Epidemiology and outcomes from severe hypoglycemia in Kuwait: a prospective cohort study

Dalal Al Hasan 1*, Ameen Yaseen 2, Mohammad Al Roudan 2 and Lee Wallis 3

Abstract

Background: The objective of this study was to describe the epidemiology of severe hypoglycaemia in Kuwait, aiming to provide a preliminary background to update the current guidelines and improve patient management.

Method: This was a prospective analysis of severe hypoglycaemia cases retrieved from emergency medical services (EMS) archived data between 1 January and 30 June 2020. The severe hypoglycaemia cases were then sub-grouped based on EMS personal initial management and compared in terms of scene time, transportation rate, complications and outcomes. The primary outcomes were GCS within 10–30 min and normal random blood glucose (RBS) within 10–30 min.

Results: A total of 167 cases met the inclusion criteria. The incidence of severe hypoglycaemia in the national EMS was 11 per 100,000. Intramuscular glucagon was used on scene in 89% of the hypoglycaemic events. Most of the severe hypoglycaemia patients regained normal GCS on scene (76.5%). When we compared the two scene management strategies for severe hypoglycaemia cases, parenteral glucose administration prolonged the on-scene time ($P = .002$) but was associated with more favourable scene outcomes than intramuscular glucagon, with normal GCS within 10–30 min ($P = .05$) and normal RBS within 10–30 min ($P = .006$).

Conclusion: Severe hypoglycaemia is not uncommon during EMS calls. Appropriate management by EMS personnel is fruitful, resulting in favourable scene outcomes and reducing the hospital transportation rate. More research should be invested in improving and structuring the prehospital management of severe hypoglycaemia. One goal is to clarify the superiority of parenteral glucose over intramuscular glucagon in the prehospital setting.

Keywords: Severe hypoglycemia, Emergency medical services, Glasgow coma scale, Random blood sugar, Kuwait
hypoglycaemia prevents life-threatening complications and eliminates unnecessary transports to emergency departments [6].

To the best of our knowledge, there have been a limited number of studies on severe hypoglycaemia in the prehospital setting [7, 8]. The goal of this study is to establish severe hypoglycaemia characteristics, complications and outcomes with the aim to deliver groundwork to revise the current severe hypoglycaemia guidelines.

Methods
Setting
Kuwait has a centralized dispatch centre for all ambulance services. For emergency calls, the universal emergency number (1–1–2) has automatic location identification with a centralized dispatch for police, fire and emergency medical services (EMS). If medical assistance is required, the call is forwarded to an EMS call-taker, who answers the call, reconfirms the address, gives first aid instructions and activates the nearest ambulance. The dispatched ambulance is staffed with two emergency medical technicians (EMTs) or 1 paramedic and 1 EMT. EMTs provide basic life support, and paramedics provide advanced life support, both based on North American resuscitation guidelines. Both EMTs and paramedics are trained in blood glucose measurement and interpretation. The national EMS provides a glucometer kit in all ambulances. For severe hypoglycaemia management, the Kuwait EMS follows different protocols than the NAEMSP recommendations for severe hypoglycaemia. In the Kuwait EMS, ‘For a patient confirmed to be hypoglycaemic and oral administration is contraindicated, the usage of 1 mg intramuscular glucagon by EMT and the usage of intravenous or intraosseous dextrose for paramedics are warranted to reverse severe hypoglycaemia in adults.’ [9] For transport decisions, the local EMS adheres to the National Model of EMS Clinical Guidelines: ‘the patient should not be transported if hypoglycaemia resolves after treatment. Release after treatment is indicated when all of the following conditions are met: random blood sugar (RBS) greater than 4.4 mol/dl (>80 mg/dl), normal Glasgow coma scale (GCS), ability to tolerate oral intake, presence of social support, and no other major compliant such as chest pain, shortness of breath, or neurological deficit.’ [10]

Design
This was a prospective analysis of severe hypoglycaemia cases retrieved from emergency medical services (EMS) archived data between 1 January and 30 June 2020. The severe hypoglycaemia cases were then sub-grouped based on EMS personal initial management and were compared in terms of scene time, transportation rate, complications and outcomes. The primary outcomes were GCS within 10–30 min and normal RBS within 10–30 min.

In this analysis we defined favourable on-scene outcomes with severe hypoglycaemia as: normal GCS after 10–30 min on scene and normal RBS on scene after 10–30 min.

Participants
The study population included all adult (> 18 years old) patients for whom EMS was activated due to severe hypoglycaemia. We defined severe hypoglycaemia as having low blood glucose levels of 3.9 mmol/dl or less (<70 mg/dl) that required assistance from another person to be treated. All patients complaining of loss of consciousness, altered mental status, dizziness, seizures, generalized weakness and cerebrovascular accident mimic symptoms plus low RBS (3.9 mmol/dl or less) were included in the analysis [3].

Data collection/measurement
Patient report forms were the only data source for severe hypoglycaemia cases. Patient report forms were completed on scene by EMS personnel and then stored in EMS audit department archived files. The researcher manually collected patient report forms from the EMS audit department archived data. All data were presented in the patient report form, including patient demographics, clinical presentation, RBS, GCS, management, transportation to the hospital and complications. We defined severe hypoglycaemia complications as head injury, cardiac arrest and airway compromise.

Patient report forms with RBS equal to or less than 3.9 mmol/dl at the initial on-scene assessment and presenting compliance with loss of consciousness, altered mental status, dizziness, seizures, generalized weakness and cerebrovascular accident mimic symptoms were included in the analysis [3].

For management, all patients were analysed for receiving intramuscular glucagon and parenteral glucose administration. Intramuscular glucagon is administered at 1 mg into the vastus lateralis muscle for a diabetic patient who is hypoglycaemic and unable to tolerate oral intake [10]. Parenteral glucose is the administration of intravenous 10% dextrose solution (150 ml) to a diabetic patient who is hypoglycaemic and not able to tolerate oral intake [4].

In relation to patient transportation to the hospital, patients were categorized as transported to the hospital, refused transportation to the hospital, and left on scene. EMS personnel leave a patient on scene if the patient meets the “treat and release” criteria [7].
Recurrent utilization of EMS was reported if the same patient repeatedly activated EMS for severe hypoglycaemia.

Severe hypoglycaemia complications included cardiac arrest, head injury or airway compromise.

In terms of primary outcomes, all cases that had a GCS of 15 at 10–30 min on scene were recorded as having a normal GCS after 10–30 min. All cases that had an RBS of 4.4 mmol/dl or more at 10–30 min on scene were documented as normal RBS after 10–30 min.

During the project, all data were kept in password-locked computer files that only the research investigator could open. No data sharing was allowed outside the context of this project.

Sample size
The sample size was determined using G power software version 3.1 with an effect size = .03, power = .80, and α = .05, and the calculated sample size was approximately 82. The relevant graph is attached in Appendix A. Using convenience sampling, all eligible severely hypoglycaemic patients treated by EMS during the study period were included.

Statistical methods
Statistical analysis was performed using Excel and Statistical Package for Social Sciences (IBM SPSS Version 23, NY, USA). Descriptive statistics such as numbers (N), percentages (%), means and SDs were determined to summarize the patients’ characteristics, dural variation, time on scene, and recurrent utilization of EMS. Severe hypoglycaemia cases were then sub-grouped based on scene management strategies and compared using Fisher’s exact test for dichotomous variables and Student’s t-test for continuous variables. Two-sided tests were applied, and a p-value ≤0.05 was interpreted as statistically significant. Missing data were retained as missing; i.e., they were not imputed or estimated.

Results
Out of the 220 severe hypoglycaemia cases, 167 met the inclusion criteria (Fig. 1); thus, the incidence of severe hypoglycaemia was 11 per 100,000 in the national EMS. Severe hypoglycaemia patients were more likely to be Kuwaiti (63%), middle-aged (51 ± 17), and male (60.5%). The peak hours for severe hypoglycaemia events were 6:00–11:59 (27%) and 12:00–17:59 (31%) (Table 1). Intramuscular glucagon was used on scene in 89% of hypoglycaemic events. Most of the severe hypoglycaemia patients regained normal GCS on scene (76.5%, Table 2).

When we compared the two on-scene management strategies for severe hypoglycaemia cases, only few participants received parenteral glucose, 4%. Incidentally we have observed the parenteral glucose administration prolonged the on-scene time (P = .002). However, it was associated with more favourable scene outcomes than intramuscular glucagon, with normal GCS within 10–30

Fig. 1 Flow chart of study population
min \((P = .05)\) and normal RBS within 10–30 min \((P = .006)\) (Table 3). There was no significant difference between the two sub-groups in terms of intervention complications and hospital transportation rate.

**Discussion**

The study describes severe hypoglycaemia epidemiology in Kuwait, which has not been described before. Our results showed that severe hypoglycaemia resulting in a request for EMS is common in Kuwait EMS emergencies. Eleven per 100,000 EMS calls were for severe hypoglycaemia. These results are lower than those reported in the current literature, 35.2 per 100,000 inhabitants [2]. These results were not predicted, especially with Kuwait’s high diabetes mellitus incidence rates [1] and Al Hasan et al.’s 2020 population-based study results. The author documented hypoglycaemia as the most common medical emergency in Kuwaiti homes [11].

This research also reports an overall favourable on-scene outcome for severe hypoglycaemia. A total of 76.5% of the patients regained normal GCS on scene, and 49.5% of them regained normal RBS on scene. To
the best of our knowledge, this is the first study to report severe hypoglycaemia on-scene outcomes.

The favourable on-scene outcomes resulted in a lower hospital transportation rate of 34%. These rates are better than those in the United States. Fifty-three percent of hypoglycaemia patients in the United States are treated and transported by EMS [6].

Our study also reports higher rates of recurrent EMS utilization for severe hypoglycaemia than recent studies, 10% [7].

We also identified intramuscular glucagon as the most frequently used agent during on-scene management to reverse severe hypoglycaemia in Kuwait. This is different from North American EMS systems, where parenteral glucose is widely used to reverse severe hypoglycaemia [6, 8]. One reason for this discrepancy is that the majority of the local EMS staff are EMTs, and parenteral glucose administration is not within the scope of EMT practice [12].

Our analysis incidentally shows more favourable on-scene outcomes with parenteral glucose in a small group of severe hypoglycaemia patients. These findings add to Kauffman 2018 et al.’s large-scale study results. Their study declared that the use of parenteral glucose was associated with lower hospital transportation rates. However, the study did not directly assess parenteral glucose on-scene outcomes. However, these results conflict with existing evidence on the intramuscular glucagon equivalency to parenteral glucose in restoring normal blood glucose and consciousness levels [13–15]. We recommend larger scale studies to confirm favourable on-scene outcomes with parenteral glucose.

The demographic characteristics of the study population (males, 60.5% mean age, 51 ± 17 years) were similar to those of other studies (males, 56.2% mean age 55 years) [7].

Our analysis illustrates similar locations for hypoglycaemic events in the present literature [8] but different peak hours: 06:00–11:59 and 12:00–17:59. These peak hours are not in line with the predicted morning elevation of blood glucose level, i.e., the “dawn phenomenon” [16, 17]. More research is required in this area.

Collectively, this cohort is the first in the current literature to report overall favourable on-scene outcomes with severe hypoglycaemia: normal GCS after 10–30 min on scene and normal RBS on scene after 10–30 min. It also contributes to identifying the role of parenteral glucose in reducing hospital transportation rates.

This research is subjected to several limitations. It compared severe hypoglycaemia on-scene outcomes between two management groups. Consequently, it did not have a randomized, controlled design. Thus, the possibility that the associations identified were related to other factors linked to both the intervention and the outcome could not be fully eliminated. Another limitation is that

| Variable                      | Intramuscular glucagon group | Parenteral glucose group | P value (CI = 95%) |
|-------------------------------|-------------------------------|--------------------------|--------------------|
| Time on scene                 | 27 ± 13 (75)                 | 43 ± 14 (100)            | .002               |
| Normal GCS with in 10–30 min  | 109 (75)                     | 7 (100)                  | .05                |
| Normal RBS with in 10–30 min  | 68 (47)                      | 7 (100)                  | .006               |
| Severe hypoglycaemia complications | 4 (2.7)                   | 1 (14)                   | .214               |
| Intervention complications    | 2 (1.4)                      | 0 (0)                    | .09                |
| Transportation to hospital    | 48 (33)                      | 2 (28)                   | .96                |

*GCS Glasgow coma scale, RBS Random blood sugar
missing cases were excluded from the analysis, which can introduce reporting bias. An additional limitation is that although the study highlighted that parenteral glucose administration reduced the hospital transportation rate, further research is required to establish the impact of parenteral glucose on hospital transportation rates.

**Conclusion**
Severe hypoglycaemia is not uncommon during EMS calls. Appropriate management by EMS personnel is fruitful, resulting in favourable scene outcomes and reducing the hospital transportation rate. More research should be invested in improving and structuring the prehospital management of severe hypoglycaemia. One goal is to clarify the superiority of parenteral glucose over intramuscular glucagon for EMS.

**Abbreviations**
EMS: Emergency medical services; GCS: Glasgow coma scale; RBS: Random blood sugar; EMT: Emergency medical technician

**Supplementary Information**
The online version contains supplementary material available at https://doi.org/10.1186/s12873-021-00457-9.

**Acknowledgements**
The authors would like to acknowledge and thank Mr. Al Jalahmah Kuwait EMS director and his team for their enormous help and support rendered in the course of gathering the necessary data for the study.

**Authors’ contributions**
Study conception and design: DA. Acquisition of data: DA, AY, MA. Analysis and interpretation of data: DA. Drafting of manuscript: DA. Critical revision: DA, LW. All authors read and approved the final manuscript.

**Funding**
This research did not require funding.

**Availability of data and materials**
The dataset used and analysed are available from the corresponding author on reasonable request.

**Declarations**

**Ethics approval and consent to participate**
The study received IRB approval from the Kuwait Ministry of Health independent ethics committee on 20 December 2019 (No. 1278). No administrative permission were need to acquire the collected data. No informed consent was sought from participants because all data were kept anonymous during this study.

**Consent for publication**
Not applicable.

**Competing interests**
The authors declare that they have no competing interests.

**Author details**
1Department of Applied Medical Sciences, College of Health Sciences, Public Authority of Applied Education and training, State of Kuwait, Kuwait City, Kuwait. 2Audit Department, Emergency Medicals Services, State of Kuwait. 3Emergency Medicine Department, University of Cape Town, Cape Town, South Africa.

**Received: 22 February 2021 Accepted: 14 May 2021**

**References**

1. International diabetes federation. IDF MENA members. In: International diabetes federation; 2020. Avilable from URL: https://idf.org/our-network/regions-members/middle-east-and-north-africa/members/38-kuwait.html2020 > [Accessed on September 5 2020].
2. Ratcliffe-Leeving SH, Mequnnin S, Reichert S, Brown J, Black I, Ryan B. Real-world crude incidence of hypoglycaemia in adults with diabetes: Results of the In Hypo-DM Study, Canada. BMJ Open Access Diabetes Res Care. 2018; 6(1–9).
3. Diabetes.co.uk. Diabetes complication; Sever hypoglycaemia: Diabetes .co.uk; 2020. Available from URL: https://www.diabetes.co.uk/severe-hypoglycaemia.html >

4. Cha S, Yun J, Lim T, et al. Severe Hypoglycaemia and Cardiovascular or All-Cause Mortality in Patients with Type 2 Diabetes. Diabetes Metab J 2016; 40(3):202–10.
5. Panska A, Carter R, Pattan V, et al. Population-based study of severe hypoglycaemia requiring emergency medical service assistance reveals unique findings. J Diabetes Sci Technol. 2012;6(1):65–73. doi:https://doi.org/10.1177/193229681100600109.
6. Kaufmann M, Nelson D, Kaushik P, Clay-Mann N, Mitchell B. Hypoglycemia emergencies: factors associated with prehospital care, transportation status, emergency department disposition, and cost. Prehospital Emergency Care. 2019;23(4):453–64. doi:https://doi.org/10.1080/10903127.2018.1528322.
7. O’Connor L, Kue R, O’Connor M. Characteristics of patients with recurrent emergency medical services utilization for symptomatic hypoglycemia in an urban setting. Prehosp Emerg Care. 2019;23(6):780–78. doi:https://doi.org/10.1177/193229681200600109.
8. Sinclair J, Austin M, Froats M, et al. Characteristics, prehospital management, and outcomes in patients assessed for hypoglycemia: repeat access to prehospital or emergency care. Prehospital Emergency Care. 2019;23(3):364–76. doi:https://doi.org/10.1080/10903127.2018.1504150.
9. Kuwait Emergency Medical Services training department. Emergency medical technician training course. Kuwait Emergency Medical Services. 2015.
10. Woodburn E, Rostyus K. Prehospital Management of Hypoglycemic Emergencies Evidence-Based Review for Collegiate-Based Emergency Medical Services’ the journal of college of emergency medical services. 2019;21(1):9–15.
11. Al hasan D, Monger E, Brightwell R. Medical emergencies requiring first aid at HomeA Population-Based Survey Study. Disaster Med Public Health Prep. 2020:1–7. doi:https://doi.org/10.1017/dmp.2020.193.
12. Kuwait Emergency Medical Services training department. Emergency medical technician job description. Kuwait Emergency Medical Services. 2015.
13. Kedia N. Treatment of severe diabetic hypoglycaemia with glucagon: an underutilized therapeutic approach. Diabetes Metab Syndr Obes. 2011;4: 337–46. doi:https://doi.org/10.2147/OMSO.S20633.
14. Patrick A, Collier a, Hepburn D, Steedman D, Clarke F, Robertson C. comparison of intramuscular glucagon and intravenous dextrose in the treatment of hypoglycaemic coma in an accident and emergency department. Emerg Med. 1990;7:73–7.
15. Khan P, Wagner N, Gabbay R. Underutilization of Glucagon in the Prehospital Setting. Ann Intern Med. 2017;168(8). doi:https://doi.org/10.7326/M17-2222.
16. Porcellati F, Lucidi P, Bolli G, Fanelli C. Thirty years of research on the Dawn phenomenon: lessons to optimize blood glucose control in diabetes. Diabetes Care. 2013;36(12):3860–2. doi:https://doi.org/10.2337/dc13-2088.
17. Holstein A, Patzer O, Machalka K, Holstein J, Stumvoll M, Kovacs P. Substantial increase in incidence of severe hypoglycaemia between 1997–2000 and 2007–2010: a German longitudinal population-based study. Diabetes Care. 2012;35(S1):972–5.

**Publisher’s Note**
Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.