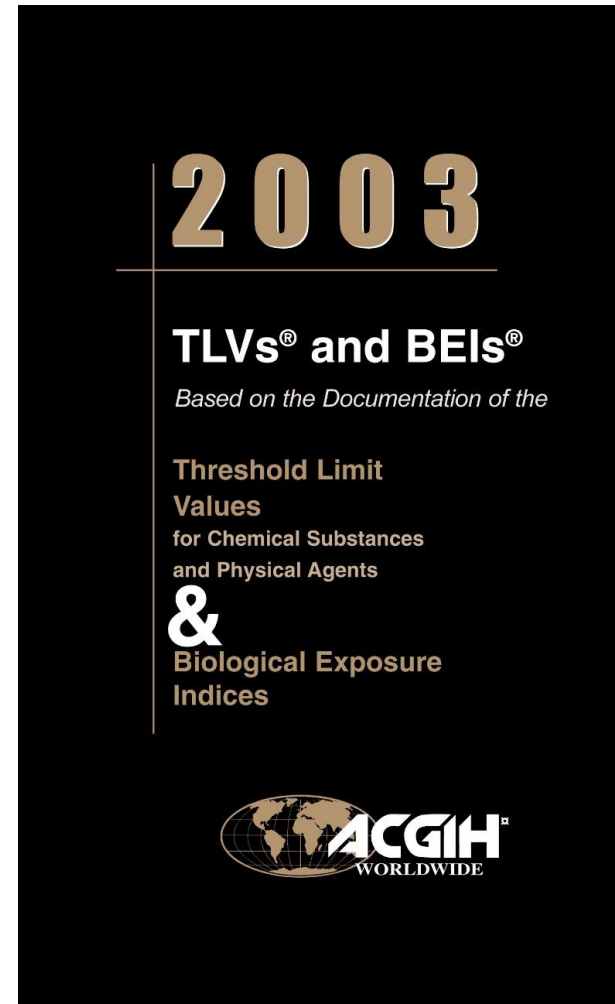
# VALORI DI RIFERIMENTO PER LA POPOLAZIONE GENERALE

Metalli	$\mu\text{g/l}$ (5° e 95° percentile)	$\mu\text{g/g}$ creatinina
Cr-U	0,05-0,35	< 0,2
Ni-U	0,1-5 $\mu\text{g/l}$	< 3,3
Cd-U	0,1 - 1,5	< 1



SIVR edizione 2011

- » **BEIs<sup>®</sup>** are intended for use in the practice of industrial hygiene as guidelines or recommendations to assist in the control of potential workplace health hazards.
- » Represent **mean levels** of determinants that are most likely to be observed in specimens collected from a healthy worker who **has been exposed to the TLV<sup>®</sup> -TWA.**



# Methodology for the Derivation of Occupational Exposure Limits

Scientific Committee on Occupational Exposure Limits  
(SCOEL)

'A **Biological Limit Value (BLV)** is a reference value for the evaluation of potential health risk in the practice of occupational health. [...] Exposure concentrations equivalent to the BLV generally do not affect the health of the employee adversely, when they are attained regularly under workplace conditions (8 hrs/day, 5 days/week), except in cases of hypersensitivity.'

Deutsche  
Forschungsgemeinschaft

**List of MAK  
and BAT Values  
2016**

***Occupational Exposure Limits*** — Three limit values are in use in Germany:

- Maximum Workplace Concentrations (“Maximale Arbeitsplatzkonzentrationen” – MAK)
- Technical Guidance Concentrations (“Technische Richtkonzentrationen” – TRK), and
- Biological Tolerance Value for occupational exposures (“Biologische Arbeitsstofftoleranzwerte” – BAT).

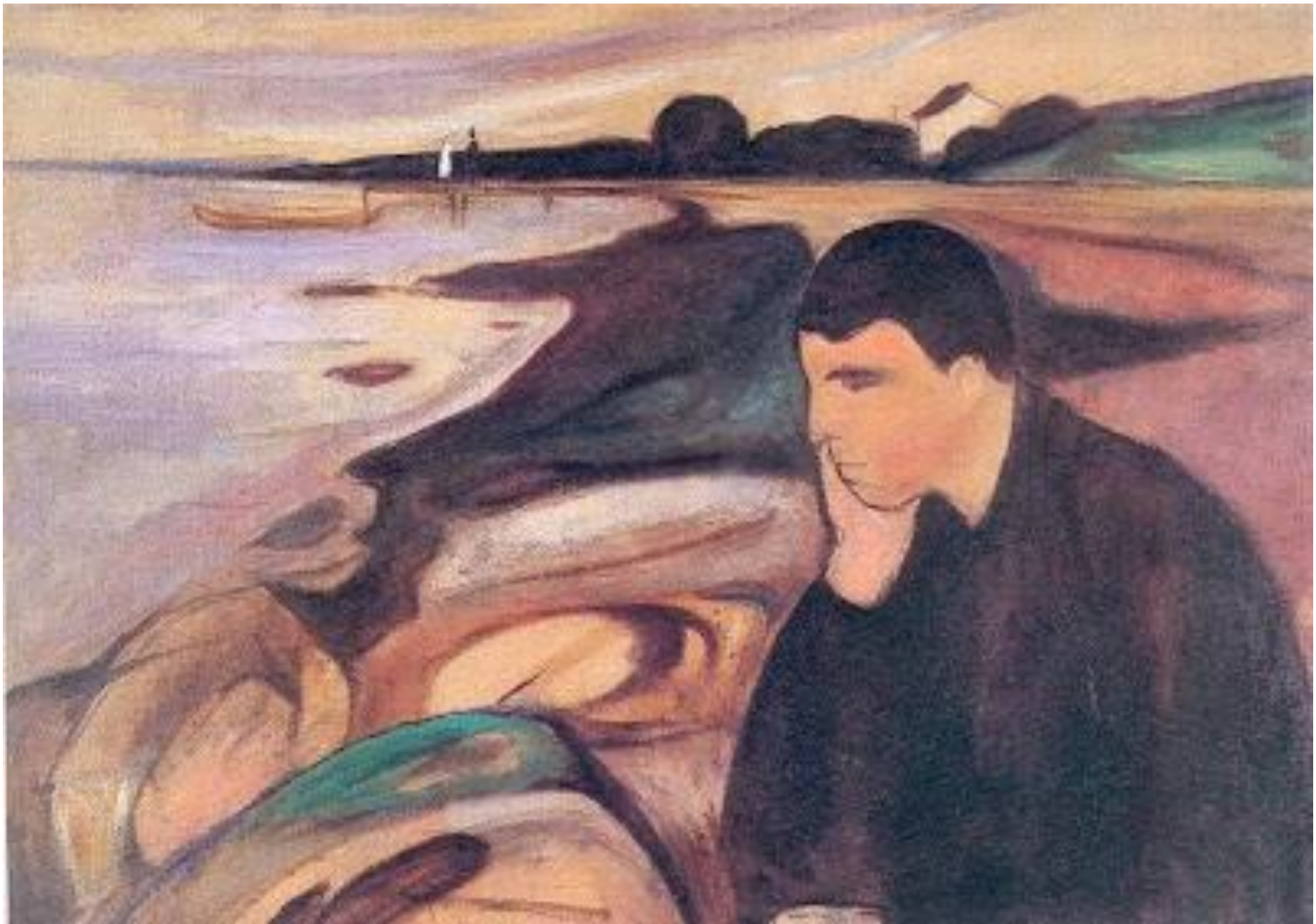
# Exposure assessment at the workplace: Implications of biological variability

Table 2

Comparison among different biological limits, e.g. BEI<sup>®</sup> (ACGIH), BAT (DFG), and BLV (SCOEL), in terms of their origin and interpretation

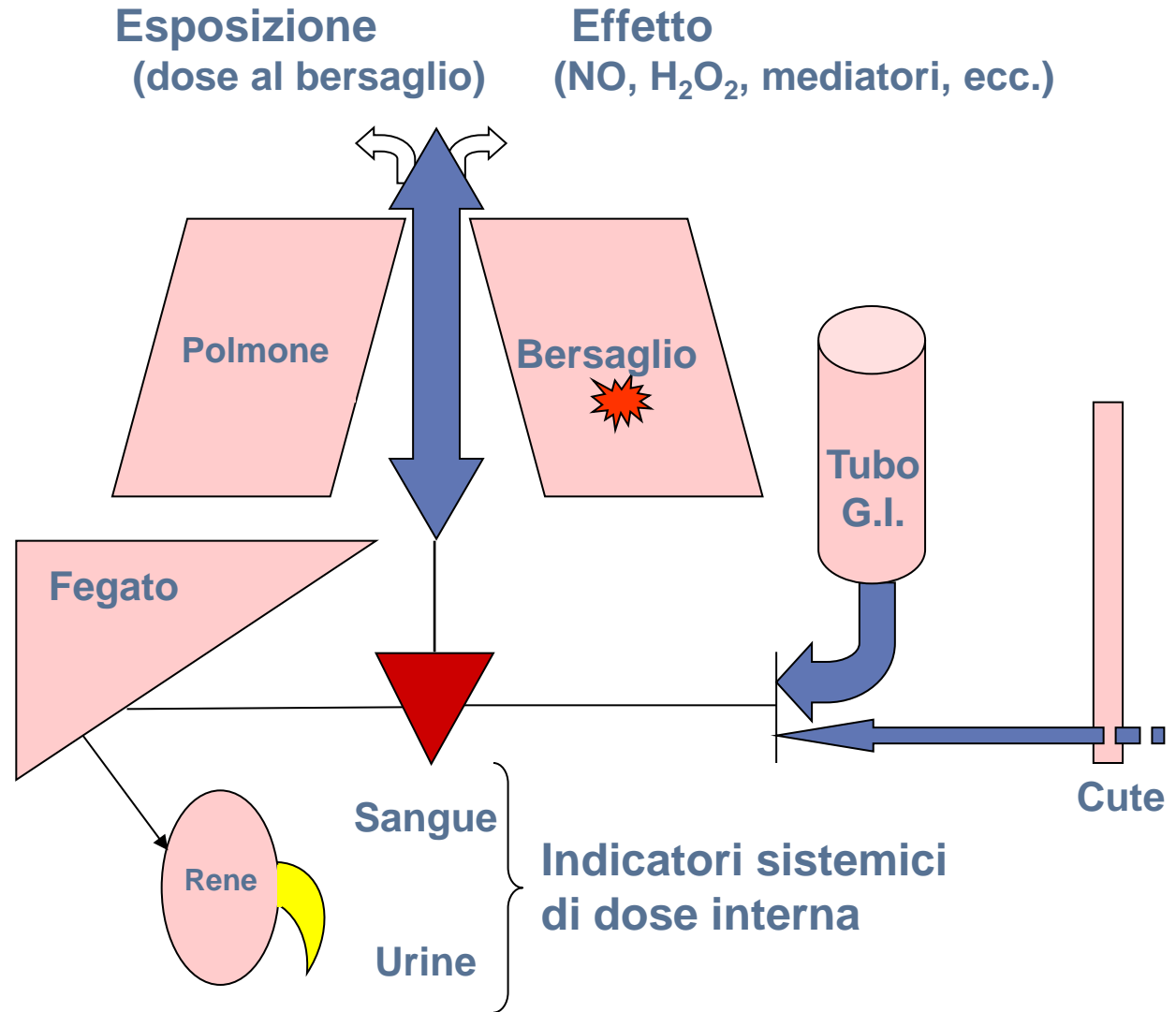
	BEI <sup>®</sup> (ACGIH)	BAT (DFG)	BLV (SCOEL)
Origin	Exposure-dose (dose-response <sup>a</sup> )	Dose-response or exposure-dose	Dose-response (OEL-dose)
Corresponds to	Mean value (NOAEL <sup>a</sup> )	Ceiling values	NOAEL, ceiling values
Interpretation	Groups (individuals <sup>a</sup> )	Individuals or groups	Individuals and groups
Criterion	TLV <sup>®</sup> -related (health-based <sup>a</sup> )	Health-based or MAK-related	Health-based
Carcinogens	Yes	No (EKA)	No



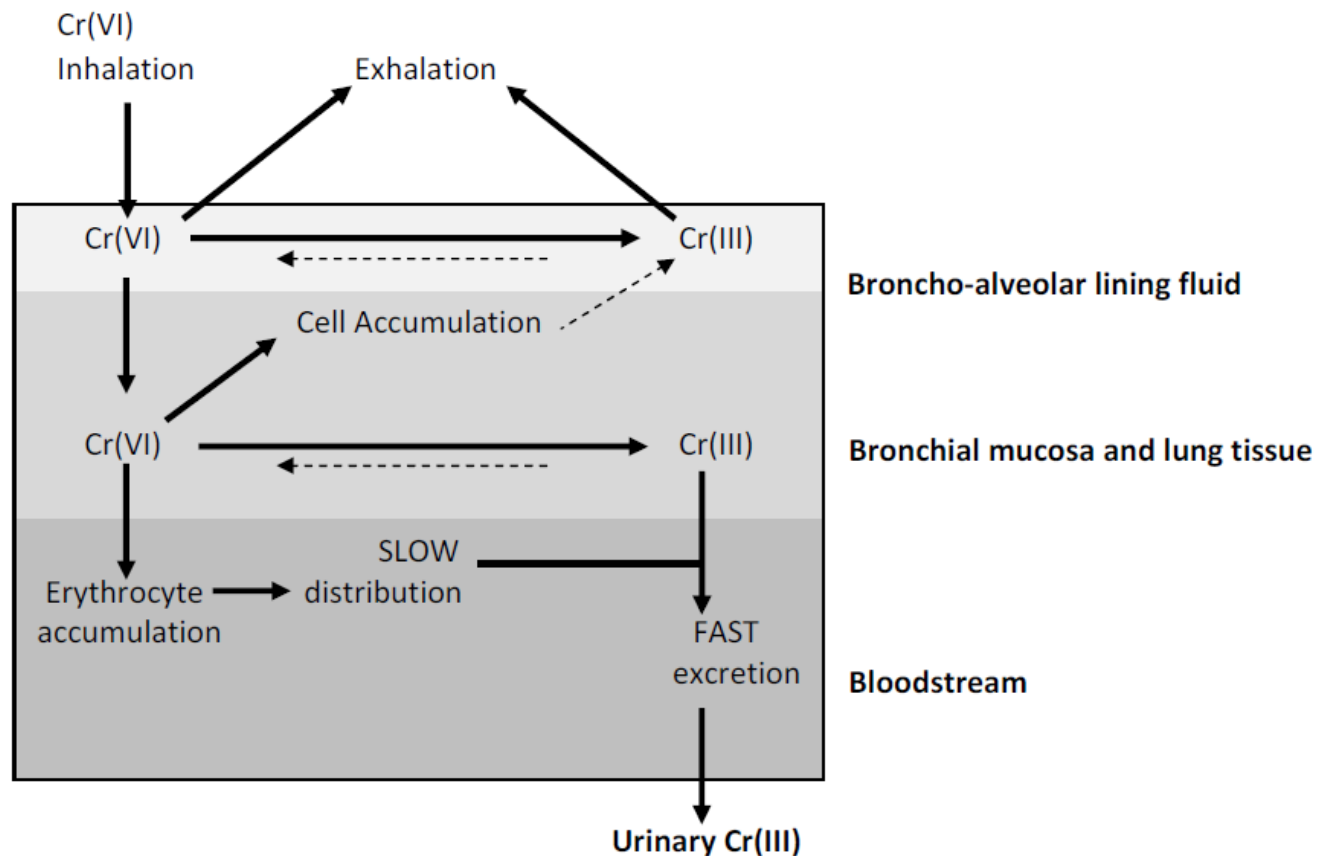


# Il significato hanno gli indicatori di dose

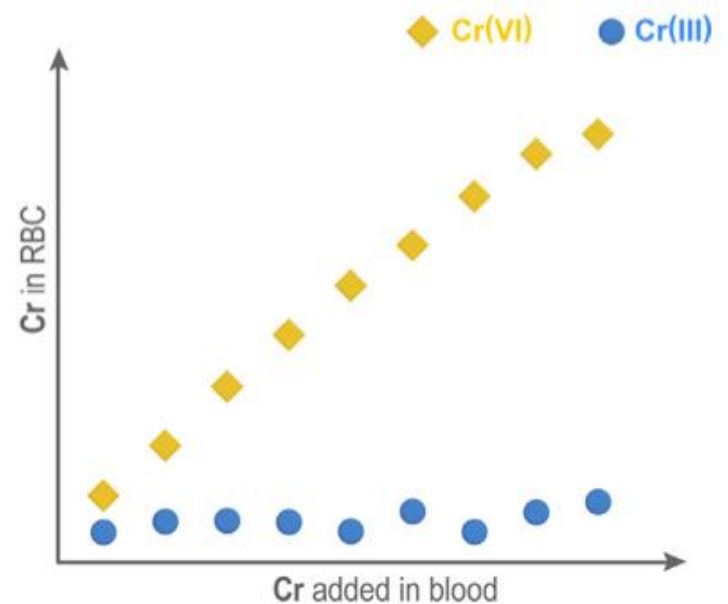
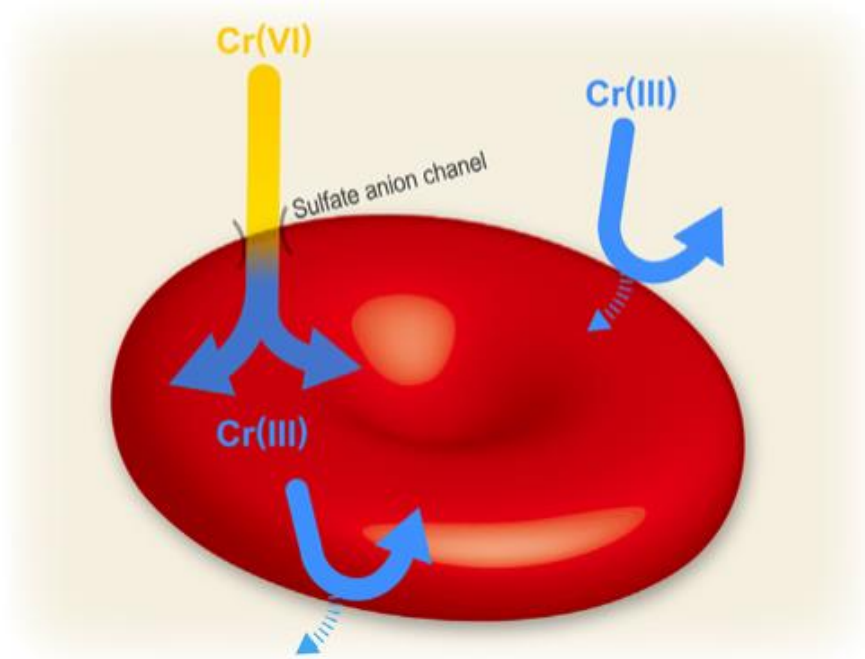
Gli effetti nocivi degli inquinanti sulle vie respiratorie possono essere dovuti alla loro **forte reattività** o **scarsa solubilità**. In entrambi i casi, la dose **ritenuta** (esposizione) sembra il determinante principale della pneumotossicità e del tumore polmonare



# Cinetica esposizione polmonare a cromo

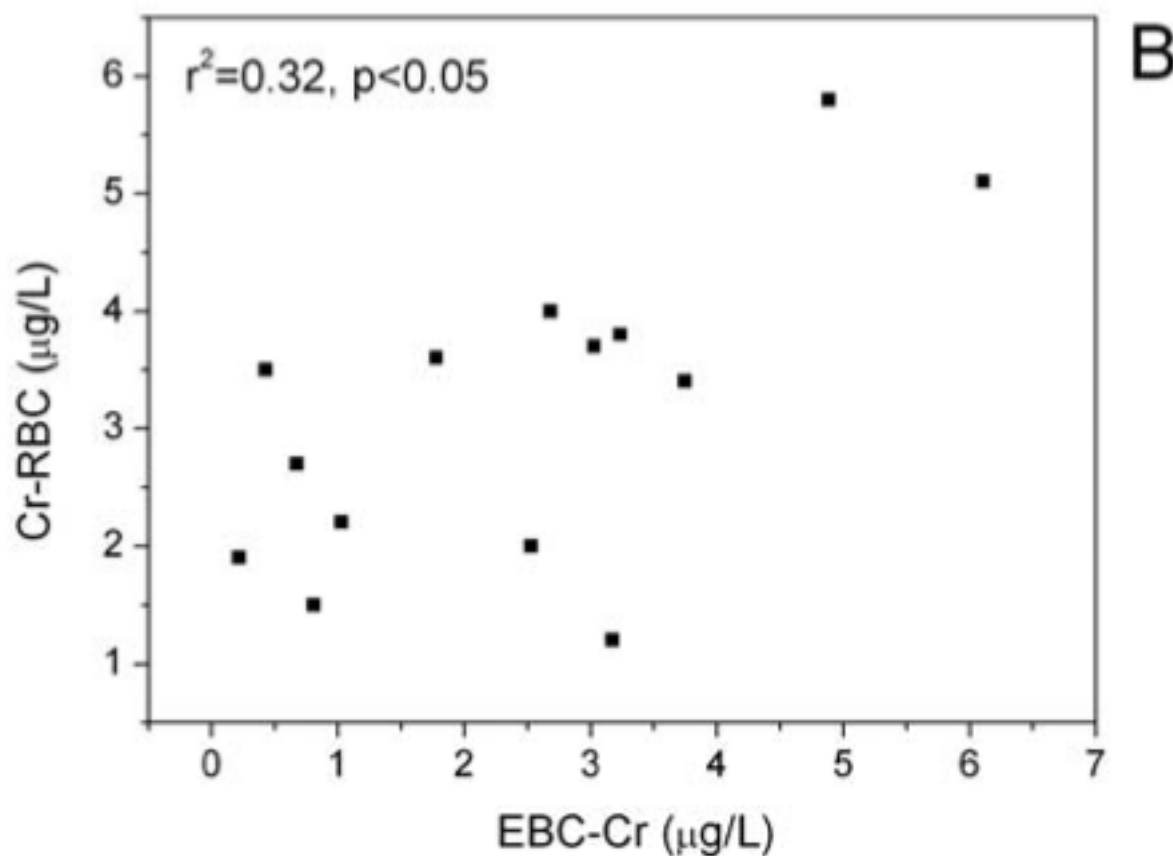


# Cr VI nei globuli rossi



# Chromium in exhaled breath condensate (EBC), erythrocytes, plasma and urine in the biomonitoring of chrome-plating workers exposed to soluble Cr(VI)

Matteo Goldoni,<sup>ab</sup> Andrea Caglieri,<sup>ab</sup> Giuseppe De Palma,<sup>c</sup> Olga Acampa,<sup>ab</sup> Petra Gergelova,<sup>d</sup> Massimo Corradi,<sup>a</sup> Pietro Apostoli<sup>c</sup> and Antonio Mutti<sup>\*a</sup>



# Risultati: Cromo nel CAE

Metalli	T <sub>0</sub>	T <sub>1</sub>	T <sub>2</sub>	p
Cr-CAE (µg/L)*	0.06 (< LoD-0.14)	0.08 (< LoD-0.22)	< LoD <sup>#</sup> (< LoD-0.10)	T <sub>2</sub> vs T <sub>1</sub> p < 0,0001 T <sub>2</sub> vs T <sub>0</sub> p = 0,023

Test non parametrico di Friedman seguito da test dei ranghi di Wilcoxon. \*I valori sono espressi come mediana (25°-75° percentile). <sup>#</sup>Lod: *Limit of detection*

**In 9 lavoratori che mostravano valori di Cr-CAE > 0.40 µg/L  
si è proceduto con la determinazione della frazione esavalente (Cr VI-CAE).**

**In 2 campioni sono emersi valori di Cr VI-CAE sopra il LOD (0.1 µg/L)**

Prepared for

**The American Welding Society**

**8669 NW 36th Street #130**

**Miami, FL 33166-6672**

Document type

**Final**

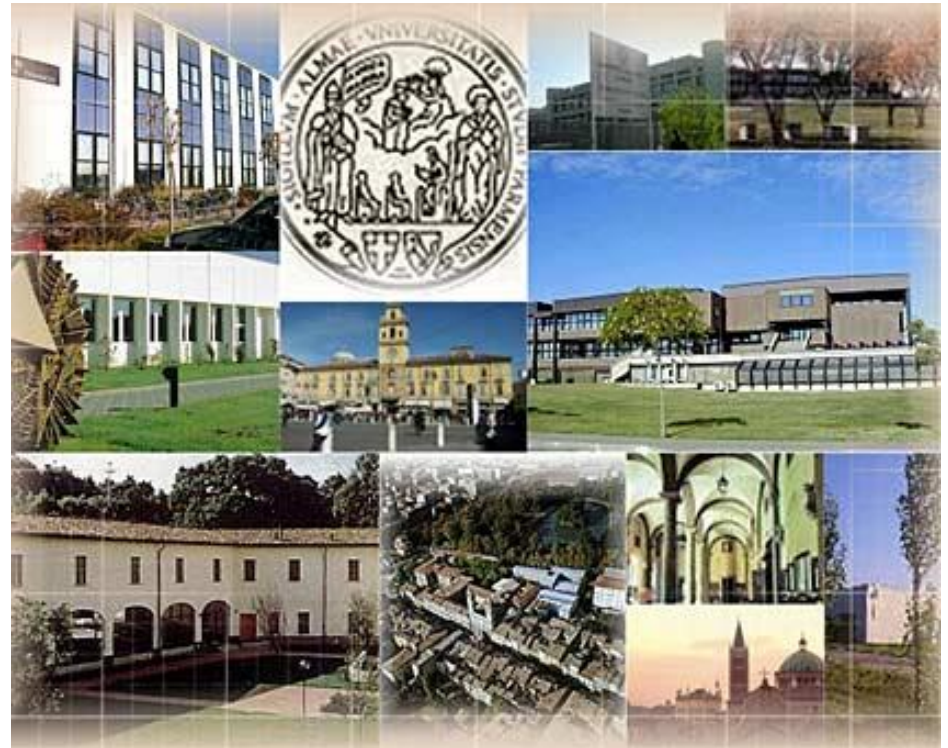
Date

**August 2017**

# **HEALTH EFFECTS FROM WELDING EXPOSURES: 2016 LITERATURE UPDATE**

# Ringraziamenti

- » Maria Grazia Riccelli
- » Andrea Sagramoni
- » Bruno Zoppi
- » Francesco Magnelli
- » Francesco Mantovani



**UNIVERSITÀ  
DI PARMA**

DIPARTIMENTO DI MEDICINA E CHIRURGIA