Comparison of serum lead level of opium dependent patients with a healthy control group

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Introduction: The prevalence of multiple symptoms, including mental illness, symptoms that mimic diseases such as pancreatitis, nonspecific abdominal pain and cholecystitis, are high in drug-dependent patients. Objectives: The aim of this study was to consider the clinical signs and serum lead levels of individuals referred to four university hospitals with various complaints. Patients and Methods: In this study, 128 sample cases in two groups, the case group and control group, have been examined. The case group included 64 persons using oral opium who were suffering from various complaints and referred to emergency department of mentioned hospitals. The control group included 64 patients without any history of addiction. Results: Mean of blood serum lead level in opium users and control group was 76.34±17.82 mg/dL and 7.68±3.72 mg/dL respectively, and the difference between these groups was significantly meaningful (P<0.001). The most prevalent complaints were abdominal pain and pulmonary complaints. Conclusion: Findings of the current study showed high levels for an average serum lead level for oral opium addicts among the studied statistical population. In addition, results show that ultimate diagnosis of lead poisoning is a factor justifying symptoms for addicts with nonspecific symptoms, the subject that confirms several previous reports. This finding asserts the necessity of screening of serum lead level for patients addicted to opium compounds in order to prevent more severe side effects.

Keywords: Lead, Abdominal pain
Oral opium

Key point
The prevalence of multiple symptoms and nonspecific abdominal pain are high in drug-dependent patients. In this study, 128 sample cases in two groups have been examined. The case group included 64 patients using oral opium who were suffering from various complaints and control group included 64 patients without any history of addiction. Average blood serum lead level was significantly higher in the case group compared to the control group (P<0.001). The most prevalent complaints were abdominal pain and pulmonary obstruction. Mean serum lead levels were high in oral opium addicts.

Objectives
Considering the frequency of patients with

were oral opium users, has definitely supported the idea that this kind of toxicity was mainly because of lead added to opium (5).

Adding lead to opium is highly hazardous and causes serious side effects, death, resulting in irreparable damages for nations as well as acute poisoning (6, 7).
different complaints including headache, non-specific abdominal pain, constipation, irritability and memory disorders among the Iranian population, this research aims to study clinical symptoms and blood lead level (BLL) of individuals referring to Rasul Akram, FiroozAbadi, Firoozgar, and Haft-e-Tir hospitals.

Patients and Method
In this case-control study, 128 sample cases in two groups, the case group and control group, have been examined. Sixty-four patients using oral opium who were suffering from various complaints and referring to emergency department of our hospitals were examined and then their serum lead level was assessed. Control group included 64 healthy individuals, while none of them were sick and were matched regarding age and gender with the first group. Their serum lead level along with other variables were also assessed.

Using checklists, information related to two groups including demographic information, health condition and life style was gathered. After completion of checklists, data were analyzed as per research purpose.

Ethical issues
Human rights were respected in accordance with the Helsinki Declaration 1975, as revised in 1983. The ethical committee of Iran University of Medical Sciences (Ethical code# IR.IUMS.FMD.REC1396.94113070121) approved the study. The informed consent was taken from the patients as well as from parents and first relatives. This paper was extracted an original research from the Ph.D., thesis in emergency medicine of Shahin Dokht Hasan Pour (Thesis #4981) in Iran University of Medical Sciences Faculty of Medical (Emergency Department).

Statistical analysis
The data were analyzed by IBM SPSS 23 statistical package (SPSS Inc, Chicago, and IL USA). Results were analyzed using descriptive statistics methods, frequencies, cumulative frequency percentage and charts. To analyze the data, t test, one-way analysis of variance (ANOVA) and chi-square were applied. To analyze the data, t test, one-way analysis of variance (ANOVA) and chi-square were applied. The correlation between BLL and drug consumption was investigated by regression analysis.

Results
Case group included 64 participants from 29 to 77 years old with average age of 46.27 ± 9.12 years old. Among them 97% were men. The most significant complaint was abdominal pain and intestinal obstruction. All cases went under endoscopy and colonoscopy.

Other symptoms of patients were; decreased appetite, chronic constipation, nausea and vomiting, memory disorder, anemia, finger paresthesia, weakness and fatigue, weight loss, bone and muscular pain. Diagnostic sonography results were normal for all cases however; four cases had gallbladder stone and sludge. Two cases underwent laparotomy, although final diagnosis for them has been ileus. Among 33 patients suffering from abdominal pain and 71.28% had anemia. Liver enzymes were elevated in 32.41% of cases. Renal function was normal for all cases.

Serum lead levels were 41 to 105 mg/dL with average of 76.34 ± 17.82 mg/dL for case group and 3 to 18 mg/dL with average of 7.68 ± 3.72 mg/dL for control group. Average blood serum lead level was significantly higher in the case group compared to the control group (P < 0.001).

Oral opium users' lead level in symptomatic patients is shown in Figure 1. Serum lead level for all patients in the case group who admitted with specific symptoms was higher than 25 mg/dL (toxic), Serum lead level for 70.96% patients with non-specific symptoms was higher than 25 mg/dL (toxic), And in control group serum lead level was below 25 mg/dL for all cases. There is a significant difference in ranges of serum lead levels in the three groups (P < 0.05; Table 1).

Daily oral opium consumption in individuals under 50 years old was 1-3 g with mean of 1.73 ± 0.23 g. On the other hand, for those aged more than 50 years old, 2-4 grams of daily consumption with average use of 2.89 ± 0.27 g was reported. There was a significant relationship between serum lead level and quantity of narcotic utilization in the case group, which indicates serum lead level has significantly increased as drug use has increased (P = 0.032).

The duration of oral opium use in the case group was from five months to 30 years with average time of 15.24 years. For the case group, no significant relationship between serum lead level and the duration of utilization was detected (P = 0.213; Figure 2).

Discussion
Lead toxicity has been recognized for thousands of years and still exists today. Lead as a heavy metal can act as an acute or chronic toxicity cause.

Inorganic lead is absorbed through the lung and gastrointestinal tract and then gets accumulated in blood, soft tissues and bone. Clinical symptoms and signs of lead...
poisoning vary from patient to patient. Lead poisoning influences central and peripheral nervous systems, kidneys performance, vascular system and digestive system and causes anemia, kidney failure, neuropathy and digestive symptoms (12).

Masudi et al reported three cases of lead poisoning in drug dependent people and therefore presented it as a new source of lead poisoning in Iran (8).

Addiction and its corresponding socioeconomic consequences is one of the most important problems in countries such as Iran. Several studies have reported lead poisoning for addicts. In addition, lead poisoning in drug-dependent patients has been observed before; e.g., Algora et al reported some instances of lead poisoning in addicts who referred to hospitals with abdominal pain and anemia (9).

Beattie et al reported 5 cases of lead and drug injection (10). Antonini et al explained a case of lead poisoning in heroin addicts (11). Lead traces in drugs has also been reported in the study by Aghae et al (12). They showed that drug dealers and smugglers may use lead as an additive to drug during production which helps them to enhance weight and gain in more benefit. Thus drug is known as a potential source for lead poisoning. In this study, patients’ BLL was higher than the control group which indicated a significantly meaningful difference between the two mentioned groups. Moreover, patients’ BLL for drug addicts was correlated with amount of drug consumption. Both groups were matched and they only differed in oral drug use; therefore, BLL increases can be as a result of drug contamination with lead. All persons had toxic levels of lead which were equivalent to 25 mg/dL and more (8).

On the other hand, the prevalence of various symptoms including mental diseases, symptoms that imitate diseases such as cholecystitis, pancreatitis, and non-specific abdominal pains are high in drug addicts (13-15).

Soltaninejad et al (2011) evaluated 25 patients seen from 2002 to 2007 with lead poisoning in the Loghman Hakim hospital poison center, Tehran, Iran. Seven of the patients had a history of opium addiction as the only probable source of lead poisoning, and none of them had occupations known to involve exposure to lead. The average age of these patients was 38.7 ± 9.3 years. The average BLL in these patients was 109 ± 37.6 mg/dL Lead poisoning should be considered in patients with a history of opium abuse who present with non-specific clinical manifestations.

Wolf’s study presented drugs as a potential source for lead poisoning. Patients’ BLL was higher than control group which was statistically significant. In addition, there was a relationship between drug-dependent patients’ BLL and drug use. Two groups were matched and the only differed in oral drug use respect, therefore BLL elevation in case group can be due to drug poisoning by lead. Around 40% of patients’ lead level was toxic which is equivalent to 25 mg/dL and more (13).

Considering the research findings of Anderson’s study, it seems that high BLL level can justify these findings for drug addicts. Hence, screening of serum lead concentrations in drug addicts, especially for those with non-specific complaints, can be useful. There was not any specific relationship between BLL and duration of drug abuse in the case group, which is expected by considering half-life of 36 days of lead and duration of drug use in individuals (at least 2 years) (15).

In moderate and chronic cases, involvement of central nervous system and symptoms associated with nephropathy have been reported by Lochitch (17). For adults, BLL elevation in mainly because of occupational contacts (18). Thanks to higher levels of workplace safety during recent years, occupational lead poisoning has decreased among adults in the world and new forms of non-occupational poisoning have emerged (19). New type of lead poisoning

| Lead level (mg/dL) | Patients referring with dominant symptoms | Patients referring with non-dominant symptoms | Control group | P Value |
|-------------------|-----------------------------------------|---------------------------------------------|---------------|--------|
| <25               | 0.00                                    | 29.04                                       | 100           | 0.016  |
| 25-45             | 12.13                                   | 41.95                                       | 0             | 0.029  |
| 45-69             | 21.21                                   | 19.35                                       | 0             | 0.041  |
| 70-99             | 57.57                                   | 9.66                                        | 0             | 0.003  |
| >100              | 9.09                                    | 0                                           | 0             | <0.001 |
resulting from opium use which is polluted with lead is gradually emerging in our region (20). Drug addiction and addiction patterns are turning into high risk methods and drugs. Drug suppliers and users may add lead for increasing the weight and for opportunistic reasons during the drug preparation process. Moreover, several cases of pathologic findings like neuropathy, abdominal pain and anemia have been observed for opiate addicts (21).

Conclusion
Findings of the current study suggested high levels of serum lead level for oral opium addicts among the studied statistical population. In addition, results show that ultimate diagnosis of lead poisoning is a factor justifying symptoms for addicts with specific symptoms, the finding that confirms several previous reports. This finding asserts the necessity of screening of serum lead level for patients addicted to opium combinations in order to prevent more severe side effects.

Drug is considered as a potential source for lead poisoning. In this study, patients’ serum lead level was higher than the control group which indicated a significantly meaningful difference between two mentioned groups. Moreover, patients’ BLL for drug addicts was correlated with amount of drug consumption. Both groups were matched and they only differed in oral drug use, therefore, BLL increase can be due to drug contamination with lead. Based on the results of this study 100% of oral opium addict individuals with specific symptoms, had toxic levels of lead which was equivalent 25 mg/dL or more. Thus screening blood lead concentration is a beneficial point for addicts especially those who have non-specific complaints.

Recommendations
• Considering the point that, costs and consequences of lead poisoning can be totally prevented through reduction and elimination of exposing lead and early diagnosis of high levels of lead in the blood. Clinicians of primary care should play an active role in prevention and early diagnosis of individuals who are exposed to lead.

• Moreover, conducting further research using larger groups of oral/inhaled opium addicts can correspond with more determination and confidence of reports on criteria.

Limitations of the study
- Non-cooperation of patients for participation in the study and data collection.
- Incomplete data required by the patients.

Authors’ contribution
ShH and GhM designed the study, observed accuracy and validity of the study. ZH and HH collected the data and follow the study. ShH, GhM and MR supervised the project. ShH and GhM wrote the paper. All authors edited and revised the final manuscript and accepted its publication.

Conflicts of interest
The authors declare that they do not have any conflict of interest.

Ethical considerations
Ethical issues (including plagiarism, data fabrication, double publication) have been completely observed by the authors.

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