# STUDYING HEALTH IMPACT OF EXPOSURE TO ORGANIC SOLVENTS ON RENAL FUNCTIONS (A CASE CONTROL STUDY) CAIRO - EGYPT.

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#### BY

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# Acknowledgement

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to dedicate
this work purely
for the sake of Allah
in the hope that
it will be
useful for mankind

#### **Abstract:**

**Introduction**: the results of the studies performed to study the effect of exposure to organic solvents on kidney in workers occupationally exposed to organic solvents over the last twenty years are contradictory. **Subjects and methods:** we studied the effect of occupational exposure to organic solvents on renal functions using both routine renal functions namely; serum urea, serum creatinine, estimated glomerular filtration rate and early renal urinary biomarkers namely; N acetyl-B-D glucosaminidase and B2 microglobulin.

This is an analytical case control study conducted on 181 male worker in paint manufacturing factory exposed to organic solvents during their work and 186 control individual never exposed occupationally to organic solvents and engaged in administrative tasks in organization outside the factory. The results of the study revealed statistically significant difference between the Urinary –N-Acetyl glucosaminidase activity (NAG index) as well as B2microglobulin (Pvalue<0.001) in the workers that exposed to organic solvents and their matched controls. The proportion of exposed workers that have abnormal NAG activity (53.6%) is higher than that in their matched controls (29.6%) and represent about two fold increase in the activity with high statistically significant difference (pvalue=0.000). **Conclusion:** results of this study point to a possibility of early renal effects, but not to a serious to influence on the routine kidney function tests at the current levels of occupational exposure to organic solvents. Recommendations: workers exposed to paints, glues, degreasing solvents, and cleaning solvents must be evaluated periodically for renal effects using Urinary NAG excretion and B2microglobulin as reliable markers of the tubulo-toxicity and non-invasive tests.

#### **Key words:**

Organic solvents – renal impairment- early renal biomarkers – NAG -B2M.

## **Contents**

Acknowledgment			
Contents			
List of Tables			
List of Figures			
List of Abbreviations			
Introduction	1		
Aim of the Study	5		
Review of literature	6		
Chapter I: chronic kidney disease	6		
Stages of chronic kidney disease	7		
Magnitude of renal disease	7		
Risk factors of chronic kidney disease	9		
Prevention of chronic kidney diseases	10		
Early detection of renal disease			
Routine laboratory renal function tests			
Early markers for renal disease			
Chapter II: Organic solvents	21		
Public health importance of organic solvents	22		
Frequency of exposure to organic solvents			
Populations at Increased Risk of Solvent-Related Diseases			
Exposure to organic solvents in different types of industry	25		
Routs of Exposure to organic solvents	26		
Chapter III: Health hazards of solvent exposure	28		
Effects on the Central Nervous System	30		
Effects on the Peripheral Nervous System	32		
Effects on the Liver	32		
Cardiovascular Effects.	33		
Dermal Effects	34		
Reproductive and Developmental Effects	34		
Carcinogenicity	35		
Specific organic solvents used in paint industry	36		
Exposure to multiple solvents	41		

Chapter IV: Health hazards of organic solvents on renal system	42
Mechanisms of solvent- induced nephrotoxicity	53
Preventive strategies to reduce solvent induced nephrotoxicity	54
Subjects and Methods	57
Results.	64
Discussion	98
Conclusion.	114
Recommendation	115
Summery	116
References.	120
Appendices	140
الملخص العربي	148

## List of tables

Table1: Socio demographic characteristics of the exposed workers to organic solvents.	
Table2: Occupational characteristics of the exposed subjects to organic	68
solvents Participated in the study.	
Table3: Comparison between the workers and control groups regarding to	70
their educational status.	
Table4: Comparison between the exposed and control groups in respect to	71
their smoking status.	
Table5: Comparison between the exposed and control groups according to	71
their family history of renal failure.	
Table 6: Comparison between the exposed and control groups according to	72
their history of hypertension.	
Table 7: Comparison between the exposed and control groups regarding to	72
their history of diabetes mellitus	
Table 8: statistical comparison between the exposed workers and their	
matched controls in respect to some personal and medical factors.	
Table 9: Distribution of workers activities in the factory as regards to their	
renal function	
Table 10: Statistical comparison between the renal functions of workers and	75
control groups.	
Table 11: Distribution of normal and abnormal renal functions among the	
studied groups.	
Table 12: Statistical comparison between the renal function of exposed and	
control groups regarding their smoking status.	
Table 13: Statistical comparison between the renal function of workers and	79
control group as regard their smoking grade.	
Table 14: Statistical comparison between the renal function of exposed and	80
control groups regarding their hypertensive status.	
Table 15: Statistical comparison between the renal function of exposed and	81
control groups regarding their diabetic status	
Table 16: Statistical comparison between the renal function of exposed and	82

control groups regarding their body mass index	
Table 17: Statistical comparison between the renal functions of the exposed	83
workers to organic solvents according to their duration of exposure	
Table 18: Correlations between age and duration of exposure with the renal	83
functions in the workers.	
Table 19: correlations between the renal functions in the workers	84
Table 20: Distribution of workers activities in the factory as regard the	88
activity of their renal biomarkers.	
Table 21: Statistical comparison between the exposed and control groups	89
regarding the activity of urinary renal biomarkers.	
Table 22: Distribution of normal and abnormal activities of studied renal	90
biomarkers among studied groups.	
Table 23: Statistical comparison between the activities of the studied urinary	91
renal biomarkers in exposed and control groups regarding their smoking	
status.	
Table 24: Statistical comparison between the studied renal biomarkers of	92
exposed and control groups as regard their smoking grade	
Table 25: Statistical comparison between the studied renal biomarkers of	93
exposed and control groups according to their hypertensive status:	
Table 26: Statistical comparison between the renal function of exposed and	
control groups according to their diabetic status	
Table 27: Statistical comparison between the renal function of exposed and	
control groups regarding their body mass index.	
Table 28: Statistical comparison between renal biomarkers activities of	95
workers as regards to different durations of exposure	
Table 29: Correlations between age and duration of exposure with the renal	96
biomarkers in the workers	
Table 30: Correlation between the studied renal biomarkers in the workers	96
Table 31: Correlations between the studied renal biomarkers and renal	97
function tests in the workers	

## List of figures

Figure I: Industrial uses of organic solvents	
Figure 1: Working activities in pachin factory	69
Figure 2: Comparison between the serum levels of renal function tests of	
exposed workers and control group	75
Figure 3: Percentage of normal & abnormal renal function tests in exposed and	
control groups	76
Figure 4: Correlation between age and GFR in exposed workers and control	
groups	84
Figure 5: Correlation between the duration of exposure of workers exposed to	
organic solvents and GFR	
	85
Figure 6: Correlation between serum creatinine and GFR in exposed and	
control groups	86
Figure 7: The activity of early urinary renal biomarkers in exposed and	
control groups	89
Figure 8: Distribution of normal & abnormal activities (%) of studied early	
renal biomarkers in workers and control groups	90
Figure 9: Correlation between the studied renal biomarkers in the exposed	
and control groups	96

### **Abbreviations list**

AAD	Alonina amina pantidasa
AAP	Alanine amino peptidase
ACGIH	American Conference of Governmental Industrial Hygienists
ALP	Alkaline phosphatase
ANOVA	Analysis Of Variance
ATN	Acute tubular necrosis
BMI	Body Mass Index
B2M	B2-Microglobulin
BUN	Blood urea nitrogen
CCIE	Chamber of Chemical Industries – Egypt
CDC	Centers for Disease Control and Prevention
CNS	Central nervous system
CRF	Chronic renal failure
ESRD	End stage renal disease
EEG	Electroencephalography
GBM	Glomerular basement membrane
GFR	Glomerular filtration rate
GN	Glomerulonephritis
IARC	International Agency for Research on Cancer
IPCS	International program on chemical safety
MDRD	Modification of Diet in Renal Disease
MEK	Methyl ethyl ketone
MN	Membranous nephropathies
MNBK	Methyl-n-butyl ketone
NAG	N-acetyl-B-D-glucosaminidase
NKF	National Kidney Foundation
NIOSH	National Institute of Occupational Safety and Health
OMAL	Ontario Association of Medical Laboratories
OSHA	Occupational Safety Health Administration
PCE	perchloroethylene
PPE	personal protective equipment
TCA	Trichloroethane
TCE	Trichloroethylene
TLV	The Threshold Limit Value

#### Abbreviations list

TWA	time weighted average

#### Introduction

Chronic kidney (CKD) disease is a major public health problem throughout the world and the prevalence of CKD is largely sustained by the inclusion of a substantial proportion of the elderly population within stage 3 CKD, according to the Kidney Disease Outcomes Quality Initiative staging system. Adverse outcomes of chronic kidney disease can be prevented through early detection and treatment (Mangione & Dal Canton, 2011).

It is clear that an individual's likelihood of developing progressive Chronic kidney disease results from complex interactions between multiple genetic (none modifiable factors) and environmental factors (modifiable factors) (Satko, et al, 2007). Identification of the modifiable and controllable risk factors e.g. exposure to nephrotoxic substances such as organic solvents, is an important first step in understanding and hopefully, revering the increasing incidence of such disease (Thoenen, 2006).

A solvent is a substance that dissolves another substance forming a solution. Solvents that contain carbon are known as organic solvents and can contain chemicals considered hazardous (**Brautbar**, 2008).

The organic Solvents represent an important group of environmental pollutants to which people are exposed daily in the household settings and workplace They are present as ingredients in paints, varnishes, lacquers, glues, adhesives, degreasers, cleaners and in the production of dyes, polymers, plastics, textiles, printing inks, agricultural products and pharmaceuticals (Bale, et.al,2011). Organic solvents are present also in detergents (citrus terpenes), in perfumes (ethanol), home deodorizers, toothpaste (formaldehyde), carpets cleaners, nail polish removers, bathroom cleaners and glue solvents (acetone, methyl acetate, ethyl acetate), in dry cleaning (e.g. tetrachloroethylene), paint thinners (e.g. toluene, turpentine), in spot removers (e.g. hexane, petrol ether), furniture oils, shoe care products (Andrews, 1989).

Exposure can occur through inhalation due to volatilization of the solvent. Primary exposure concerns result from inhalation exposure from occupational work such as metal degreasing operations and/or residential exposures resulting from off gassing from the factories using these agents or from groundwater contamination. Many varied health effects including toxicity to the liver, kidney, lungs, and carcinogenic effects have been reported in the literature. ( **Bale, et al, 2011**). Millions of workers are exposed to organic solvents worldwide at different workplaces. Depending on the vapor pressure of the individual solvents, workers may be exposed by inhalation. Exposure concentrations may be very high, at least for some procedures (e.g. cleaning). In addition, dermal exposure may contribute considerably to the overall exposure. Exposure may occur despite the use of protective clothing, because solvents may penetrate gloves or cloth. Solvent mixtures are frequently used. Therefore, effects may also be caused by combined exposures. Threshold limits in the air, established by several national and international institutions, enable control of a large number of solvents in air at the workplace. (**Mangelsdorf, 2009**).

Among the health hazards of exposure to organic solvents, are renal problems. The severity of these effects is largely dependent on the compound and the level of exposure. (Gunnar et al, 2008).

In recent years case reports, case studies and experimental animals studies demonstrated that solvents are causing a variety of kidney disorders including acute and chronic kidney failure. The major routes of absorption of these compounds are through the lungs and the skin. Once in circulation, these compounds are concentrated in the kidney by mechanism remains uncertain but may be related to a solvent-induced disruption of the structural and functional integrity of the limited membranes of the kidney cells. A number of studies have demonstrated a significant association between organic solvents exposure and the development of glomerulonephritis through immune mediated mechanisms. (**Brautbar**, 2008).

Workers with a history of exposure to paints, glues, degreasing solvents, and cleaning solvents must be evaluated for renal impairment. Such patients may not know the risks of their exposure and may actually not understand the relationship between an