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# **Original Article**

# An Assessment of the Modulatory effect of Zinc on Humoral Immune Parameters in Welders and Cement Block Moulders in Ogbomosho, Nigeria

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#### **ABSTRACT**

Welding and cement block moulding (CBMs) are increasing in the developing countries due to increase in small scale industries resulting in excessive exposure to lead (Pb) and cadmium (Cd). Welding is associated with simultaneous release of zinc (Zn) fumes unlike cement block moulding. Unlike Cd and Pb, zinc enhances immune response while Pb and Cd are immunosuppressants. The possible modulation by Zn of the immunosuppressive effect of Pb and Cd is unexplored. Thirty welders, thirty CBMs and thirty unexposed participants were studied. Cadmium, Pb and Zn were determined using atomic absorption spectrophotometry technique. Immunoglobulins, IgA, IgG and IgM were assayed using ELISA. Total lymphocyte count (TLC) and monocyte (TMC) were determined using standard haematological methods. Lead and Cd were significantly higher in welders and CBMs than in controls. Cadmium was significantly higher in welders than in CBMs Interestingly, IgG and IgM in welders were significantly higher than in CBMs; IgG being approximately two fold in welders. Monocytes were significantly higher in CBMs than in welders. These data suggest antagonism of Pb and Cd to Zn by the lower levels of zinc in the exposed groups and the humoral immunomodulatory effect of zinc, by the higher levels of immunoglobulins in welders with higher zinc levels

Key words: Cadmium, Lead, Zinc, Immunomodulation, Welders, Cement block moulders.

#### RÉSUMÉ

Les activités du soudage de métaux ainsi que celles du moulage en ciment pour la fabrication des blocs de construction se multiplient dans les pays en développement suite à une augmentation du nombre des petites et moyennes entreprises. Ces activités aboutissent à une libération exagérée du plomb (Pb) et du cadmium (Cd). En plus, le soudage est associé à une libération simultanée des vapeurs du Zinc (Zn) contrairement au moulage en ciment. Zinc améliore la réponse immunitaire des individus pendant que les Pb et Cd sont des immunosuppresseurs. La modulation possible par Zn de l'effet immunosuppresseur du Pb et du Cd est inexplorée. Trente soudeurs, trente fabricants du moulage en ciment et trente participants non exposés (témoins) ont été étudiés. Les taux de Cd, Pb et Zn ont été mesurés à l'aide de la technique de spectrophotométrie d'absorption atomique. Immunoglobulines : IgA, IgG et IgM ont été analysées en utilisant ELISA. Le nombre total des lymphocytes et celui des monocytes ont été déterminés à l'aide de méthodes hématologiques standard. Plomb et Cd ont été significativement plus élevés chez les soudeurs et les fabricants du moulage en ciment que chez les témoins. Cadmium était significativement plus élevé chez les soudeurs que chez les fabricants du moulage en ciment contrairement à Pb. Remarquablement, Zn était significativement plus élevé chez les soudeurs que chez les fabricants du moulage en ciment. Faits intéressants, IgG et IgM chez les soudeurs étaient significativement plus élevées chez les fabricants du moulage en ciment que chez les soudeurs. Les résultats suggèrent l'antagonisme du Pb et du Cd au Zn avec des niveaux inférieurs du Zn dans les groupes exposés ainsi que les effets immunomodulateurs du Zn observés à travers des taux très élevés d'immunoglobulines chez les soudeurs ayant des concentrations importantes du Zn.

Mots clés: Cadmium, Plomb, Zinc, Immunomodulation, Soudeurs, fabricants du moulage en ciment.

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#### **INTRODUCTION**

n an attempt to raise the standard of living and improve their economies, developing countries are emphasizing both small and medium scale enterprises (SME) (1) on the understanding that SMEs play a substantial role in employment generation and national development (1). Welding and cement block moulding are among the most prevalent of these small-scale industrial activities but associated with excessive occupational exposure to toxic substances such as Pb, Cd and many others, which occur due to lack of proper workplace control measures.

The upsurge observed in welding and cement block moulding activities in these countries may be related in part to the relatively low level of education required in these occupational activities. A country like Nigeria is a good case in point to assess the potential human health effects posed by these two occupations where there may be exposure to toxic metals with one also associated with high degree of exposure to an essential metal such as is seen in welding where there is simultaneous exposure to both toxic metals and the essential metal, zinc (Zn) which has substantial effect on immune function.

Welding involves joining metals at very high temperature using electrodes which improve the joining process, leading to release of fumes rich in both toxic metals and zinc (2). The fume consists of very small particles, including ultrafine matter (2). Exposure to welding fumes is affected by several factors, including the welding process itself, work place characteristics, and use of protective measures (3).

Welding fume generated during the welding process contain at least 13 metals, including manganese (Mn), beryllium (Be), cadmium (Cd), chromium (Cr), cobalt (Co), copper (Cu), iron (Fe), lead (Pb), mercury (Hg), molybdenum (Mo), nickel (Ni), zinc (Zn), antimony (Sb), and vanadium (V) (4). Of these metals, lead, cobalt, chromium (IV) and Cadmium have been found to have critical adverse effect on the immune system (5, 6, 7).

Several researchers have shown that inhalation of cement dust containing a number of toxicants perturb both haematological and immunological parameters (8, 9). Ogunbileje et al. (10) demonstrated that metals such as lead, mercury, cadmium, chromium, nickel, copper, zinc, calcium, iron, manganese and chromium (VI) are present in cement dust.

Although certain properties differentiate between welding fumes and cement dust (aerodynamic size), there is the likelihood that both may have common effects and mechanism of toxicity on humoral immune system due to relative similarity in their immunotoxic metal composition (4, 11). Studies by Steel and Sanderson (1966) and Noweir (1988), show that metal composition of welding fumes and

cement dust are relatively similar with higher zinc content in welding fumes than in cement dust (12, 13).

Growing studies on metals have revealed numerous essential bio-metal functions and presence of excess biologically active metals can disrupt the normal physiological processes in the body. Chemical elements such as Zn are essential for life while Cd and Pb have no known beneficial effects in animals and humans. Cadmium, the common component of welding fumes and cement dust is confirmed to be carcinogenic, and exhibit toxic effects on many body systems, including the immune, nervous, endocrine, renal, musculoskeletal, and cardiovascular systems (14). In vivo studies have shown that Pb is an immunotoxicant, depressing humoral immunity (15,16), increasing host susceptibility to bacterial (17,18) and viral infections (19).

While lead and cadmium impair immune system (5, 6, 7), Zn at physiological dose plays a vital role in enhancing immune system. It is known that zinc affects multiple aspects of the immune system, from the barrier of the skin to gene regulation within lymphocytes (20). Zinc is crucial for normal development and function of cells involved in nonspecific immunity such as neutrophils and natural killer cells (20). Zinc also affects the development of acquired immunity by affecting certain functions of T lymphocytes such as activation, Th1 cytokine production, and B lymphocyte development (20). Similarly, lymphocyte development and antibody production, particularly immunoglobulin G, depend on physiological role of Zn (21). The activities of macrophage, a pivotal cell in many immunologic functions, such as intracellular killing, cytokine production and phagocytosis require Zn (21). Inhalation of Zn fumes during welding of galvanized metals is analogous to Zn supplementation above the recommended daily allowance in welders, while in block moulding there is little or no Zn or lower Zn exposure due to its very low concentration in cement dust. The implications of this supplementation and its consequent effect on humoral immunity based on the physiological role of Zn have not been well explored.

In this study, we investigated the humoral immune response to welding fume laden with Pb and Cd but also rich in Zn and cement dust exposure which is also rich in Pb and Cd with relative paucity of the immune response enhancing prime micronutrient Zn. The immunological response was assessed by examining the levels of immunoglobulins (IgG, IgA and IgM), total lymphocyte count, and total monocyte count between the exposed groups and the control group. Data were further stratified by zinc status to determine whether zinc modified the immunological parameters. This was conducted by comparing the immunological index in welder with those of cement block moulders with lower Zn level.

Previous study by Boshnakova et al. has shown that welding fume exposure significantly reduce IgG levels (22). Hanovcova et al. reported similar findings of alterations in certain parameters of humoral immunity which include decrease in IgG and IgA levels (23). In contrast, Tuschl et al. observed no effect on immunoglobulin levels in a study involving 30 welders who worked in a plant where levels of welding fumes typically exceeded Austrian Working Limits of 5mg/m3 (24). We hypothesized that Zn status may significantly modify immune function in welders with higher Zn levels.

# **MATERIALS AND METHODS**

# Selection of Subjects:

A total of ninety (90) participants were selected for the study consisting of sixty males both exposed groups (18-40 years): 30 welders and 30 cement block moulders (CBM) who have been occupationally exposed a for minimum of five years, were recruited and thirty (30) age and sex matched apparently healthy civil servants who were neither welders nor moulders of cement block served as control (18-42years). The participants were selected for the study after a detailed interview using a structured questionnaire with details about personal data and health, residence, surrounding environment, smoking habits, length of time in occupational and daily hour of exposure, age, body mass index (BMI) as well as activity of serum alanine aminotransferase to exclude liver pathology and serum creatinine level to exclude renal disease. These formed part of the eligibility or inclusion and exclusion criteria. As far as it could be ascertained no subject selected for control was occupationally or by hobby exposed to the metals of interest.

Prospective control subjects residing in polluted industrial areas where there could be possibility of increase passive exposure were also excluded from the study. Importantly also, vestigial passive exposure may be similar to both control and test group which cancels out. The socioeconomic status of participant was essentially between low and medium; most were uneducated and control subjects were also largely selected to match the study subjects.

Only subjects who were not smokers and who have no history or signs and symptoms suggestive of haematological, infectious, liver and kidney disease were eligible and recruited into the study. However, participants with symptoms that are common clinical manifestation among workers involved in dusty occupations such as coughing and sneezing were recruited into either of the exposed groups. This was established by a careful history and complete physical examination.

# Sample Collection & Laboratory Studies:

Blood sample (10ml) was drawn from the cubital vein of each participant. An aliquot of 3ml of blood sample was dispensed into lithium-heparin bottle for whole blood analyses of lead and cadmium. Another aliquot of 3ml of blood was dispensed into bottle containing EDTA for total lymphocyte and monocyte count, while 4 ml of blood were dispensed into plain tubes to yield serum for determination of total protein, albumin, immunoglobulins (IgG, IgM, IgA) as well as zinc level.

Lead, cadmium, and zinc were determined using flame Atomic Absorption Spectrophotometer (FAAS) (205 Buck Scientific, Germany). Blood lead and cadmium were determined by the method of Bannon et al. (25) and Lauwerys et al. (26) respectively, while plasma zinc was determined by the method of Smith et al. (27). Total protein (Tp) and albumin (Alb) were determined with standard biuret and bromocresol green methods (BCG) respectively while total globulin level was computed from the values of total protein and albumin. Total lymphocyte and monocyte counts were determined using Auto Haematology Analyzer (HA6700 Model, Hawksley and Son Ltd, London). Serum immunoglobulin levels of IgM, IgG and IgA were determined by ELISA technique using kits from

# Statistical Analysis:

Statistical analysis was performed using SPSS version 21. The results were reported as mean ± standard deviation. Shapiro–Wilk's and Komolgorov Smirnov tests were used to verify normality. Since most data were not normally distributed, non-parametric equivalent of their parametric test were used and the data concerned were ranked before the analysis. Either an analysis of variance or Kruskal–Wallis ANOVA test was used for multiple comparisons of data. Comparisons between the exposed groups, either of the exposed group and control group were conducted using the Student's t-test for parametric data and the Mann-Whitney test for non-parametric data. A value of p<0.05 was considered to be statistically significance.

#### **Ethical Considerations:**

The University of Ibadan/University College Hospital (UI/UCH) Joint Ethics Review Committee (UI/EC/05/01) approved this study. Additionally, written informed consent (approved by the UI/UCH Ethics Committee) was obtained from each subject.

## **RESULTS**

The age range of 90 study participants was 18-40 years and they were all males. Average work history was 5 years for welders and cement block moulders. Daily hour of occupational exposure to fumes and cement dust and indicators of nutritional status such as body mass index

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(BMI), total protein, albumin and globulin were similar among the studied population (Table 1). Table 2 summarizes the central elements in the study population. Cadmium as well as Pb levels were significantly higher in welders and cement block moulders than in control subjects, while zinc concentrations were significantly lower of Pb did not differ significantly between the two exposed groups. Cadmium level was significantly higher in welders than in CBMs. Remarkably the level of zinc was observed to be significantly higher in welders than in CBMs (p < 0.001) (Table 2). Total lymphocyte count was similar in both exposed groups (Table 3) but significantly higher in both exposed groups than in the control group (Tables 4&5). Total monocyte count was significantly higher in CBMs than in welder and control groups (Tables 3&5). Interestingly, the levels of the immunoglobulins (IgG, IgM and IgA) were observed to be higher in welders than in CBMs and the control group. IgG and IgM were significantly higher in welders than in CBMs while only IgA and IgM were significantly higher in welders compared to control group (p < 0.05) in each case (Table 3). The IgG level was significantly higher in the control group compared to CBMs (p < 0.05). Remarkably, IgG level in welders demonstrated approximately 2-fold increase compared with that of cement block moulders. Serum IgA levels appeared to be higher in welders than in CBMs however, this difference did not reach statistically significant difference.

Table 1: Occupational exposure and nutritional indices of studied population

Parameters	Welder	CBMs	Control	P-value	
	Mean ±SD, Mean Rank	Mean± SD, Mean Rank	Mean± SD, Mean Rank		
Occupational exposure (Hours/day)	$9.0 \pm 2.4$	$9.1 \pm 2.6$	-	0.918	
BMI (kg/m <sup>2</sup> )	$25.0 \pm 3.7$	$23.8 \pm 2.4$	25. 6± 3.0	0.065	
Protein (g/l)	$85.0 \pm 4.0, 51.7$	$83.0\pm3.0$ , $45.6$	82.0± 2.0, 39.2	0.179	
Albumin (g/l)	$44.0 \pm 6.0, 48.3$	$44.0 \pm 5.0,  44.7$	44.0± 4.0, 43.5	0.754	
Globulin (g/l)	$41.0 \pm 2.0, 50.5$	$39.0 \pm 1.0, 46.5$	38.0± 2.0, 39.5	0.259	

P< 0.05 is statistically significant.

Table: 2 Lead, cadmium, zinc levels and immunological parameters in the study population and control

Parameters	Welders	CBMs	Control	P-value	Expected
	Mean ±SD, Mean Rank	Mean± SD, Mean Rank	Mean± SD, Mean Rank		normal value
Monocyte				0.001	0.1-1.0
$(10^3/\mu l)$	$0.2 \pm 0.1, 25.1$	$0.5 \pm 0.2, 64.5$	$0.3 \pm 0.2, 47.0$		
Lymphocyte (10 /µl)	2.5±0.6	2.6± 0.8	$2.2 \pm 0.5$	0.003	1.0-5.0
IgA(mg/dl)	$440.0 \pm 120, 58.9$	$380.0 \pm 190, 35.6$	360.0± 80, 42.0	0.002	81-211
IgG (mg/dl)	1530.0±560,59.6	810.0 ±350, 27.8	1290.0±590,49.1	0.001	688-1251
IgM(mg/dl)	160.0 ±30.0	130.0 ±30.0	140.0 ±30.0	0.001	65-132
Zinc ( µg/dl )	$76.6 \pm 8.3, 45.0$	$65.6 \pm 10.1, 23.6$	$93.7 \pm 16.3, 67.9$	0.001	
Cadmium					
(µg/dl)	$0.1 \pm 0.0, 68.1$	$0.01 \pm 0.0, 52.2$	$0.002 \pm 0.0, 16.3$	0.001	
Lead( μg/dl)	$5.0 \pm 2.1,60.0$	$4.4 \pm 2.3, 51.1$	$2.3 \pm 1.0, 25.4$	0.001	

P< 0.003; 0.002 & 0.001 statistically significant CBMs = Cement block moulders

Table 3: Lead, cadmium, zinc levels and immunological parameters in the two exposed groups

Parameters	Welders	CBMs	P-value
	Mean± SD, Mean Rank	Mean± SD, Mean Rank	Expected
			normal value
Monocyte (10 /μl)	0.2 ±0.1, 18.2	$0.5 \pm 0.2,$ 42.8	0.001 0.1-1.0
Lymphocyte (10 /µl)	2.5±0.6	2.6± 0.8	0.651 1.0-5.0
IgA(mg/dl)	$440.0 \pm 120,$ 37.5	$380.0 \pm 190,$ 23.5	0.002 81-211
IgG (mg/dl)	1530.0±560, 41.1	810.0 ±350, 19.9	0.001 688-1251
IgM(mg/dl)	160.0 ±30.0	130.0 ±30.0	0.001 65-132
Zinc ( µg/dl )	$76.6 \pm 8.3,$ 39.6	$65.6 \pm 10.1,$ 21.4	0.001
Cadmium (µg/dl )	$0.1 \pm 0.0$ , $38.3$	$0.01 \pm 0.0,$ 22.7	0.001
Lead( μg/dl)	$5.0 \pm 2.1,$ 33.8	$4.4 \pm 2.3$ , 27.2	0.135

P<0.002 & 0.001 is statistically significant.

Table 4: Lead cadmium, zinc levels and immunological parameters in welders and control group

Parameters	Welders	Control	P-value
	Mean±SD, Mean Rank	Mean± SD, Mean Rank	Expected
			normal value
Monocyte (10 <sup>3</sup> /μl)	$0.2 \pm 0.1$ , 22.3	$0.3 \pm 0.2,$ 38.7	0.001 0.1-1.0
Lymphocyte	2.5±0.6	$2.2 \pm 0.5$	0.018 1.0-5.0
(10 /µl)			
IgA(mg/dl)	$440.0 \pm 120,$ 36.9	$360.0\pm 80,$ 24.1	0.004 81-211
IgG (mg/dl)	1530.0±560, 34.0	1290.0±590, 27.0	0.116 688-1251
IgM(mg/dl)	160.0 ±30.0	140.0 ±30.0	0.006 65-132
Zinc ( µg/dl )	$76.6 \pm 8.3,$ 21.0	$93.7 \pm 16.3,$ 40.1	0.001
Cadmium (µg/dl )	$0.1 \pm 0.0,$ 45.3	15.8 0.002±0.0,	0.001
Lead( μg/dl)	$5.0 \pm 2.1,$ 41.7	$2.3 \pm 1.0,$ 19.4	0.001

P< 0.02; 0.004; 0.006 & 0.001 statistically significant

Table 5: Lead, cadmium, zinc levels and immunological parameters between CBMs and control group

CBMs		Control		P-value	
Mean± SD, Mean Ran	ık	Mean± SD,	Mean Rank		Expected
					normal value
$0.5 \pm 0.2,$ 3	7.2	$0.3 \pm 0.2$ ,	23.8	0.003	0.1-1.0
2.6± 0.8		$2.2 \pm 0.5$		0.018	1.0-5.0
$380.0 \pm 190,$ 2	7.6	$360.0\pm 80$ ,	33.4	0.197	81-211
$810.0 \pm 350$ , 2	3.3	1290.0±590,	37.7	0.001	688-1251
130.0 ±30.0		$140.0 \pm 30.0$		0.121	65-132
$65.6 \pm 10.1$ , 1	7.7	93.7±16.3,	43.3	0.001	
	45.0			0.001	
$0.01 \pm 0.0$ ,		$0.002\pm0.0$ ,	16.0		
$4.4 \pm 2.3$ , $3^{\circ}$	9.5	$2.3 \pm 1.0$ ,	21.6	0.001	
	Mean $\pm$ SD, Mean Ran $0.5 \pm 0.2$ , 3 $2.6 \pm 0.8$ $380.0 \pm 190$ , 2 $810.0 \pm 350$ , 2 $130.0 \pm 30.0$ $65.6 \pm 10.1$ , 1 $0.01 \pm 0.0$ ,	Mean± SD,     Mean Rank $0.5 \pm 0.2$ , $37.2$ $2.6 \pm 0.8$ $380.0 \pm 190$ , $27.6$ $810.0 \pm 350$ , $23.3$ $130.0 \pm 30.0$ $65.6 \pm 10.1$ , $17.7$ $45.0$ $0.01 \pm 0.0$ ,	Mean $\pm$ SD,       Mean Rank       Mean $\pm$ SD,       Mean $\pm$ SD,	Mean $\pm$ SD,       Mean Rank       Mean $\pm$ SD,       Mean Rank $0.5 \pm 0.2$ , $37.2$ $0.3 \pm 0.2$ , $23.8$ $2.6 \pm 0.8$ $2.2 \pm 0.5$ $380.0 \pm 190$ , $27.6$ $360.0 \pm 80$ , $33.4$ $810.0 \pm 350$ , $23.3$ $1290.0 \pm 590$ , $37.7$ $130.0 \pm 30.0$ $140.0 \pm 30.0$ $65.6 \pm 10.1$ , $17.7$ $93.7 \pm 16.3$ , $43.3$ $45.0$ $0.002 \pm 0.0$ , $16.0$	Mean $\pm$ SD,       Mean Rank       Mean $\pm$ SD,       Mean Rank $0.5 \pm 0.2$ , $37.2$ $0.3 \pm 0.2$ , $23.8$ $0.003$ $2.6 \pm 0.8$ $2.2 \pm 0.5$ $0.018$ $380.0 \pm 190$ , $27.6$ $360.0 \pm 80$ , $33.4$ $0.197$ $810.0 \pm 350$ , $23.3$ $1290.0 \pm 590$ , $37.7$ $0.001$ $130.0 \pm 30.0$ $140.0 \pm 30.0$ $0.121$ $65.6 \pm 10.1$ , $17.7$ $93.7 \pm 16.3$ , $43.3$ $0.001$ $0.01 \pm 0.0$ , $0.002 \pm 0.0$ , $16.0$

P< 0.018; 0.003 & 0.001 statistically significant CBMs = Cement block moulders

## **DISCUSSION**

Globally, workplace hazards are recognized to be deleterious to the health of workers and the attendant public health concern has considerable medical and social impact. Welding and cement block moulding are common vocations and mostly embraced by those in the medium socioeconomic class in the developing countries. The rise in number of practitioners of these vocations coupled with the very few control measures has brought with it the possibility of excessive exposure to toxicants associated with different degrees of severity. In this present study, occupational exposure to toxicants, mainly cadmium and lead, present in welding fumes and cement dust in welders and cement block moulders, was evident by the significantly higher blood Pb and Cd levels than in the occupationally unexposed or control group. This appears also alluded by the significantly higher zinc level in the control than the occupational groups. An adverse effect of this exposure is that it may at least in part have contributed to the observed depression of immunological response indices in one of the occupationally exposed groups.

The significantly lower zinc level, an essential and prime micronutrient, in the exposed groups compare to control group is of central importance in this study. Given the key roles zinc plays in a number of biological processes including genomic activities embracing synthesis of indices

of immune function. The non-significant change in the nutritional indices such as body mass index, total protein and albumin in the exposed groups compare to

metabolic antagonism between the toxic metals, lead and cadmium. The synergistic inhibiting effect of Cd and Pb induced metallothionein synthesis may be operant (28). This might have also played a role in the observed significantly reduced zinc level in the exposed groups. Aside from exposure to the metallothionein inducer, Cd is a well-known metabolic antagonist of Zn (29). Thus the observed lower level of zinc in the exposed groups could also in part be attributed to the higher Cd level. Our result is consistent with the reports of Dioka et al. (30) and Ayatollahi (31) who observed decrease in serum Zn level in artisans who were occupationally exposed to low Pb level. Other investigators, Mehdi et al. (32) and Chiba et al. (33) have reported non-significant differences in the Zn levels between Pb occupationally exposed groups and control group. The explanation for these discrepancies is not clear, but may involve a combination of nutritional status duration of occupational exposure and the constituents of the metals involved. Li et al. (34) have demonstrated that long - term low level exposure to the welding fume increases serum concentrations of manganese, iron lead, while it decreases serum zinc level among the career welders.

The significantly increased Zn level in welders compared to CBMs may be due to the higher Zn in welding some of

which contain galvanized Fe. Several studies have shown that welders are exposed to high level of Zn in welding fumes. Steel and Sanderson (12) investigated the toxic fumes produced from various types of coated welding electrodes and found Zn concentration produced by burning such electrodes to range from 1.07 to 2.76. The elevated Zn level in the welders compared to the cement block moulders is analogous to supplementation with Zn to welders. Noweir (13) showed that cement dust contains 12.25µg zinc/g of cement concentration, a concentration that is lower than the level of zinc reported by Steel and Sanderson (12) in welding fume. This may corroborate the higher level of zinc observed in welder compared to CBMs being largely due to that inhaled during welding process.

Importantly, the observed higher levels immunoglobulins in welders compared to CBMs may be ascribed to the significant increase in zinc level in welders. Supporting a modulatory role of zinc in humoral immune response. Although previous reports in occupationally exposed individuals to welding fumes have been equivocal, Borska et al. (35) reported decreased levels of serum IgG, IgA, and IgM in welders with blood Pb level greater than the recommended WHO safe limit (WHO recognized toxic level, 10µg/dl). The low levels of Pb (5μg/dl) in our study may explain why there was no immunosuppression in both exposed groups.

The observed higher levels of IgM and IgG in welders is of great biological significance. It appears to support a role of Zn in immune response and that this principle could be applied to modulate the immunotoxic effects of exposure to toxic metals, which may be, dose dependent.

While IgM is associated with primary immune response, IgG plays an important role in secondary immune response. Therefore, the increased IgG and IgM may be indicative of immune enhancement induced by higher Zn level in welders compared to CBMs. One implication of this observation is that there is increased IgG in turn due to raised Zn level in the welders' group. The report of Ayatollahi (31) demonstrated lower IgG levels but IgA and IgM were unchanged in car battery workers with low Pb and low zinc exposure. This appears consistent, as there was no Zn supplementation to boost humoral immune response. Cement block moulders in this study also had their IgG levels significantly lower compare to control and welder group while IgA and IgM were not significantly different from control group. The report of Ayatollahi (31) and our findings in CBMs appear to suggest that workers engage in occupations that expose them to Pb and low Zn may have low level of IgG, probably implying to supplement the workers with zinc. It also appears to underscore that zinc has profound effect on IgG synthesis than on other immunoglobulin classes. Thus, the immunostimmulatory effect of Zn on IgG synthesis was demonstrated in welder with higher Zn level compare to CBMs.

The significantly higher humoral immune parameters, IgA and IgM in welders compared to the control group despite lower but optimum Zn level in welders. This may suggest a role for stoichiometric relationship between the toxic and essential metals, implying that the Zn intake during the welding process, not only modified the immunoglobulin levels but competed and overturned the immunoinhibitory effect of toxic metals on humoral immune response.

The elevated lymphocyte count in welders in our study is similar to the report of Hanovcova et al. (23) who observed increased lymphocyte counts in welders compared to unexposed populations. This may play a role in the argument on immune response enunciated above. Some investigators such as Shenker et al. (36); Gaworski and Sharma (37); McCabe and Lawrence (38) have reported the possible immunostimulatory effect of low blood Pb level on lymphocyte proliferation. Many of these reports did not consider the possible role of zinc. The main message of this study gaps not withstanding is that zinc has a modulatory effect on immune function particularly in humoral immune system. This as might be expected arises from the diversity of the function of Zn with a wide range of effects including protein synthesis of which the humoral immune system is dependent (39).

## CONCLUSION

Although previous reports in oc

The data from this study suggest that exposure to welding fumes and cement dust perturbed humoral immune system but modulated by the higher Zn level in the welders. The higher Zn level in the welders and the corresponding higher lymphocyte and approved a specific and approved a role Zn in i levels in the same group appear to argue strongly for a modulatory role for Zn in immune response in these workers. This appears supported by the findings in the CBMs with low Zn levels exhibiting significantly decreased IgG level compared to the welders. One implication of these data is that the immunosuppressive roles of toxic metals such as Pb and cadmium may be modulated by increase zinc intake. An indirect support for this appears to be demonstrated by the antagonism of Pb and Cd to Zn by the lower levels of zinc in the exposed groups than in controls and associated indices of immune response and may explain why there could be varied susceptibility to illness in occupationally exposed individuals. The data imply that cement block workers may benefit from zinc supplementation.

#### **Recommendations**

Every measure should be taken to reduce exposure to metal fumes; metal fume exposure, particularly in the rapidly developing nations is inevitable but can be modulated by nutritional means. Owing to the observed positive modulating effect of Zn in this study, increased intake of this micronutrient may serve as host resistance to the deleterious effect of Pb and Cd resemble the humoral i

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metal fumes) on the immune system. Zinc therefore may be considered a factor in public health that deserves greater recognition and the need for a separate advised dietary intake (ADI) in polluted communities.

#### **Conflict of Interest**

Authors declare that they have no competing interest.

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