



Heavy Metal Contributions on Human Skin Disease near Cement Plant: A Systematic Review

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Abstract

AIM: The aim of this systematic review was to investigate heavy metal roles at skin disease of humans in the cement industrial area.

METHODS: All research reporting a specific heavy metal, which could be increasing the risk of the disease, cause cement induced to skin. Fourteen selected articles from our inclusion criteria rated using eight quality-appraisal criteria derived from Strengthening the Reporting of Observational Studies in Epidemiology checklist and following preferred reporting items for systematic reviews and meta-analyses methodology.

RESULTS: We performed a systematic review of published articles between 2009 and 2019 from ten case-control, two cohort, one ecological study, and one cross-sectional study showed the heterogeneity of the study design. The studies are focusing on the relationship between heavy metal and skin disease that appeared in cement plant areas such as dermatitis, skin cancer, eczema, and other allergies. Chromium, nickel, cobalt, zinc, cadmium, mercury may present and play an important role for human skin disease.

CONCLUSION: Heavy metals have been found in the skin of human living and work around the cement industry. Chromium, cobalt, and nickel are almost certainly present in every case of dermatitis and eczema. Signs and mild symptoms of skin disease include redness, itching, starch, and dryness. Elements such as cadmium, lead, and mercury usually are found in chronic skin disease (psoriasis and skin cancer) on a biologic test. These findings should raise an important concern and recommendation policy for controlling raw materials contains heavy metal in cement industrial areas and proper protection for workers and non-worker.

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Introduction

Heavy metals have a long half-life and found in the body through the skin, food, or drink [1]. Heavy metal released from industrial activity, coal combustion, and mining. Moreover, if heavy metal particulates are carried by the wind, the distribution is able to reach a far place from the source while the wind blows. Heavy metal is one of the most serious environmental pollutants because of its toxicity, abundant, easily accumulated by various plants and organisms [2]. Exposure to heavy metal are associated with cancer [3], [4], obesity [5], infant mortality [6], cognitive problem, and skin damage [7].

Industries have been considered as a source of heavy metal. The main problem that some industries, and more specifically, cement industries, still have to deal with is the requirement to determine and control the content of some chromium, nickel, and cobalt compounds due to their allergenic effects on human health [8]. Some important studies mentioned the harmful effects of Cr (VI) on human health, mainly skin in contact with cement such as dermatitis [7], [9]. Dermatitis is an inflammatory skin condition which will be found around the cement industry area with symptoms

of itching, redness, burning, and skin irritation from the contact of cement corrosive components. Furthermore, increasing heavy metal concentration near the plant related with extreme skin disease, for example, skin cancer and mortality from cement manufacturing activities [10].

Skin damage of cement industrial can be caused through physical insults, biologic cause, and long-high concentration of chemical exposure [11]. For workers, occupational exposure to heavy metal occurs mainly through inhalation and dermal absorptions in the working environment. Similarly, residence in near of an industrial activities affected by the toxicity of excess material from factory for many years [12]. The skin can be one of the most vulnerable organs of heavy metals penetration into human body surfaces. Disorders and signs of the skin disease occur in a short and long time period depending on the concentration, heavy metal characteristics, and duration of exposure. Redness was an initial symptom that a human skin sign and being a possible alert of a skin problem. However, legislative initiatives may be necessary to protect all humans near to cement industrial area for a threshold level concern and protective plan [13].

Materials and Methods

Eligibility criteria

The criteria were determined based on the format (P = Population, I = Intervention, C = Comparison, O = Outcome, and S = Study Design).

P = Patients at any age, sex, and activity indicates skin disease near the cement plant

I = Heavy metal exposure and induced to skin

C = No exposure; occupational risk; concentration

O = Heavy metal contributions and roles on skin disease

S = Observational studies

Search strategy

This literature review assessed and arranged using the 2015 preferred reporting items for systematic reviews and meta-analysis (PRISMA) guidelines. List of relevant words was selected and determined to get peer-view articles on an appropriate topic. A computerized literature searches in four international online databases: DOAJ, PubMed, SpringerLink, and Google Scholar. All published papers and reports associated with the subject of this study from January-2009 to December-2019. We performed a systematic search using Mesh terms including: "Heavy metal exposures," "dermatitis," "cement industry," "mercury," "lead," "arsenic," "chromium," "hexavalent chromium," "skin lesions," and "skin cancer." The "AND" and "OR" operators were used. The combination of keywords used in this research was (dermatitis) OR (skin cancer) OR (eczema) AND (heavy metal) AND (industry) OR (cement) AND (chromium).

Inclusion and exclusion criteria

Articles taken from each database are imported into the Mendeley Library. For additional relevant publications that might be missed, we look for bibliographic references from all articles that met the inclusion criteria. The research inclusion criteria, focusing on cement plant area (residences, workers, and patients); an observational study; heavy metal exposure to skin; published in 2009–2019; English articles; and peer view journal. We excluded studies for following reasons; if the data contain less information in methods, no original data included qualitative studies, non-epidemiological research, articles published in predatory publication, small amount of sample (<50), unmentioned locations, and industrial health effect to skin except cement industry. Three reviewers extracted the results and read all the unique papers and assess them with exclusion and inclusion criteria. The titles and abstracts of papers were read, and all papers were unrelated to research objective (heavy metal related to skin disease) were excluded from further analysis (189 papers).

Data extraction and quality assessment

Data on the following characteristic were screened by reviewers using eight indicators of quality-appraisal criteria derived from Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) checklist. The checklist consists of 22 items that are used to evaluate external validity (based on potential selection bias) and internal validity (based on the potential measurement and confounding bias) from observational studies. After the initial assessment of all studies reviewed based on 22 STROBE items, these items then grouped into eight quality assessment criteria (sample size, sampling methodology, response rate, outcome measures, statistical analyses, control for confounding, study limitation, and ethical consideration). STROBE checklist (0–8) becomes the standard criteria for selecting articles. Studies that scored <3 considered to be low quality, 3–6 medium, and 7–8 as a good quality study. Articles rate with <7 will be removed and if they gained a good rate (7–8), they would be taken. Fourteen studies were selected based on the quality criteria. There were five studies with a rate of 8 and 9 studies with a rate of 7. The diagram of paper selection is shown in Figure 1.

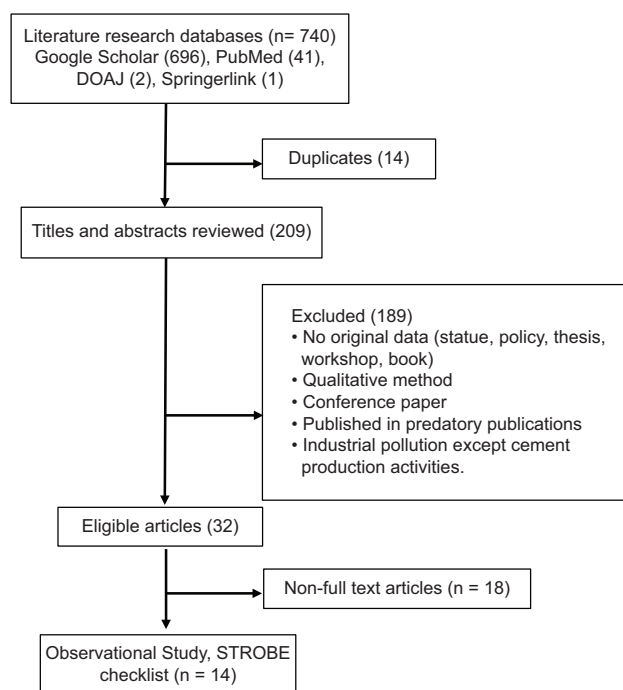


Figure 1: Preferred reporting items for systematic reviews and meta-analyses flow diagram of paper selection

Results

Skin disease

Exposure of heavy metals from coal combustion and high-temperature processes in the industry will release harmful particles (gas, dust, and metals) [14]. Cement dust has a significant correlation with health

outcomes. Minor health problems such as coughing [15], itching, and red eyes are mild symptoms that will arise due to body response to environmental quality. The industry is the sector that associated with population health problems. Dust from industries accumulates and frequently gets in contact with skin constantly through hands and forearms [16]. Alkalinity and corrosive materials in cement show effect and allergic to skin. Small outdoor jobs gained the highest number of dermatitis than other divisions in cement factories that usually done mechanically, especially indoor jobs. The less contact to skin, the fewer cases will appear.

In the cement industry, workers face the danger of routine exposure from allergens in their work. One of the most susceptible damage organs in humans is skin. Therefore, protection gloves that cover the surface of the skin are crucial. Dust in the cement industry is different from the dust around the roadside. The dust in cement contains silica [17], [18] and heavy metals [19]. Long term exposure affected the epidermal skin layer, protein metabolism, and DNA damage [10]. Dermatitis is a skin disease that is always found in workers in the cement industry, where most chromium, nickel, and cobalt will be found [20]. Irritant

and allergic contact dermatitis are considered the most frequent occupational health hazards in cement workers.

Metal sensitivity

Exposure to heavy metals has been extensively studied and reported as evidence for disorders of human skin in the vicinity of the cement industry. Dermatitis is a disease from external exposure from allergens, for example, chemicals [21]. If it is no further contact from an allergens, the redness, crusts, and itching will disappear. Common sources of exposure to the skin are tanned leather and cement. Dermatitis resulting from exposure to Cr(VI) can occur either by direct dermal contact or by ingestion [22]. Chromium found on the skin of patients with dermatitis proven by a skin patch test. Metals that cause systemic contact dermatitis are aluminum, chromium, cobalt, copper, gold, mercury, nickel, and zinc [23]. The safest limit of chromium exposure in the cement industry is 1 mg/Nm³ [24]. Chromium, in hexavalent form, is both toxic and carcinogenic. A Cr (IV) allergic patients also give a positive reaction to cobalt in a workplace. Based on Table 1, this theory related

Table 1: Skin disease reporting due to heavy metal

Author/year	Study design	Quality of assessment	Study group (N of subject, variables/risk factor)	Prevalence of skin disease	Diagnosis of skin disease (observed symptoms; analysis method)	Location/Site
Lejdning <i>et al.</i> (2018)	Case control study	7	6482 patients (construction activities); leisure activities and occupational cement contact; age	157 Cr(IV) and Cobalt-Allergic female and 76 Cr(IV) and Cobalt Allergic male related to dermatitis	Eczema, allergic contact to dermatitis; Skin patch test	Southern Sweden
Cha <i>et al.</i> (2011)	Case control study	7	374 respondents; age, drinking alcohol, urinary level of Cr	17 in Sinpyeong, 44 in SsangYong, 28 in Ipseok and 13 in Kwangkyeong (control). Chromium and mercury high concentration in exposure group of dermatitis disease	Burns sensitization and skin ulcer; Urinary analysis	Yeongwol, Jaecheon, and Wonju City, South Korea
Koh <i>et al.</i> (2011)	Cohort study	7	5596 respondents (cement workers); age, sex	1 observed case for skin cancer	n/a	6 Portland cement factories in South Korea
Perez <i>et al.</i> (2015)	Ecological study	7	36667 observed death by cancer (cement workers); specific rate by age group	25 observed death related to skin cancer (14 men, 11 women)	n/a	Spain
Kridin <i>et al.</i> (2016)	Cohort study	8	4846 patients (cement workers); sex, occupational allergic to cement and age	146 positive for potassium chromate, 46 chromate allergic patients OACD, 27/46 had cement induced chromate OACD	Metal allergy (Cobalt, Nickel, Chromate); Skin patch test	Israel
Afridi <i>et al.</i> (2010)	Case-control study	8	241 respondents (live in town since birth and were 25–55 years old)	Increasing Cd, Cr and Ni concentrations made 124 exposed to Psoriasis, 63 male and 61 female	n/a; Biological sample (scalp hair, blood, and urine)	Hyderabad, Pakistan
Elhosary <i>et al.</i> (2014)	Case-control study	7	65 males (14–65 years); coffee/tea drinking, smoking, residence, education	Cr levels of two exposure group were higher, about (22.7%) of cement and (35%) of tannery workers had severe skin manifestations. ; n/a	fissured skin and chrome ulcers; Laboratory investigation were carried out by blood and urine sample	Egypt
Thomas <i>et al.</i> (2015)	Case-control study	8	250 respondents; n/a	49 of 250 patients reacted to one or more bone cement constituents, 2 patients in potassium dichromate, 1 patient in cobalt chloride, 11 to nickel sulfate	Pain, recurrent effusions, reduced range of motion, eczema or loosening; skin patch test	Germany
Wong <i>et al.</i> (2014)	Case control study	8	3685 respondents; n/a	47 (41%) cases of chromium OACD caused by cement were identified	Allergic reaction; skin patch test	Australia
Schwensen <i>et al.</i> (2014)	Case-control study	7	1471 blue-collar workers, DISCO 88; MOALHFA index, sex, age	64 over 1468 had positive allergy to potassium chromate, 68 were allergic to cobalt chloride, 178 were allergic to nickel sulfate	contact allergy, sensitization; Skin patch test	Denmark
Thyssen <i>et al.</i> (2010)	Case-control study	8	22506 patients with dermatitis; MOALHFA index, sex	1281 nickel allergic patients, 425 cobalt allergic patients, 275 chromate allergic patients	redness, infiltration, irritant response; skin patch test	Denmark
Le <i>et al.</i> (2010)	Case-control study	7	110 patients with eczema and 41 patients with miscellaneous skin conditions (SCORAD, NESS); age, sex, dietary intake	44 patients with eczema and 24 in non-eczema had a low level of copper	n/a; Serum level by pediatric dermatology clinic	Hong Kong
Thyssen <i>et al.</i> (2009)	Case-control study	7	16228 patients, MOALHFA index; age, test year, sex	100 patients had positive test reaction to chromium	Chromium allergy to skin; skin patch test	Denmark
Wang <i>et al.</i> (2011)	Cross-sectional study	7	97 cement workers; sex	24 males were allergic to potassium chromate, 4 males were allergic to nickel sulfate	Thickened, hyperkeratosis, scaling, dryness, erythema, fissure, pigmentation, scratch, ulceration, erosion, edema, itching, vesicles; Skin patch test	Tainan City, Taiwan

n/a: Not available. OACD: Occupational allergic contact dermatitis.

to Lejding (2018), Schwensen (2014), Thomas (2015), and Thyssen (2010) research. Chromium in cement to <2 ppm through the addition of ferrous sulfate to cement has seen a significant decrease in the incidence of chromium allergy [25]. It means, reducing and controlling chromium in the cement industry are required.

Nickel allergy is the most common cause of Allergic Contact Dermatitis (ACD), and its incidence is thought to be increasing by the growth of industries. Sensitized individuals who are exposed to nickel on their skin or mucosal surfaces generally have a predictable localized response, including erythema, vesicles, scaling, and pruritus [23]. Furthermore, the cement factory in the USA has been lead to human skin disease and cobalt exposure recorded about 0.02–0.5 ppm in different countries [26]. Cobalt allergy may also occur in the construction industry, primarily through skin contact with cement.

Mercury, one of the most toxic heavy metals also found in serum samples of skin chronic disease detection. Body burden of mercury associated with eczema [27]. Raw materials in cement contain mercury, lead, cadmium from fuel, and coal [28]. These heavy metals determined by metallothionein analysis in serum, hair, and nail as a biomarker for metal detection. In Table 1, Afridi (2010) found a high concentration of Cd, Cr, and Ni and the high number of psoriasis in males and females in Pakistan. Psoriasis is a chronic skin condition, unnoticeable, and long-lasting. These carcinogenic elements can trigger skin disease.

MOAHLFA (Male, Occupation, Atopic dermatitis, Hand eczema, Leg dermatitis, Facial dermatitis, and Age > 40 years) index used by Schwensen (2014) and Thyssen (2010) to determine associations characteristics of patients of allergic contact dermatitis [34]. Skin patch tests commonly used for all patients with Atopic Dermatitis [35]. Homogeneous redness and infiltration in the entire test area were scored as a 1+ reaction. Homogeneous redness, infiltration, and vesicles in the test area were scored as a 2+ reaction and homogeneous redness, infiltration, and coalescing vesicles in the test area as a 3+ reaction. A 1+, 2+, or 3+ reading was interpreted as a positive response. An irritant response, a doubtful (+?), or a negative reading was interpreted as a negative response [36].

Risk factors

The working area is a crucial and determining factor in the emergence of skin disease. Human activities will affect the health outcome and illness. In the cement industry, workers in the industry are the riskiest population for skin diseases due to allergen induction. The most common skin disease related to heavy metal in cement was dermatitis. Lejding (2018) investigated 6482 patients working in the construction and cement sectors in Sweden, 157 women and 76 men were associated with heavy metal chromium and cobalt exposure. Chromium and cobalt were also found on the skin of workers in Tainan City, Taiwan with 23 men and one woman allergic to potassium chromate, four women allergic to nickel sulfate over 97 cement workers [29].

Research in Denmark found that 64 over 1468 blue-collar workers had a positive allergy to potassium chromate, 68 to cobalt chloride and 178 were allergic to nickel sulfate [30]. Kridin (2016) reported 46 chromate allergic patients of occupational allergic contact dermatitis from 4846 cement workers in Israel and increase currently. In terms of skin cancer, mortality recorded in two countries. One case of skin cancer of cement producing workers revealed in South Korea. This case happened to a worker who had been worked for 30 years. Furthermore, In Spain, an ecological study caused 14 cancer incidence in male and 11 female cement workers. All of the patients with skin cancer worked in a cement plant [31].

Residence or non-workers in the cement industry face a huge problem. In the study of heavy metal exposure of Cd, Cr, Ni, and Zn in biological samples of psoriasis for living human near cement in Pakistan, the result was significantly higher for residence than control. Moreover, other diseases experienced by residents living next to the cement plant were eczema and dermatitis. Topsoil heavy metal able to mobilize based on the wind direction and dissolved in consumed water [32]. Patients with skin diseases took a higher heavy metal concentration than the control/un-exposure group in their skin surface and biological samples (hair, blood, and nails), and it accumulates for a long period. It was found that chromium levels in blood and urine from cement workers were higher than control in Egypt [33].

Table 2: Prevalence of skin disease from cement industry in 2009 to 2019

Author	Total sample	Skin disease cases	Amount of cases (%)	Heavy metal	Observation study
Lejding <i>et al.</i> (2018)	6482	233	3.59	Cr (IV)	10 years
Cha <i>et al.</i> (2011)	374	102	27.27	Cr (IV), Hg, Cd, Pb	n/a
Koh <i>et al.</i> (2011)	5596	1	0.02	Cr (VI)	8, 15 years
Perez <i>et al.</i> (2015)	36667	25	0.07	Cr (IV), Mg	9 years
Kridin <i>et al.</i> (2016)	4846	146	3.01	Cr (VI)	14 years
Afridi <i>et al.</i> (2010)	241	124	51.45	Cr (III), Cr (IV), Cr (VI), Ni, Cd	n/a
Elhosary <i>et al.</i> (2014)	65	27	41.54	Cr (IV)	1 year
Thomas <i>et al.</i> (2015)	250	49	19.60	Cr (IV)	3 years
Wong <i>et al.</i> (2014)	3685	47	1.28	Cr (IV)	20 years
Schwensen <i>et al.</i> (2014)	1471	310	21.07	Cr (IV)	9 years
Thyssen <i>et al.</i> (2010)	22506	1981	8.80	Cr (VI), Ni, Co	33 years
Le <i>et al.</i> (2010)	151	68	45.03	Zn, Cu	1 year
Thyssen <i>et al.</i> (2009)	16228	7	0.04	Cr (III), Cr (IV)	22 years
Wang <i>et al.</i> (2011)	97	28	28.87	Cr (VI)	n/a
Total	98659	3148			

n/a: Not available

The prevalence of skin diseases caused by heavy metals described that the amount of case-control studies was greater than others (Table 1) and being the most common methods ($n = 10$). The complete data from all reviewed papers of prevalence of skin disease in 2009–2019 are described in Table 2 and Figure 2. There was 98659 samples from 10 years studies and 3148 cases had positive tested to heavy metal and indicated as skin disease.

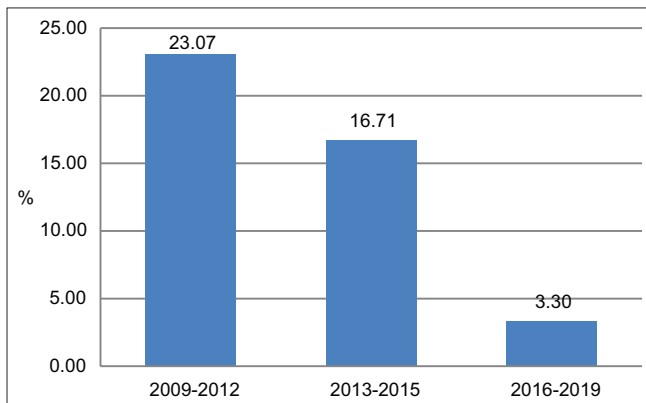


Figure 2: The columns represent 3 years period (e.g., 2009–2012, we could assume that the last column was 3 years because no research found in 2019) and proportions of patients who tested positive for skin disease due to heavy metal exposure near the cement plant

Strengths and limitations

Long-term consequences of human exposure to cement dust should be investigated. The number of papers and the specific risk factors was not significantly mapping the issues. Skin disease association with heavy metal in the cement area is less than lung inflammations, cardiovascular, and cancer studies. Therefore, we observed a positive association between chromium-cobalt allergies with dermatitis. For specific elements such as mercury and cadmium, the exposure may be classified as a chronic level. These two elements accumulated in the organism body and made an important role in environmental biomonitoring but did not explain the process to lead skin cancer.

A total 14 studies shown a diverse location (Sweden=1, South Korea=2, Spain=1, Israel=1, Pakistan=1, Egypt=1, Germany=1, Australia=1, Denmark=3, Hong Kong=1, and Taiwan=1). Our review was collect various methods and recent data in skin disease focusing on the cement industry. Some studies did not mention the specific symptoms relates to skin cancer but obtained good data for mortality rate. Only the age and occupation variables showed high relevance with the incidence of skin disease, in contrast to gender variables, which produced different results for each study — heavy metal exposure to a human living in a cement industrial area associated with the number of skin disease. The result of this systematic review mentioned chromium, nickel, and cobalt was a major element appearing on skin patch test and other tests. Elements such as cadmium, lead, and mercury usually are found in chronic skin disease on a biologic

test. These findings should raise an important concern and recommendation policy for controlling heavy metal in industrial area for worker and non-worker. Further detailed work and research, however, to better characterize these associations, risk factors, and to assess causality, are needed.

Conclusion

Heavy metal exposure to a human living in a cement industrial area is associated with the case of skin disease. The result of this systematic review mentioned chromium, nickel, and cobalt was a major element appearing on skin patch test and other tests. Signs and mild symptoms of skin disease include redness, itching, starch, and dryness. Elements such as cadmium, lead, and mercury usually are found in chronic skin disease on a biologic test. These findings should raise an important concern and recommendation policy for controlling heavy metal in industrial area for worker and non-worker. Further detailed work and research, however, to better characterize these associations, risk factors, and to assess causality are needed.

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