Review Paper



Differentiation of Suspected Methanol Poisoning From Ethanol Poisoning: A Review Study and Consensus Statement

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ABSTRACT

Background: Ethanol and methanol poisoning pose significant clinical challenges due to their overlapping clinical presentations. This literature review and consensus statement aim to provide evidence-based guidelines for differentiating these two poisonings.

Methods: We conducted a comprehensive literature review of relevant literature, including individual patient data (IPD) meta-analysis, to assess clinical, laboratory, and diagnostic findings in cases with ethanol and methanol poisoning. A literature review was also performed to gather case series of electrocardiograms of ethanol and methanol poisoning. We synthesized the evidence and convened an expert panel to develop consensus-based recommendations.

Results: Based on the literature review and IPD analysis, cases of methanol poisoning often present later than 24 hours after alcohol consumption, emphasizing the importance of assessing patients with delayed referrals. Methanol-poisoned patients are frequently awake and responsive after 12 hours of ingestion, but symptoms alone do not reliably differentiate between the two poisonings. Declines in visual acuity and symptoms are strongly associated with methanol poisoning and should prompt immediate hospitalization and ophthalmic examination. Oliguria and anuria are more common in cases of methanol poisoning, warranting detailed urinary symptom assessment and close monitoring of urinary output. Electrocardiograms are essential for all alcohol poisoning patients, with non-sinus rhythm suggesting methanol poisoning. Arterial blood gas analysis, with low pH values, favors methanol poisoning. However, blood

Keywords:

Methanol, Poisoning, Ethanol, Intoxication

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sugar levels alone cannot definitively determine the type of poisoning. Laboratory point-ofcare tests to diagnose methanol in serum are urgently needed.

Conclusion: This literature review and consensus statement offer guidance for healthcare providers facing cases of alcohol poisoning where point-of-care testing methods of blood alcohol concentrations are unavailable. A comprehensive assessment, considering clinical history, physical examination, laboratory data, and specialized evaluations such as ophthalmic examination and electrocardiograms, is essential for accurately differentiating ethanol and methanol poisoning.

Introduction

ontaminated alcoholic beverages and industrial or household products can contain methanol and cause methanol toxicity in case of human consumption [1]. Illicit production is responsible for most methanol poisoning outbreaks [2], which have been reported in the USA [3], Malaysia [4], Norway [5], Libya, Kenya [6], and Iran [7]. Methanol poisoning can cause various symptoms, such as nausea, vomiting, headache, dizziness, abdominal pain, and visual disturbances. In severe cases, it can cause metabolic acidosis, central nervous system depression, coma, seizures, and even death [8]. The treatment for methanol poisoning involves administering ethanol or fomepizole [9]. At the same time, as there are few cases of this toxication worldwide and most are limited to Asian countries with boundaries on alcohol roles, the current practices are not well structured. While some outdated guidelines are available, like American [10], Iranian health authorities have faced this thread. Here, we aim to prepare a guideline based on the expertise of Iranian experts in the field. Differentiating ethanol from methanol poisoning is challenging for physicians in Iran due to the lack of available methanol detection measures in most centers [11].

Scope and purpose

The overall objective of the Iranian methanol poisoning guideline is to establish an evidence-based guide for the prevention, diagnosis, and treatment of methanol poisoning. This instruction will be developed based on the appraisal of guidelines for research & evaluation (AGREE) recommendations for writing clinical guidelines [12]. Expected benefits include early diagnosis and improved treatment. This study does not focus on preventative measures or legislation to stop a methanol poisoning outbreak.

Based on this, the population, intervention, comparison, outcome (PICO) method was used to structure study questions.

Population (P):

What are the differences between ethanol and methanol poisoning based on clinical presentation at arrival and time of consumption to referral?

What are the differences in physical examination findings between patients with ethanol and methanol poisoning on arrival?

What are the differences in laboratory results between patients with ethanol and methanol poisoning on arrival?

Intervention (I):

What are the most effective diagnostic interventions for individuals with methanol poisoning?

Comparison (C):

How does diagnosis differ from ethanol and methanol poisoning?

Outcome (O):

How reliable is evidence to differentiate the methanol poisoning from methanol poisoning?

Materials and Methods

Stakeholder involvement

A team of experts was enrolled in this study. The steering group members provided strategic guidance in the subject area. Focus group discussions were conducted with these stakeholders. The research team was responsible for selecting and reviewing the evidence. Some other experts formulated the final recommendations for synthesizing them into actionable guidance.

Specialists in relevant medical fields, such as toxicology, emergency medicine, and critical care, are a primary target audience. Health policymakers and regulators may also be part of the audience.

Search methods

This section will determine the methodologies used to integrate evidence for study questions. For questions about symptoms and clinical presentations, as epidemiologists only report the proportion of symptoms that could not be used as a source for differentiating patients, our objective would be gathering individual cases for individual patient data (IPD) meta-analysis with the primary aim of comparing ethanol and methanol poisoning symptoms, stratified based on the time from toxication. So, the inclusion criteria for studies were methanol poisoning in individual reported cases. Performing casecontrol studies based on IPD meta-analysis is previously used in the literature [13]. A MeaSurement tool to assess systematic reviews (AMSTAR2) guidelines were observed when performing the IPD meta-analysis [14]. The search was conducted using the following databases: PubMed, Embase, Scopus, and Google Scholar. PubMed search strategy would be as follows: ("Methanol poisoning" OR "methyl alcohol poisoning") AND ("clinical presentations" OR "symptoms" OR "signs" OR "manifestations") AND ("case series" OR "case report"); and same for ethanol poisoning: ("Ethanol poisoning" OR "ethyl alcohol poisoning") AND ("clinical presentations" OR "symptoms" OR "signs" OR "manifestations") AND ("case series" OR "case report").

Two independent reviewers screened the search results based on titles and abstracts. Full texts of potentially relevant articles were retrieved for further assessment. Any discrepancies in inclusion were resolved through discussion. Data were extracted from the included case reports using a standardized data extraction form. Two independent reviewers delicately extracted data and compared results.

In the case of ethanol poisonings, studies with concomitant intoxications or interacting medications were not sought. Only case reports of severe ethanol poisoning were included where admission was required, and three were high values of blood alcohol concentrations.

Demographic data included age and gender, while clinical data encompassed clinical status (e.g. coma, somnolent, lethargic/drowsy, awake), vital signs (e.g. heart rate, respiratory rate, systolic and diastolic blood pressure, temperature), and blood gas analysis, specifically pH levels. Clinical symptoms such as unconsciousness, abdominal pain, nausea and vomiting, diarrhea, decreased level of consciousness (LOC), aggressive behavior, visual symptoms, dyspnea, chest pain, headache (H/A), and the presence of asymptomatic cases were also recorded.

For questions about ECG findings of ethanol and methanol poisoning cases, our approach primarily focuses on a literature review. To achieve this, our inclusion criteria were centered on methanol poisoning ECG findings reports. There was a previous systematic review and meta-analysis; our study will be updated on the mentioned study [15]. This methodology follows a sub-group meta-analysis of proportional data. Data extraction included both demographic and clinical aspects. Demographic data included age and gender, and ECG findings were extracted.

CAse REport (CARE) guidelines were used to assess the risk of bias in case reports and case series studies [16]. The Newcastle-Ottawa scale was used to determine the quality of non-randomized studies [17].

Evidence was synthesized based on the statistical analyses in SPSS software, version 21. Descriptive statistics were used to summarize the data, including Mean±SD for continuous variables and frequency and percentage for categorical variables. Differences between the ethanol and methanol poisoning groups were assessed using appropriate statistical tests. For continuous variables (e.g. age), the independent samples t test or Mann-Whitney U test was employed based on the data distribution. Categorical variables (e.g. gender, clinical status) were compared using the chi-square test or Fisher exact test when appropriate. Subgroup analyses were performed for variables of interest, such as age groups or time since poisoning. Logistic regression analysis was conducted to investigate associations between specific clinical symptoms or outcomes (e.g. mortality) and poisoning type, adjusting for potential confounding variables. Receiver operating characteristic (ROC) analysis assessed the discriminatory power of pH levels in differentiating between severe ethanol and methanol poisoning. A significance level of P<0.05 was used to determine statistical significance. We used the Python programming language, along with the Pandas, Seaborn, and Matplotlib libraries, to create a heatmap using Seaborn.

Synthesized evidence was evaluated using the grading of recommendations assessment, development, and evaluation (GRADE) tool [18].

Results

Table 1 provides included studies in the individual patient level (IPD) meta-analysis related to symptoms and clinical presentation in cases of alcohol poisoning. There were 18 case reports of severe ethanol poisoning and 5 case series of methanol poisoning. The Table lists vari-

Table 1. Included studies in IPD meta-analysis of symptoms and clinical presentation

Author(s)	Study Design	Cases (No.)	Type of Poisoning	Country	(RoB)
Atassi et al. [19]	Case report	1	Ethanol	United States	++
Elliott & Hunter [20]	Case report	1	Ethanol	United States	+++
Pereska et al. [21]	Case report	1	Ethanol	Macedonia	+
Morgan et al. [22]	Case report	1	Ethanol	United States	++
Johnson et al. [23]	Case report	1	Ethanol	United States	++
Berild & Hasselbalch [24]	Case report	1	Ethanol	Denmark	+++
Sanap & Chapman [25]	Case report	1	Ethanol	Australia	+
Quispe et al. [26]	Case report	1	Ethanol	Valladolid	+
Nagashima et al. [27]	Case report	1	Ethanol	Japan	+
Wilson & Waring [28]	Case report	1	Ethanol	United Kingdom	+
Doyon & Welsh [29]	Case report	1	Ethanol	United States	+
Roberts et al. [30]	Case report	1	Ethanol	United Kingdom	++
Meyer et al. [31]	Case report	1	Ethanol	United Kingdom	++
Henry-Lagarrigue et al. [32]	Case report	1	Ethanol	United States	++
Bookstaver et al. [33]	Case report	1	Ethanol	France	+++
Tavolacci et al. [34]	Case report	1	Ethanol	Columbia	+
Brvar & Bunc [35]	Case report	1	Ethanol	France	+++
Piccini & Zaas [36]	Case report	1	Ethanol	Slovenia	+++
Alqurashi et al. [37]	Case series	6	Methanol	Saudi Arabia	+
Eskandrani et al. [38]	Case series	9	Methanol	Saudi Arabia	++
Kabli et al. [39]	Case series	9	Methanol	Saudi Arabia	++
Amin et al. [40]	Case series	8	Methanol	Bangladesh	+++
Liu et al. [41]	Case series	50	Methanol	United States	+++

RoB: Risk of bias; IPD: Individual patient data.

+: Low; ++: Moderate; +++: High.

ous case reports and case series, the number of cases, the country of origin, and the assessed risk of bias for each study.

Table 2, on the other hand, presents an overview of additional studies, each with its number of cases, study design, and risk of bias assessment. These studies encompass a range of study designs, including case-control, crossover, cross-sectional, prospective cohort, and retrospective studies, contributing to a comprehensive analysis.

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In our study comparing ethanol poisoning (n=18) and methanol poisoning (n=82) patients, ethanol poisoning patients had a similar mean age (45.55±20.53 years) compared to methanol poisoning patients (37.67±12.693 years), with no statistically significant (P=0.134). Gender distribution showed a higher proportion of males in both groups (61.11% in ethanol, 76.83% in methanol), but the difference was not statistically significant (P=0.234). In terms of clinical status, ethanol poisoning patients were more likely to present in an uncon-

Table 2. Additional studies for the analysis of electrocardiogram findings

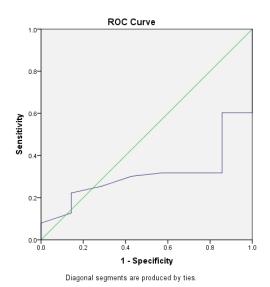
Author(s)	No.	Study Design	Risk of Bias
Cardy et al. [42]	10	Case-control	+
Priest et al. [43]	37	Case-control	+
Uyarel et al. [44]	10	Crossover study	+
Aasebo et al. [45]	84	Case-control	+
Drooshi et al. [46]	250	Cross-sectional	+
Brunner et al. [47]	3028	Prospective cohort	+
Total	3419		
Nikoo et al. [48]	356	Cross-sectional	+
Tabatabaiei et al. [49]	114	Cross-sectional	+
Jaff et al. [50]	9	Retrospective	+
Sanaei-Zadeh et al. [51]	42	Retrospective	+
Total	521		International Journal of

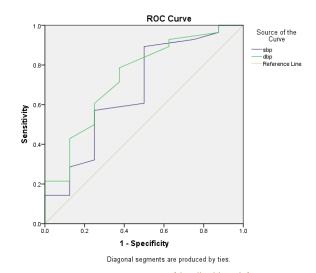
+: Low; ++: Moderate; +++: High.

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scious state (66.67%) compared to methanol poisoning patients (15.85%), and this difference was statistically significant (P=0.087). Other clinical parameters, such as heart rate (HR), respiratory rate (RR), systolic blood pressure (SBP), and temperature, showed no differences between the two groups (P>0.05). However, diastolic blood pressure (DBP) was significantly lower in etha-

nol-poisoning patients (P=0.024). Notably, the pH levels were significantly lower in the methanol poisoning group (7.082857 ± 0.254) compared to the ethanol group (7.26 ± 0.104) (P=0.002). Additionally, the mortality rate was significantly higher in the methanol poisoning group (39.02%) compared to the ethanol group (5.56%) (P=0.0001) (Table 3).





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between methanol and ethanol

Figure 1. ROC analysis of pH (left side) and blood pressure (right side) for differentiation between methanol and ethanol poisoning

ROC: Receiver operating characteristic.

Table 3. Characteristics of the study populations

-		Mean±SD/No. (%)			
Characteristics —		Ethanol Poisoning (n=18)	Methanol Poisoning (n=82)	Р	
Age (y)		45.55±20.53	37.67±12.693	0.134*	
Gender	Male	11(61.11)	63(76.83)	0.234	
Gender	Female	7(38.89)	19(23.17)	0.234	
	Coma	12(66.67)	13(15.85)		
Status	Somnolent	3(16.67)	3(3.66)	0.087	
Status	Lethargic/Drowsy	2(11.11)	3(3.66)		
	Awake	1(5.56)	12(14.63)		
	Weight (kg)	58.5±10.847	NR	-	
HR	(bpm) [35 data]	92.14286±25.3	93.60714±20.11	0.871	
RR (bre	aths/min) [20 data]	17.25±7.80	24.5±10.9	0.234	
SBP (nm Hg) [36 data] 104.37±33.85		126.4286±28.82	0.075	
DBP (mm Hg) [36 data]	61.375±19.07	78.07143±17.16	0.024	
Temperature (°C) [16 data]		35.09±1.93	36.68333±1.135	0.09	
	Bs [10 data]	133±36.51	108.9000±36.08	-	
	pH [70 data]	7.26±0.104	7.082857±0.254	0.002	
Death		Death 1(5.56)		0.0001	

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Abbreviations: HR: Heart rate; RR: Respiratory rate; SBP: Systolic blood pressure; DBP: Diastolic blood pressure; Bs: Blood sugar; NR: Not reported.

There are statistically significant differences in the distribution of the presenting pattern of poisoning among different types of alcohol poisoning categories based on chi-square (P=0.047). It seems that methanol poisoning patients are more frequently presenting awake and responsive than ethanol poisoning patients, later than 12 hours after first ingestion (P=0.025) (Table 4).

Significant differences were observed between the two groups in several key symptoms. Notably, a significantly higher proportion of patients with ethanol poisoning presented with unconsciousness (66.67%) compared to those with methanol poisoning (14.63%) (P<0.001). Additionally, the prevalence of aggressive behavior was significantly higher in the methanol poisoning group (29.27%) compared to the ethanol poisoning group (5.56%) (P=0.038). Visual symptoms were absent in the ethanol poisoning group but were present in 26.83% of methanol poisoning cases, showing a significant differ-

ence (P=0.010). Other symptoms, such as abdominal pain, nausea and vomiting, diarrhea, decreased LOC, dyspnea, chest pain, and headache, did not exhibit statistically significant differences between the two groups (P>0.05) (Table 5).

Based on a ROC analysis of pH for differentiation of severe ethanol poisoning and methanol poisoning, pH had an area under the curve (AUC) of 0.310 (P=0.10). It cannot be utilized for differentiation, as well as the systolic blood pressure with an AUC of 674 (P=0.128) (Figure 1). Meanwhile, diastolic blood pressure had an AUC of 0.739 (P=0.049), showing promising differentiating performance.

Discussion

Based on the provided literature review, IPD metaanalysis, and literature review of studies on electrocar-

Table 4. Crosstabulation of clinical presentation, timing of referral, and type of toxicity

Status		As Time Passed the Poisoning (h)			
		<12	12-24	24-72	- Р
Coma	Ethanol	7	1	1	0.508
	Methanol	6	0	2	
Somnolent	Ethanol	2	0	0	
	Methanol	0	0	0	-
Lethargic/Drowsy	Ethanol	3	0	0	
	Methanol	1	0	0	-
Awake	Ethanol	0	1	0	0.025
	Methanol	0	0	4	
Total	Ethanol	10	2	1	0.047
	Methanol	7	0	6	0.047

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diogram findings, we will offer recommendations for differentiating ethanol poisoning from methanol poisoning.

The low quality of evidence suggests that these recommendations should be considered, but they may benefit from further research and validation. Our first recommendation is in line with the body of literature. Williams et al. reported that most patients with methanol poisoning start having symptoms 24 hours later, and mixed ethanol poisoning might delay the symptom's manifestation [55]. A literature review and meta-analysis of 2327 patients in 67 studies showed that the time from methanol

Table 5. Comparison of symptoms between ethanol and methanol poisoning patients

Communica and	No.			
Symptoms	Ethanol Poisoning (n=18)	Methanol Poisoning (n=82)	Р	
Unconsciousness	12(66.67)	12(14.63)	<0.001	
Abdomen	2(11.11)	7(8.54)	0.257	
Nausea and Vomiting	3(16.67)	16(19.51)	0.847	
Diarrhea	1(5.56)	0(0)	0.969	
LOC decrease	3(16.67)	25(30.49)	0.384	
Getting aggressive	1(5.56)	24(29.27)	0.038	
Visual symptoms	0(0)	22(26.83)	0.010	
Dyspnea	0(0)	7(8.54)	0.128	
Chest pain	0(0)	3(3.66)	0.463	
Headache	0(0)	1(1.22)	0.987	
Asymptomatic	0(0)	10(12.2)	0.201	

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Q1. What are the differences between ethanol and methanol poisoning based on the clinical picture on arrival and time of drinking to referral?

1. Based on the review of results of individual patient level (IPD) meta-analysis, a higher percentage of methanol poisoning cases present later than the first day passing the drink compared to ethanol poisoning cases.

Recommendation: Healthcare providers should carefully assess patients referred to the hospital for more than 24 hours for alcohol consumption.

Quality of evidence: 0000

2. Methanol poisoning patients are more frequently presenting awake and responsive than ethanol poisoning patients, later than 12 hours of first ingestion.

Recommendation: Being awake and responsive could not be observed as a good clinical picture, and all patients with a history of alcohol consumption should be further evaluated in cases of symptoms, timings, and clinical findings.

Quality of evidence: ⊕⊕00

3. Our IPD analysis did not show robust evidence for symptomatology, and there is no subjective sign for differentiating methanol poisoning from ethanol poisoning, except for the visual impairment symptoms.

Recommendation: A detailed history of exposure, timing, and sources of alcohol is needed to interpret the signs of the patient to decide on the diagnosis of ethanol or methanol poisoning.

Quality of evidence: 0000

4. Symptoms of oliguria and anuria are frequently seen in methanol poisoning cases. In contrast, ethanol poisoning may rarely manifest with these symptoms (only in some cases reports of rhabdomyolysis after ethanol poisoning [52] and concurrent NSAID and ethanol consumption [53] or chiroptic chronic alcohol users [54]).

Quality of evidence: 0000

Recommendation: A detailed history should be obtained from patients with alcohol poisoning focusing on the urinary symptoms, and urinary output should be closely documented. Further laboratory exams should be considered to manage any acute renal failure.

Symbol: OOO: High; OOO: Moderate; OOO: Low; OOO: Very low.

poisoning to hospital referral ranges between 3 to more than 48 hours [56]. While there is a substantial number of studies on ethanol and methanol poisoning individually, there is limited direct comparative research on the symptomatology of these two types of poisoning cases in human patients. Most studies separately focus on each poison's characteristics, treatment, and outcomes. However, there was an animal model study that found that methanol is more lethal than ethanol, particularly when ingested in its pure form [57]. In the case of the clini-

cal picture on arrival, Methanol's central nervous system (CNS) depressive effects are primarily attributed to its metabolites. However, ethanol has direct effects on neurotransmitters and causes direct CNS depression [58].

Ocular examination with assessments of visual acuity, pupillary abnormalities, fundoscopic examinations, and monitoring for visual symptoms is critical for all patients who have a history of alcohol exposure. For alert patients, a detailed examination is needed, including assessing visual acuity, vision and color vision testing

Q2. What are the differences in physical examinations of patients with ethanol and methanol poisoning on arrival?

1. Vital signs evaluation is a critical step in facing alcohol poisoning patients, while normal blood pressure could not be related, as even methanol poisoning cases might have a higher DBP than ethanol poisoning cases but have similar systolic blood pressures.

Recommendation: A normal blood pressure reading alone may not reliably distinguish between methanol and ethanol poisoning cases.

Quality of evidence: 0000

2. Visual acuity decreases or visual symptoms are directly associated with methanol poisoning.

Recommendation 1: Any patient reporting visual disturbances or experiencing a decline in visual acuity after alcohol consumption should be hospitalized promptly and be treated for potential methanol exposure.

Recommendation 2: Ocular examination should not be limited to patients with ocular symptoms. All patients with suspected methanol poisoning should undergo a detailed ophthalmic examination.

Quality of evidence: 0000

3. Based on our literature review of observational studies, it's evident that ethanol poisoning cases predominantly show symptoms related to sinus rhythm abnormalities, particularly Sinus bradycardia, and tachycardia, while methanol poisoning cases are more characterized by abnormalities in QTc interval duration, ST-segment depressions/elevation, and T-wave changes. Additionally, methanol poisoning cases exhibit a broader range of ECG abnormalities compared to ethanol poisoning cases.

Recommendation: Electrocardiogram is required for all patients with alcohol poisoning referring to hospitals. In patients with non-sinus rhythm, physicians should be suspected of methanol poisoning.

Quality of evidence: 0000

Notes: High; $\Theta\Theta\Theta$ O: Moderate; $\Theta\Theta$ OO: Low; Θ OOO: Very low.

Q3. What are the differences in laboratory analyses of patients with ethanol and methanol poisoning on arrival?

1. Based on our IPD meta-analysis, atrial blood gas analyses with low PH values favor methanol poisoning, while our data was not satisfactory in bringing a clinically usable cut-off point. Based on the literature, ethanol poisoning mostly manifests as mild metabolic acidosis, while acidosis in methanol poisoning is severe [11].

Recommendation: An atrial blood gas analysis is crucial laboratory data needed for all alcohol poisoning patients.

Quality of evidence: 0000

2. Ethanol poisoning is known to be associated with hypoglycemia [11, 63], but we only found 10 cases with available blood sugar levels for analysis.

Recommendation: Literature shows more frequent hypoglycemia in ethanol poisoning and hyperglycemia in methanol poisoning; this is not reliable as no case-control or comparative studies exist. While blood sugar assessment is a necessary part of alcohol poisoning assessment, we cannot use it as a diagnostic tool.

Quality of evidence: 0000

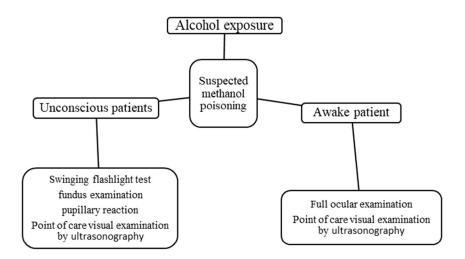
3. Methods of diagnosing methanol poisoning in blood are recently developed but are not widely available or approved for clinical application.

Quality of evidence: 0000

Notes: High; OOOO: Moderate; OOOO: Low; OOOO: Very low.

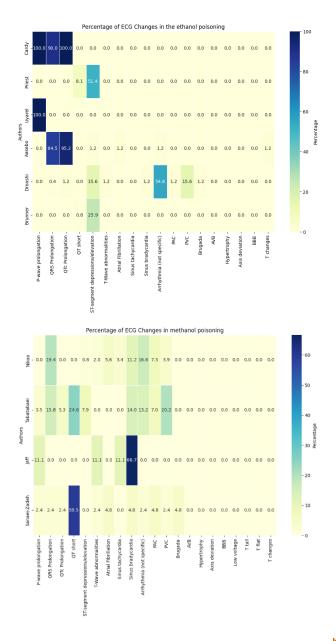
chart, eye movements, measuring intraocular pressure, pupillary reactions, and conducting a fundoscopic examination. For unconscious patients, a swinging flashlight test, a fundus examination, and pupillary reaction examinations should be performed at the bedside [59]. Point-of-care ocular ultrasonography can be used for both types of patients [60]. Studies have proposed daily examinations for patients hospitalized with methanol poisoning [59]. Optic disc atrophy was found in all one hundred patients reviewed in a retrospective study [61]. In case of positive findings, magnetic resonance imaging should be used to assess optic fiber nerves and potential brain edema [62] (Figure 2).

Ten observational studies were included studies in the literature review of ECG findings, with 3419 ethanol poisoning and 521 methanol poisoning cases. In the groups of ethanol poisoning (n=3419) and methanol poisoning (n=521), the most frequent ECG finding varies significantly. In the ethanol poisoning group, the most frequent finding is sinus bradycardia, with a prevalence of 63.73%, followed by sinus tachycardia at 24.10%. In contrast, the methanol poisoning group exhibits a higher prevalence of QTc prolongation at 19.58%, followed by ST-segment depressions/elevation and T changes at 14.59% and 8.06%, respectively (Figure 3).



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Figure 2. Schematic diagnostic approach for visual assessment of suspected cases



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Figure 3. ECG findings of methanol and ethanol poisoning cases from the IPD meta-analysis

Notes: Each cell in the heatmap represents the intersection of a row and column and is colored based on its value. The color intensity reflects the magnitude of the value. Numbers indicate the actual values for that combination of author and ECG change.

Methanol is metabolized in the body to formaldehyde and formic acid, which can disrupt glucose metabolism and increase blood sugar levels. Hypoglycemia is not commonly associated with methanol poisoning. Methanol poisoning more often leads to hyperglycemia and is known to be a prognostic factor in methanol poisoning [64]. Even in some circumstances, it might lead to diabetic ketoacidosis [65, 66].

Researchers have developed a simple and rapid diagnostic test for methanol poisoning using an enzyme called formate oxidase (FOX), which can detect formate in the blood. The test showed high sensitivity and specificity, with no false positives detected, making it a promising tool for point-of-care diagnosis, especially in low-resource settings where laboratory equipment may not be readily available [67]. Researchers have developed a bedside test strip that can detect formate, a toxic metabolite of methanol, from a single drop of blood, providing

a rapid and simple diagnosis of methanol poisoning. In a clinical trial, the test strip correctly identified a 61-yearold patient with methanol poisoning, allowing for prompt treatment and confirming the potential of this technology to revolutionize the management of methanol poisoning worldwide [68]. In another study, a commercially available dipstick test, originally designed to detect ethanol in saliva, was found to rapidly and reliably detect methanol in the serum of mice at concentrations as low as 5 mg/ dL, offering a potential rapid diagnostic tool for emergency physicians [69]. Kubáň and colleagues developed a rapid and direct method for detecting formate in blood serum using capillary electrophoresis with contactless conductometric detection, which can diagnose methanol intoxication in approximately 1 minute with high sensitivity [70]. Gas chromatography/mass is also used to detect and quantify methanol in blood [71]. But these methods are not broadly investigated for clinic or are so expensive.

Ethanol's impact on blood sugar levels involves inhibiting the release of glucose from the liver and enhancing insulin secretion, which can lead to a drop in blood sugar levels. A study [72] shows that hypoglycemia can occur in children following ethanol ingestion. Research in rural Uganda [73] has shown instances of alcohol-related hypoglycemia, particularly in communities where alcohol consumption is prevalent.

Conclusion

In conclusion, the differentiation between ethanol and methanol poisoning presents a clinical challenge, and several key clinical and laboratory factors must be considered. Methanol poisoning cases tend to present later than the first day after alcohol consumption and are often characterized by awake and responsive patients, especially after 12 hours of ingestion. Notably, symptoms alone do not provide a reliable means of differentiation, emphasizing the importance of obtaining a detailed history of exposure, timing, and alcohol sources. Physical examinations may not reliably distinguish between the two types of poisoning, with vital signs such as blood pressure showing limited discriminatory value. Visual acuity and ophthalmic examinations are critical in cases of suspected methanol poisoning. Electrocardiograms are essential for all alcohol poisoning patients, particularly those with non-sinus rhythm, as methanol poisoning is associated with a broader range of ECG abnormalities. Laboratory analyses, including arterial blood gas measurements and blood sugar assessments, are valuable in the evaluation, but blood sugar levels alone cannot definitively diagnose the type of poisoning. A comprehensive evaluation combining clinical history, physical examination, and laboratory data is crucial for accurately differentiating ethanol and methanol poisoning.

Study limitations

The study's reliance on case reports and case series of severe ethanol and methanol poisoning may introduce selection bias, as these cases may not be representative of all alcohol poisoning presentations. This may lead to an overemphasis on statistical significance rather than clinical relevance. Furthermore, the sample size of ethanol poisoning cases (n=18) was relatively small compared to methanol poisoning cases (n=82), which may limit the power to detect significant differences between the two groups.

Ethical Considerations

Compliance with ethical guidelines

This article is a review with no human or animal sample.

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Authors' contributions

All authors equally contributed to preparing this article.

Conflict of interest

The authors declared no conflict of interest.

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