

# Outcome Prediction in Acutely Intoxicated Patients Admitted to Intensive Care Unit

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## Abstract:

**Background:** Acute poisoning represents a significant proportion of intensive care unit admissions. Even though the overall mortality may be low, they may consume considerable intensive care unit (ICU) resources. Early diagnosis and rapid initiation of appropriate therapy in emergency department and ICU are critical for decreasing hospital morbidity and mortality in poisoned patients. The objective of this study is to determine predictors of outcome of acutely intoxicated patients in intensive care unit which may improve the course of management and decide the pathway of care.

**Methodology:** This is an observational cross sectional retrospective study of 321 acutely intoxicated patients admitted to intensive care unit of Poison Control Center (PCC) in hospitals of Ain Shams University. Information was collected from the sheets and computerized data base of the patients after obtaining the permission of the director of PCC and the regional ethics committee. The results were revised, coded and organized for statistical analysis.

**Results:** The total number of acutely intoxicated patients admitted to ICU of PCC in hospitals of Ain Shams University in the selected period of study was 321 patients. There were 265 patients (82.6%) survived with no complications, 30 patients (9.4%) survived but developed complications and 26 patients (8%) died. The study showed statistically significant difference between uncomplicated, complicated and dead cases as regards mode of toxicity, causative Agent, heart rate, systolic blood pressure, respiratory rate, skin discoloration, Glasgow coma scale (GCS), blood pH, emesis or lavage, activated charcoal, dialysis, endotracheal intubation, mechanical ventilation and dopamine therapy.

**Conclusion:** It could be detected by statistical analysis that causative agent, heart rate, systolic blood pressure, respiratory rate, blood pH, mechanical ventilation and dopamine therapy were significant outcome predictors of acutely poisoned patients in intensive care unit. From the previous results, our study recommends that these predictors of outcome should be assessed routinely and as early as possible to evaluate the severity, improve the course of management and deciding the pathway of care.

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## Key words

Predictors of outcome, ICU, GCS, complications.

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## Introduction

Acute poisoning represents a significant proportion of intensive care unit admissions and even though the overall mortality may be low, they may consume considerable intensive care unit resources (Singh et al., 2011).

Toxicologic conditions are encountered in patients with acute poisoning in intensive care unit due to intentional or unintentional exposure to therapeutic or illicit drugs. Additionally, toxicities related to medical interventions may occur in hospitalized patients (Philip and Janice, 2008).

Intensive care unit management of poisoned patient requires rapid diagnosis and supportive care while in some cases providing specific antidotal treatment (Donna et al., 2011).

Accurate prognostic information for critically ill patients could help clinicians with decisions like whether and when patients might benefit from intensive care. From the perspectives of families, prognostic information is welcome in discussions about the benefits of intensive care and completeness of this information is

an independent determinant of family satisfaction (Heffner and Barbieri, 2000; Heyland et al., 2002).

### Aim of the work

The aim of this work is to determine the predictors of outcome of acutely intoxicated patients in intensive care unit of the Poison Control Center in hospitals of Ain Shams University.

### Methodology

This is an observational cross sectional retrospective study of 321 acutely intoxicated patients in ICU of PCC in hospitals of Ain Shams University in the period from 1/7/2011 to 31/1/2012. An approval from the Ethical Committee and the permission of the director of PCC were taken. Information was collected from the sheets and computerized data base of the patients with the confidentiality of these records. The basis of acute poisoning diagnosis was positive history, clinical pictures and initial laboratory tests specific to certain poisons.

### The following data was collected from the sheet of each patient:

- (A) Demographic Data.
- (B) Toxicological data.
- (C) Physical findings on admission.
- (D) Reported investigational data and treatment received
- (E) Outcome of the patients:

The patients were classified according to the outcome into 3 groups' uncomplicated, complicated and dead.

### Statistical Analysis

The obtained results were revised, coded and organized for statistical analysis by SPSS. Mean and standard deviation was done for numerical data. Frequency and percentage were obtained for non-numerical data.

Comparison between outcome groups was tested using KruskalWallis Test. Linear regression analysis was used to identify significant predictors of outcomes. P-value was considered statistically significant if  $< 0.05$ .

### Results

The total number of acutely intoxicated patients admitted to ICU of PCC in hospitals of Ain Shams University in the selected period of study was 321 patients. In the current study, 36.7% of patients were 15 to  $< 25$  years and 6.9% of them were 5 to  $< 15$  years old, 52.6% of

patients were females and 47.4% were males (table, 1). Most of poisoning cases were due to pharmaceutical agents (36.5%), followed by organophosphorus compounds (15.3%), tramadol (14.3%), snake bites (5.7%), carbon monoxide (4.9%) and corrosives (4%), other toxic agents (19.3 %) (kerosene, scorpion, alcohol, hydrogen sulphide, ciguatera, PPD{ Paraphenylenediamine}) (table, 2). As shown in table (3) 52% of cases were suicidal, route of exposure in 84.7% of patients was oral and the mean delay time of patients was 6.09 hours while 54.9% of cases showed delay time 2 - 6 hours.

The outcome of the patients as shown in table (4) was as the following: 265 patients were survivors with no complications representing about (82.6%), 30 patients were survivors with complications representing about (9.4%) and 26 patients died representing about (8%). The most frequently encountered complications were dysphagia, bleeding and anemia in (26.6%), followed by intermediate syndrome in (23.3%), cognitive dysfunction in (16.6%), rhabdomyolysis in (10%), renal failure in (6.6%), then, hepatotoxicity in (3.3%), disseminated intravascular coagulation in (3.3%), myopathy in (3.3%), pneumonia in (3.3%) and pneumothorax in (3.3%) (table, 5). The study showed statistically significant difference between uncomplicated, complicated and dead cases as regards mode of toxicity, causative Agent, heart rate, systolic blood pressure, respiratory rate, skin discoloration, Glasgow coma scale (GCS), blood pH, emesis or lavage, activated charcoal, dialysis, endotracheal intubation, mechanical ventilation and dopamine therapy (table, 6). No statistically significant differences in age, sex, route of exposure, delay time, body temperature, random blood glucose, serum potassium levels, multiple doses activated charcoal, alkalization of urine and antidotes administration among uncomplicated, complicated and dead cases. Table (7) showed that predictors of outcome were heart rate, systolic blood pressure, respiratory rate, causative agent, blood pH, mechanical ventilation and dopamine therapy.

**Table (1) statistical analysis: Number and percentage of the studied patients as regard sociodemographic data.**

		Number	Percentage %
Age group (years)	<5	68	21.2%
	5-	22	6.9%
	15-	118	36.7%
	25-	58	18%
	35-	24	7.5%
	$\geq 45$	31	9.7%
Sex	Male	152	47.4%
	Female	169	52.6%

**Table (2) statistical analysis: Number and percentage of the studied patients as regard causative agents of poisoning.**

The causative agents	Number	Percentage %
Pharmaceutical agents	117	36.5%
Organophosphates	49	15.3%
Tramadol	46	14.3%
Snake bite	18	5.7%
Carbon monoxide	16	4.9%
Corrosives	13	4%
Other toxic agents	62	19.3%
Total	321	100%

**Table (3) statistical analysis: Number and percentage of mode of toxicity, route of exposure and delay time in the studied patients.**

		Number	Percentage %
Mode of toxicity	Accidental	116	36.2%
	Suicidal	167	52%
	Therapeutic error	38	11.8%
Route of exposure	Oral	272	84.7%
	Inhalation	20	6.3%
	IM or SC	4	1.3%
	Bite or Sting	22	6.8%
	Dermal	3	0.9%
Delay time	<2	58	18%
	2-6	176	54.9%
	>6	87	27.1%
	Mean $\pm$ SD	6.09 $\pm$ 1.5	
	Range	1 – 120	

IM: Intramuscular –SC: Subcutaneous

**Table (4) statistical analysis: Number and percentage of the outcome of the studied patients.**

Outcome	Frequency	Percentage
Survived (with no complications)	265	82.6%
Survived (with complications)	30	9.4%
Died	26	8%
Total	321	100%

**Table (5) statistical analysis: Number and percentage of the studied patients as regard complications**

Complications	Number	percentage
Dysphagia, bleeding and anemia	8	26.6%
Intermediate syndrome	7	23.3%
Cognitive dysfunction	5	16.6%
Rhabdomyolysis	3	10%
Renal failure	2	6.6%
Hepatotoxicity	1	3.3%
Myopathy	1	3.3%
DIC	1	3.3%
Pneumonia	1	3.3%
Pneumothorax	1	3.3%
Total	30	100%

DIC: Disseminated intravascular coagulation

**Table (6) statistical analysis: Kruskal Wallis test of mode of toxicity, causative Agent, heart rate, systolic blood pressure, respiratory rate, skin changes, GCS, blood pH, emesis or lavage, activated charcoal, dialysis, endotracheal intubation, mechanical ventilation and dopamine therapy in relation to outcome of the studied patients.**

Kruskal Wallis test		Uncomplicated n.= 265		complicated n. = 30		Dead n. = 26		P- value
		N	%	N	%	N	%	
Mode of toxicity	Accidental	86	32.4%	17	56.7%	13	50%	0.043*
	Suicidal	147	55.5%	10	33.3%	10	38.5%	
	Therapeutic error	32	12.1%	3	10%	3	11.5%	
The causative agent	Organophosphates	35	13.2%	7	23.3%	7	26.9%	0.000*
	Other agents	230	86.8%	23	76.7%	19	73.1%	
Heart rate	Normal	193	72.9%	23	76.7%	13	50%	0.039*
	Abnormal	72	27.1%	7	23.3%	13	50%	
Systolic B.P	Normal	239	90.2%	22	73.4%	12	46.2%	0.000*
	Abnormal	26	9.8%	8	26.6%	14	53.8%	
Respiratory rate	Normal	223	84.2%	19	63.4%	6	23.1%	0.000*
	abnormal	42	15.8%	11	36.6%	20	76.9%	
Skin discoloration	Pallor or cyanosis	16	6%	3	10%	12	46.2%	0.000*
	None	249	94%	27	90%	14	53.8%	
GCS	GCS ≤ 8	69	26%	8	26.7%	21	80.7%	0.000*
	GCS > 8	196	74%	22	73.3%	5	19.3%	
Blood pH	None	166	62.6%	12	40%	0	0%	0.000*
	applied Normal	48	18.1%	8	26.6%	1	3.8%	
	applied Acidosis	48	18.1%	8	26.6%	24	92.4%	
	applied Alkalosis	3	1.1%	2	6.8%	1	3.8%	
Emesis or lavage	applied	134	50.6%	6	20%	5	19.2%	0.005*
	None	131	49.4%	24	80%	21	80.8%	
Activated charcoal	applied	95	35.9%	0	0%	1	3.8%	0.000*
	None	170	64.1%	30	100%	25	96.2%	
Dialysis	applied	2	0.75%	1	3.3%	2	7.7%	0.018*
	None	263	99.2%	29	96.7%	24	92.3%	
ETI	applied	53	20%	8	26.6%	24	92.3%	0.000*
	None	212	80%	22	73.4%	2	7.7%	
Mechanical Ventilation	applied	25	9.4%	8	26.6%	20	76.9%	0.000*
	None	240	90.6%	22	73.4%	6	23.1%	
Dopamine therapy	applied	1	0.38%	4	13.3%	10	38.4%	0.000*
	None	264	99.6%	26	86.7%	16	61.6%	

\*P is considered statistically significant if  $\leq 0.05$ , GCS: Glasgow Coma Scale, ETI: Endotracheal intubation

**Table (7) statistical analysis: Linear regression analysis of statistically significant parameters in relation to outcome.**

Linear regression analysis	Unstandardized C.		Standardized C.	P- value
	B	Standard Error	Beta	
Mode of toxicity	-.033	.037	-.038	0.376
Causative Agent	-.027	.005	-.264	0.000*
Heart rate	.116	.056	.089	0.039*
Systolic blood pressure	-.208	.074	-.125	0.005*
Respiratory rate	-.146	.063	-.103	0.022*
Skin discoloration	.075	.093	.037	0.419
Conscious level	.004	.011	.027	0.739
Blood pH	.085	.038	.129	0.024*
Emesis or lavage	-.053	.030	-.072	0.078
Activated charcoal	.050	.060	.038	0.405
Dialysis	.220	.195	.046	0.260
Endotracheal intubation	-.004	.085	-.003	0.963
Mechanical ventilation	.314	.097	.196	0.001*
Dopamine therapy	.874	.123	.311	0.000*

C: Coefficients, \*P is considered statistically significant if  $< 0.05$ .

## Discussion

Acute poisoning is a frequent etiology of admission to emergency departments (ED) and always requires treatment in the intensive care unit (ICU). Early diagnosis and rapid initiation of appropriate therapy in ED and ICU are critical for decreasing hospital morbidity and mortality in patients with acute poisoning (Islambulchilar et al., 2009).

In this study 265 patients (82.6%) improved and were discharged without complications, 30 patients (9.4%) survived but with complications and 26 patients (8%) died during treatment in the ICU. These results differ from those reported by Hassanian et al., (2007) where ICU mortality in poisoned patients was 18.6%. On the other hand, two studies done in Germany and Hong Kong reported that mortality of poisoning in ICU were respectively 0.7 and 3% (Schwake et al., 2009). Lam et al., (2010) attributed these different results to the extreme variation in reported mortality and criteria of ICU admission across hospitals and countries.

In our study, the most frequently encountered complications were dysphagia, bleeding and anemia followed by intermediate syndrome and cognitive dysfunction. The most frequent agents involved were corrosives, organophosphorus compounds and carbon monoxide. Taghaddosinejad et al., (2012) reported that the most common complications of severe poisoning cases in ICU were coma, rhabdomyolysis and aspiration pneumonia where the most frequent agents involved were benzodiazepines and tricyclic antidepressant.

The present study showed that there was statistically significant difference between the 3 groups in the mode of toxicity where nearly half of complicated cases and dead cases were accidental. These findings are similar to those of Lee et al., (2008) who stated that there was statistically significant difference between survivors and non-survivors as regard mode of toxicity. Agran et al., (2001) and Khadka and Ale, (2005) explained the high incidence of complicated and dead cases in accidentally intoxicated patients as accidental poisoning is common in children under five years. In the current study, statistically significant difference was among uncomplicated, complicated and dead cases as regard causative agents. Organophosphorus compounds and carbon monoxide carried the highest mortality risk (26.9%) followed by corrosives. This is in accordance with Goel and Aggarwal, (2007) and Malangu and Ogunbanjo, (2009) who reported that more than half of dead cases in their study due to poisoning with Organophosphorus compounds and carbon monoxide. Davies et al., (2008) mentioned that organophosphorus poisoning had high mortality, most patients died from cardiorespiratory failure and many patients had cardiorespiratory arrests after admission. Significant exposures of Carbon monoxide cause hypotension, dysrhythmia, ischemia, infarction, and, in extreme cases, cardiac arrest (Kao and Nanagas, 2004). The reverse was reported by Lund et al., (2012) where more than half of

the mortality cases caused by substances of abuse and Litovitz et al., (2001) mentioned that drugs like analgesics, sedatives and antidepressants were associated with high mortality rate. Disparities between the results of these studies are most likely due to the difference in access, as well as the ages of affected victims (Saddique, 2001).

As regards heart rate, systolic blood pressure, respiratory rate and skin discoloration there was statistically significant difference between the 3 groups in the present study. These results go with those of Yu et al., (2012) who mentioned that there were significant differences between survivors and non-survivors as regard heart rate, systolic blood pressure, and respiratory rate. He denoted that the patients with extremely abnormal vital signs had the greatest risk of in-hospital mortality. Also, Baumann and Strout, (2007) found that the Emergency Severity Index triage score, which incorporates vital signs into its algorithm, accurately assesses the risk of patients for hospitalization and mortality. The decrease in blood pressure can then cause blood vessels to contract resulting in a pale or bluish skin color and seizures in advanced cases with organ failure, unconsciousness and death (Hildebrandt et al., 2002).

As regard GCS there was statistically significant difference among uncomplicated, complicated and dead patients. There were 80.7% of dead cases and 26.7% of complicated cases had GCS less than or equal to 8. This was similar to results observed by Basar et al., (2011) where they found that the mean GCS values were 4 for patients who had died and 13 for discharged patients and there was a statistical correlation between GCS values and mortality, and low GCS values indicate the potential for respiratory insufficiency development and bad prognosis in acutely organophosphorus (OP)-poisoned patients admitted within 24 hours after exposure. Also Budhathoki et al., (2009) stated that GCS < 8 had been more associated with mortality in poisoned children. Moreover Russell and Shobhan, (2009) found that a GCS  $\leq$  8 is a useful indicator for the requirement of endotracheal intubation in toxic coma.

As regard blood pH, the present study revealed that there was significant difference between uncomplicated, complicated and dead cases. Similarly, Louriz et al., (2009). Singh et al., (2011) mentioned that presence of acidosis was related to poor outcome in aluminum phosphide poisoning. Also, Hong et al., (2000) mentioned that patients with metabolic acidosis were also related to higher paraquat fatality and Hampson and Hauff, (2008) stated that in carbon monoxide-poisoned patients the mortality was related to the severity of metabolic acidosis.

Statistically significant difference was among uncomplicated, complicated and dead patients in the present study as regard decontamination, where emesis or gastric lavage was used in half of uncomplicated cases and 19.2% of dead cases. Regarding activated charcoal,

35.9% of uncomplicated cases and 3.8% of dead cases received activated charcoal. Gastric decontamination including lavage and activated charcoal administration are known to decrease the absorption of some ingested poisons provided they are done within one-hour post ingestion (Chiu et al., 2011).

Statistically significant difference was among uncomplicated, complicated and dead patients as regard usage of dialysis in the current study. Dialysis may be indicated for life-threatening ingestions involving water-soluble substances of low molecular weight and severe cases of poisoning (Zimmerman, 2003).

There was statistically significant difference among non complicated, complicated and dead patients as regard emergency treatment, where endotracheal intubation and mechanical ventilation were needed in the majority of dead cases. 38.4% of dead cases, 13.3% of complicated cases needed dopamine therapy. These results are similar to Singh et al., (2011) reported that most of dead poisoned cases in ICU cases received inotropic support and needed mechanical ventilation.

In the current study, linear regression analysis identified that causative agent, heart rate, systolic blood pressure, respiratory rate, blood pH, mechanical ventilation and dopamine therapy are significant predictors of outcome while mode of toxicity, skin discoloration, coma scale, emesis or lavage, activated charcoal, dialysis and endotracheal intubation failed to be a predictor. These results coincide with those of Lam et al., (2010) and Malangu and Ogunbanjo (2009) who mentioned that the causative agent vary from place to place, and over time, due to the availability and use of different chemicals and other poisoning agents and doubling the incidence of pesticide poisoning in developing countries during the past decades which accounts for a large number of fatal outcomes despite increased ICU facilities. Also, our results coincide with those of Hsin et al., (2008) who reported that abnormal vital signs were significant predictors of poisoning-related fatalities. Similarly, Jayashree and Singhi, (2011) mentioned that hypotension at admission was the most significant predictor of death in children admitted to the ICU with acute poisoning. Hu et al., (2010) mentioned that factors such as hypotension and respiratory failure upon presentation can predict overall poisoning related fatality in emergency department poisoned patients. And, Hampson and Hauff, (2008) mentioned that arterial blood pH was significant predictors of death in carbon monoxide poisoning. Nejad et al., (2012) mentioned that there was negative relation between mechanical ventilation need and outcome in aluminum phosphide intoxication. Rocker et al., (2004) mentioned that the use of inotropic agents or vasopressors was associated with a higher risk of death in ICU than no use of inotropic agents or vasopressors. Risk stratification and detection of high risk patients are very important in provision of health care with the limited resources in the emergency department (Lee et al., 2008).

## Conclusion

The causative agent, heart rate, systolic blood pressure, respiratory rate, blood pH, mechanical ventilation and dopamine therapy were significant predictors of outcome of acutely intoxicated patients in intensive care unit. Their routine assessment as early as possible can be beneficial to assess the poisoning severity, improve the management course and decide the pathway of care.

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## الملخص العربي

### التنبؤ بالنتيجة لمرضى التسمم الحاد بوحدة العناية المركزة بمركز علاج التسمم بجامعة عين شمس

اسماء احمد عساف و منال على عبد الكريم<sup>١</sup> و مروة احمد حسب النبی<sup>٢</sup>

**مقدمة البحث:** يشكل التسمم الحاد نسبة كبيرة من حالات وحدة العناية المركزة وعلى الرغم من أن معدل الوفيات عموماً قد يكون منخفض إلا أن هذه الحالات قد تستهلك معدل كبير من موارد وحدة العناية المركزة. ويعتبر التشخيص المبكر والبدء السريع للعلاج المناسب في قسم الطوارئ ووحدة العناية المركزة هاما جدا لخفض معدلات الاعتلال والوفيات لمرضى التسمم في المستشفيات وكذلك التكلفة.

**الهدف من الدراسة:** صممت هذه الدراسة لتحديد التنبؤ بالنتيجة لمرضى التسمم الحاد بوحدة العناية المركزة التي قد تساعد في تحسين منهج الإدارة و تنظيم العمل بالرعاية.

**طريقة البحث:** تم عمل هذه الدراسة بأثر رجعي لمرضى حالات التسمم التي تم ادخالها وحدة العناية المركزة في مركز علاج التسمم بمستشفيات جامعة عين شمس في الفترة من ٢٠١١/٧/١ إلى ٢٠١٢/١/٣١ . وكان إجمالي عدد المرضى ٣٢١ مريض. وتم تقسيم المرضى الى ٣ مجموعات (الوفيات، شفاء بدون مضاعفات ، و شفاء بمضاعفات). وقد تم جمع البيانات من ملفات المرضى وقاعدة البيانات الالكترونية بالمركز .

**نتائج البحث:** قد كشفت نتائج البحث ان ٢٦٥ (٨٢,٦٪) شفاء مع عدم وجود مضاعفات، ٣٠ مريضا (٩,٤٪) على قيد الحياة ولكن بمضاعفات و ٢٦ (٨٪) من المرضى توفي. وقد بينت هذه الدراسة وجود فروق ذات دلالة إحصائية بين الحالات التي خرجت بدون مضاعفات وحالات المضاعفات والوفيات من حيث أسلوب التسمم، العامل المسبب للتسمم، معدل ضربات القلب، ضغط الدم الانقباضي، معدل التنفس، تغيرات الجلد، مقياس غلاسكو للغيبوية، درجة الحموضة في الدم، غسيل المعدة ، استخدام الفحم المنشط ، استخدام الغسيل الكلوي ، تركيب انبوية حنجرية ، التنفس الصناعي والعلاج بالدوبامين.

**استنتاج وتوصيات البحث:** بناء على ذلك نستنتج ان العامل المسبب للتسمم، معدل ضربات القلب، ضغط الدم الانقباضي، معدل التنفس ، درجة الحموضة في الدم، التنفس الصناعي والعلاج بالدوبامين منبئات هامة للنتيجة لمرضى التسمم الحاد بوحدة العناية المركزة . وتوصى الدراسة بتقييم منبئات النتيجة بصورة دورية ومبكرة لتحديد شدة خطورة التسمم وتحسين فترة العلاج وتقرير طريق الرعاية.

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٢ قسم الطب الشرعي والسموم الإكلينيكية-كلية الطب -جامعة سوهاج - مصر.