# Assessment of nursing perceptions of three insulin protocols for blood glucose control in critically ill patients

Avaliação da percepção de enfermeiros sobre três protocolos para controle glicêmico em pacientes críticos

Thiago Domingos Corrêa<sup>1</sup>, Francisca Pereira de Almeida<sup>1</sup>, Alexandre Biasi Cavalcanti<sup>1</sup>, Adriano José Pereira<sup>1</sup>, Eliézer Silva<sup>1</sup>

## ABSTRACT

Objective: To evaluate nurses' perception regarding three different blood glucose control protocols for critically ill patients. Methods: As part of a randomized trial comparing three blood glucose control protocols in critically ill patients (Computer-Assisted Insulin Protocol, Leuven Protocol, and conventional treatment), all nurses participating in the study were asked to fill in a questionnaire to assess their perceptions of efficacy, complexity, feasibility, and safety (as to the occurrence of hypoglycemic episodes), an to indicate which protocol they would like to see adopted as the standard one in the Intensive Care Unit they worked. Results: Sixty nurses answered the questionnaires. Computer-Assisted Insulin Protocol was considered the most efficient protocol to maintain blood glucose levels within the target range by 58% of the nurses, compared to 22% for Leuven Protocol (p<0.001) and 40% for conventional treatment (p=0.04). Computer-Assisted Insulin Protocol was considered easier to use than Leuven Protocol (p < 0.001) and as easy as conventional treatment (p = 0.78). Out of the nurses, 37% considered Computer-Assisted Insulin Protocol more feasible than Leuven Protocol and conventional treatment. A total of 51% of nurses chose Leuven Protocol as the protocol more often associated with hypoglycemia, while 27% chose Computer-Assisted Insulin Protocol and 8% conventional treatment. Finally, 56% of the nurses selected Computer-Assisted Insulin Protocol as the protocol they would like to see adopted as the standard one in the Intensive Care Unit they were based, as compared to 22% that selected Leuven Protocol and 15% that selected conventional treatment. Conclusion: Computer-Assisted Insulin Protocol was considered more efficacious, easier to use and safer than Leuven Protocol by nurses. The complexity and feasibility of Computer-Assisted Insulin Protocol were considered similar to conventional treatment. Most nurses chose of Computer-Assisted Insulin Protocol as the protocol they would like to see adopted in their Intensive Care Units.

**Keywords:** Blood glucose/metabolism; Insulin; Hyperglycemia; Hypoglycemia; Metabolism

## **RESUMO**

Objetivo: Avaliar a percepção de enfermeiros a respeito de três protocolos para controle glicêmico em pacientes críticos. Métodos: Como parte complementar de um estudo randomizado comparando três protocolos de controle glicêmico em pacientes críticos (Protocolo de Insulina Assistido por Computador, Protocolo de Leuven e tratamento convencional), todos os enfermeiros participantes do estudo foram convidados a preencher um questionário a fim de avaliar suas percepcões a respeito da eficácia, complexidade, viabilidade e segurança (em razão da ocorrência de episódios de hipoglicemia), e indicar qual dos três protocolos eles gostariam que fosse adotado na Unidade de Terapia Intensiva em que trabalhavam. Resultados: Os questionários foram respondidos por 60 enfermeiros. O Protocolo de Insulina Assistido por Computador foi considerado o protocolo mais eficiente para a manutenção do nível de glicemia dentro da faixa-alvo por 58% dos enfermeiros, comparado a 22% para PL (p<0,001) e 40% para TC (p=0,04). O Protocolo de Insulina Assistido por Computador foi considerado mais fácil de ser utilizado em relação ao Protocolo de Leuven (p<0,001) e tão fácil quanto o tratamento convencional (p=0,78). Dentre os enfermeiros, 37% consideraram o Protocolo de Insulina Assistido por Computador mais viável do que o Protocolo de Leuven e o tratamento convencional. O Protocolo de Leuven foi escolhido por 51% como o mais frequentemente associado à hipoglicemia, enquanto 27% dos enfermeiros elegeram o Protocolo de Insulina Assistido por Computador e 8% o tratamento convencional. Finalmente, 56% dos enfermeiros escolheram o Protocolo de Insulina Assistido por Computador como o protocolo que gostariam que fosse adotado na Unidade de Terapia Intensiva em que trabalhavam

Study carried out at the Intensive Care Unit, Hospital Israelita Albert Einstein – HIAE, São Paulo (SP), Brazil.

<sup>&</sup>lt;sup>1</sup> Intensive Care Unit, Hospital Israelita Albert Einstein – HIAE, São Paulo (SP), Brazil.

Corresponding author: Eliézer Silva – Avenida Albert Einstein, 627 – Morumbi – Zip code: 05651-901 – São Paulo (SP), Brazil – Phone: (55 11) 2151-1500 – Fax: (55 11) 3746-9411 – E-mail: eliezer@einstein.br Received on: Jan 26, 2012 – Accepted on: Jul 26, 2012

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em comparação a 22% para o Protocolo de Leuven e 15% para o tratamento convencional. **Conclusão:** Protocolo de Insulina Assistido por Computador foi considerado pelos enfermeiros mais eficaz, fácil de utilizar e mais seguro do que o Protocolo de Leuven. A complexidade e viabilidade do Protocolo de Insulina Assistido por Computador foram consideradas similares ao tratamento convencional. A maioria dos enfermeiros gostaria que o Protocolo de Insulina Assistido por Computador forse o protocolo adotado na Unidade de Terapia Intensiva em que trabalhavam.

**Descritores:** Glicemia/metabolismo; Insulina; Hiperglicemia; Hipoglicemia; Metabolismo

## INTRODUCTION

Tight blood glucose control with continuous intravenous insulin administration may reduce morbidity and mortality in critically ill patients<sup>(1-3)</sup>. Since the publication of the reference randomized controlled trial of Van den Berghe et al.<sup>(1)</sup>, intensive insulin therapy strategy has been recommended based on guidelines from several medical associations<sup>(4,5)</sup>.

Different results regarding efficiency and safety have been published related to the implementation of various insulin protocols in Intensive Care Units (ICUs)<sup>(1,6-8)</sup>. The best results were found with nursedriven protocols, which quickly led to more effective blood glucose control in critically ill patients, compared to physician-led management<sup>(9,10)</sup>. Therefore, highly involved and motivated ICU nurses are essential for successful implementation of an efficient and safe protocol<sup>(10-12)</sup>. One of the most important obstacles in the implementation of a tight glucose control algorithm is the increased nursing workload imposed by frequent blood glucose determinations and insulin adjustments<sup>(11,12)</sup>. Nursing staff concerns regarding hypoglycemia risk and a lack of knowledge about normal blood glucose maintenance benefits may also hinder protocol acceptance<sup>(12,13)</sup>.

Some studies were performed to evaluate the workload, time, and costs related to algorithms for tight glycemic control. Only a few studies examined the perception of nurses regarding intensive insulin therapy<sup>(12,14-16)</sup>. However, we are not aware of any studies that have compared nurses' views on three different protocols implemented simultaneously as part of a randomized controlled trial.

## **OBJECTIVE**

Our aim was to evaluate nurses' perception of the efficacy, complexity, feasibility, and safety (measured by the occurrence of hypoglycemic episodes), as well as

which protocol they would you like to see adopted as the standard in their ICU, among three different blood glucose control protocols administered to critically ill patients as part of a prospective, randomized, controlled, multicenter trial<sup>(17)</sup>. The three insulin protocols evaluated in the randomized controlled trial were Computer-Assisted Insulin Protocol (CAIP)<sup>(17)</sup>, the Leuven Protocol (LP)<sup>(1,2)</sup>, and conventional treatment (CT).

# **METHODS**

## **Participants**

A randomized controlled trial was conducted to compare three insulin administration protocols in intensive care patients<sup>(17)</sup>. When this study was complete, a questionnaire was given to all nurses to assess their perceptions about efficacy, complexity, feasibility, and safety (measured by the occurrence of hypoglycemic episodes), as well as which protocol they would you like to see adopted as the standard in their ICU (Supplemental Digital Content / Methods).

The study was conducted in five ICUs from different Brazilian organizations: *Hospital Estadual Mário Covas*, a 32-bed closed teaching ICU in a 321-bed hospital; *Hospital Israelita Albert Einstein* (HIAE), a 30-bed open teaching ICU in a 450-bed hospital; *Hospital Municipal São José*, a 8-bed, teaching closed ICU in a 200-bed hospital; *Hospital Dona Helena*, a 7-bed, non-teaching, closed ICU in a 120-bed hospital; and *Centro Hospitalar* UNIMED, an 8-bed, non-teaching, closed ICU in a 140-bed hospital.

Adult medical patients admitted to the ICU were eligible for the study if they had at least one blood glucose measurement  $\geq 150 \text{ mg/dL}$  plus one of the following: 1) mechanical ventilation for an acute process, with an expected duration  $\geq 24$  hours; 2) trauma; 3) burns; and/ or 4) systemic inflammatory response syndrome (SIRS, modified criteria), with at least three of the following: a) a core temperature  $\geq 38^{\circ}$ C or  $\leq 36^{\circ}$ C; b) heart rate of  $\geq 90$  beats per minute, except in patients with a medical condition or who were receiving a medication known to prevent tachycardia; c) respiratory rate  $\geq 20$  breaths per minute or a partial pressure os carbonic dioxide (PaCO<sub>2</sub>)  $\leq 32$ mmHg; and/or d) white blood cell count  $\geq 12,000/\text{mm}^3$  or  $\leq 4,000/\text{mm}^3$ , or >10% immature neutrophils.

Patients were excluded if they were: younger than 21 years old, surgical patients, admitted because of diabetic ketoacidosis or a non-ketotic hyperosmolar state, or in a state in which death was perceived as imminent. The study protocol and consent form were approved by the ethics review board of each institution. The study was performed in accordance with ethical standards stated in the Declaration of Helsinki. Written consent was obtained from every patient or the next of kin when the patient was unable to give it.

#### Insulin protocols (Supplemental Digital Content/Methods)

The three insulin protocols evaluated in the randomized controlled trial were: CAIP, a protocol developed by the authors, based on continuous infusion of insulin with adjustments guided by a hand-held device or desktop software with target glucose levels between 100-130mg/dL<sup>(17)</sup>. LP is the standard strict blood glucose control protocol described in two large clinical trials<sup>(1,2)</sup>. It is a continuous insulin infusion protocol with glucose target levels between 80 and 110mg/ dL. CT involves intermittent subcutaneous insulin administration according to a sliding scale starting with levels above 150mg/dL, with glucose target levels for glucose levels between 80 and 150mg/dL. A computer-generated centralized, blinded allocation sequence, with blocks of six patients and stratified by center was performed. All nurses involved in the study were trained in all protocols before enrollment<sup>(17)</sup>.

## **Data collection**

A waiver for informed consent was obtained from the hospital Institutional Review Board to distribute surveys to critical care nurses.

At the end of the randomized study, we sent the questionnaires to the research coordinator of each of the five centers involved in the study. Then, the questionnaires were delivered by a person not involved in the study to the nurses who were involved in the administration of all three protocols, who filled them out without the presence of the investigators at the nursing station or in the break room, during normal shift hours. To protect the subjects' anonymity, the survey did not ask for any personal identifiers (name, age, ICU, shift etc.), and nurses were instructed to complete the questionnaire without including any identifying information. This questionnaire, developed by the investigators before the randomized controlled trial began, was first tested by six critical care nurses not involved in this study, who provided feedback regarding its validity.

The questionnaire consisted of 10 multiple choice questions (Appendix 1). The first part of the questionnaire (questions A1-A4) asks about the efficacy of the glycemic control: questions A1-A3 allowed the nurses to evaluate the performance of the CAIP, LP, and CT protocols regarding their efficiency in maintaining glucose levels in the target range (% of time that blood glucose levels were under control), and question A4 asked which protocol (CAIP, LP, or CT) was the most efficient for the control of blood glucose levels. The second part (questions B1-B4) evaluated protocol complexity (time spent to execute the protocol tasks) and feasibility: questions B1-B3 asked the nurses to evaluate the complexity of CAIP, LP, and CT, and question B4 asked which protocol (CAIP, LP, or CT) was the most feasible. The third part, question C1, was about safety and asked which protocol had the most episodes of hypoglycemia (blood glucose  $\leq 40 \text{mg/dL}$ ). The final section of the survey (question D1) asked which of the three protocols (CAIP, LP, and CT) the nurses would like to see adopted as the standard protocol in their ICUs.

## **Statistical analysis**

Comparisons were made between CAIP *versus* LP and CAIP *versus* CT. Categorical variables were displayed as absolute and relative frequencies. Nominal variables (with more than two categories) were displayed as proportions with 95% confidence intervals (95%CI). Comparisons of proportions were made using a  $\chi^2$  test or Fisher's exact test, when appropriate. P-values <0.05 were considered statistically significant. The Statistical Package for Social Sciences (SPSS) version 18.0 (SPSS Inc<sup>®;</sup> Chicago, IL, USA) was used for statistical analysis.

## RESULTS

From August to December 2006, 60 nurses from the 5 institutions involved in the study filled out the questionnaires. Out of total of 600 questions, only 2 were left unanswered for an overall response rate of 99.7%.

Nurses' perceptions of the efficacy of protocols are presented in table 1. CAIP was considered efficient for maintaining glucose levels in the target range more than 75% of the time by 35/60 (58%) nurses. A total of 13/60 (22%) and 24/60 (40%) nurses considered LP and CT, respectively, efficient in maintaining glucose levels under control (p<0.001 for CAIP *versus* LP and p=0.04 for CAIP *versus* CT). When considering the protocols' efficiency to maintain glucose levels in the target range more than 90% of the time, 25% (15/60) of the nurses chose CAIP as the most efficient, followed by CT with 13% (8/60) and LP with 5% (3/60) of the nurses (p < 0.001 for CAIP *versus* LP and p=0.16 for CAIP *versus* CT) (Table 1).

 Table 1. How nurses evaluate the performance of Computer-Assisted Insulin

 Protocol, Leuven Protocol, and conventional treatment regarding their efficiency

 in maintaining glucose levels within the target range (% of time that blood glucose

 levels were under control)

Efficiency	Study protocols			
Efficiency (% of the time)	CAIP n (%)	LP n (%)	CT n (%)	
>90*	15 (25.0)	3 (5.0)	8 (13.3)	
About 75**	20 (33.3)	10 (16.7)	16 (26.7)	
About 50	9 (15.0)	13 (21.7)	18 (30.0)	
About 25	8 (13.3)	26 (43.3)	11 (18.3)	
<10	8 (13.3)	8 (13.3)	7 (11.7)	

\* more than 90% of the time: p<0.001 for CAIP versus LP and p=0.16 for CAIP versus CT; \*\* more than 75% of the time: p<0.001 for CAIP versus LP and p=0.04 for CAIP versus CT.

CAIP: Computer-Assisted Insulin Protocol; LP: Leuven Protocol; CT: conventional treatment.

In answering the question, "which protocol do you believe is the most efficient?", 34/59 (58%; 95%CI:44-70) of the nurses chose CAIP as the most efficient, 9/59 (15%; 95%CI: 7-27) chose LP, 8/59 (14%; 95%CI: 6-25) chose CT, and 8/59 (14%; 95%CI: 6-25) believed no difference existed between them.

Nurses' evaluation of the complexity of the protocols is presented on table 2. CAIP was classified as difficult or very difficult to use by 7/60 (11.6%) of nurses, as compared to 23/60 (38.3%) for LP and 8/60 (13.3%) for CT (p<0.001 for CAIP *versus* LP and p=0.78 for CAIP *versus* CT).

Table 2. Nurses' evaluation of the complexity of the protocols

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Complexity	CAIP n (%)	LP n (%)	CT n (%)	
Very easy	34 (56.7)	14 (23.3)	38 (63.4)	
Easy	19 (31.7)	23 (38.3)	14 (23.3)	
Difficult	3 (5.0)	16 (26.7)	6 (10.0)	
Very difficult	4 (6.6)	7 (11.7)	2 (3.3)	

CAIP: Computer-Assisted Insulin Protocol; LP: Leuven Protocol; CT: conventional treatment.

The nurses were asked "which protocol was the most feasible?" and they answered as follows: 22/60 (37%; 95%CI: 24-50) elected CAIP, 13/60 (22%; 95%CI: 12-34) chose LP, 20/60 (33%; 95%CI: 22-47) chose CT, and 5/60 (8%; 95%CI: 3-18) believed no difference existed between them.

Out of the nurses, 51% (30/59; 95%CI: 37-64) chose LP as the one most often associated with hypoglycemia

(blood glucose  $\leq$ 40mg/dL), while 27% (16/59) chose CAIP (95%CI: 16-40), 8% (5/59; 95%CI: 3-19) chose CT, and 14% (8/59; 95%CI: 6-25) believed no difference existed between them.

Finally, the nurses were asked which protocol they wished to be implemented in their ICUs. Out of the nurses, 56% (34/60; 95%CI: 43-69) selected CAIP as the protocol they would like to be adopted in their ICUs, followed by LP with 22% (13/60; 95%CI: 12-34) and CT with 15% (9/60; 95%CI: 7-27). Among nurses involved in this study, 7% (4/60; 95%CI: 2-16) believed no difference existed between them.

### DISCUSSION

In 2001, a large randomized controlled trial by Leuven demonstrated that the normalization of blood glucose levels using an intensive insulin infusion protocol improved clinical outcomes in patients admitted to a surgical ICU<sup>(1)</sup>. Since this publication, intensive insulin therapy strategy has been recommended by guidelines from several medical associations<sup>(4,5)</sup>.

Although nurses agree with the need for glycemic control and believe that such intervention is beneficial for patient care, they also recognize the increased work effort associated with maintaining tight glycemic control<sup>(10)</sup>. Reaching blood glucose goals requires extensive efforts from nurses, including frequent bedside capillary glucose monitoring and the implementation of a variety of insulin infusion protocols, with different degrees of complexity. Moreover, the prevalent fear of hypoglycemia among hospital staff further hinders the widespread acceptance of intensive blood glucose control protocols administered to critically ill patients<sup>(10,11)</sup>.

In the present study, nurses considered CAIP more efficacious in maintaining blood glucose levels within the target range, easier to use, and safer than the LP. In fact, nurses' perceptions corresponded to the randomized controlled trial findings in which the mean of patients' median blood glucose was 125.0 ( $\pm 17.7$ )mg/dL in CAIP and 127.1 ( $\pm 32.2$ )mg/dL in LP (p=0.34). In the CAIP group, 21.4% of patients had at least one episode of hypoglycemia  $\leq 40$ mg/dL, as compared to 41.4% in LP (p=0.02)<sup>(17)</sup>.

When CAIP was compared to CT, a statistically significant difference regarding efficacy in maintain blood glucose levels in the target range for more than 75% of the time was found. Both CAIP and CT were considered equally easy to use and feasible. Finally, the majority of nurses chose CAIP as the protocol they

would like to be adopted as the standard algorithm in their ICUs. We believe that CAIP was considered easier to use than LP because, in CAIP, adjustments are quick and easily completed using the computer software, while, with LP, nurses must consult a written and relatively long protocol<sup>(1,2,17)</sup>.

The reason why LP was considered less safe than the other protocols may be due to the fact that nurses observed a higher incidence of hypoglycemia with this algorithm. However, the fact that, with LP, decisions regarding insulin adjustments are often left to the discretion of the nurses may have contributed to the feeling of less safety.

Blood glucose control in intensive care should be a nurse-driven task<sup>(10-12,15,16)</sup>. Nurse-driven protocols achieve the best results in terms of glucose control and safety<sup>(9-11)</sup>. For this reason, an accurate understanding of the insulin algorithms by nurses is essential. In particular, they should consider whether the algorithm is efficacious in terms of maintaining a controlled blood glucose level, safe and easy to use. We were able to demonstrate that CAIP was the protocol that best met these criteria.

We believe that our results can be generalized for most ICUs. The survey was performed in the setting of a trial involving five mixed ICUs. Eligibility criteria for the trial were very pragmatic and included critically ill hyperglycemic patients, who are ordinarily managed with strict blood glucose control. Finally, the algorithms evaluated are in current use (LP or CT) or may be easily implemented in the ICU routine (CAIP).

Several limitations exist in this study. First, the sample size of nurses is relatively small. Nevertheless, we observed statistically significant differences for most of the comparisons that were carried out. Second, the CAIP protocol was developed by one of the clinical trial authors. However, the study was carried out in five centers, and most nurses were not aware of this fact. Nevertheless, this may have influenced nurses' answers. Although we were not able to estimate the size of this possible bias, we do believe it would not have changed the main conclusions of the study. Third, the difference in the target blood glucose range between CAIP (100-130 mg/dL), LP (80-110 mg/dL) and CT (80-150 mg/dL) must have influenced the nurses' perceptions about the efficacy, complexity, and feasibility of the protocols. We cannot estimate to what extent the target blood glucose range independently influences the nurse's perception.

This study has several strengths. The glucose control protocols were evaluated simultaneously

in the setting of a randomized controlled trial in five different ICUs. This increases the confidence that differences in nurses' evaluations are due to the protocols' properties instead of an imbalance in patient characteristics (e.g., the sickest group of patients being treated with any specific protocol). All nurses involved in the administration of the protocols filled out the questionnaires, with a very low frequency of unanswered questions. Only few studies examined nurses' perceptions as to intensive insulin therapy<sup>(10,12,14-16)</sup>. We are not aware of any study comparing nurses' views of three different protocols implemented simultaneously. Based on our results, it may be interesting to involve the nursing team in an insulin protocol implementation process, and it also may help to support future improvement actions.

# CONCLUSIONS

The CAIP was more efficacious, easier to use, and safer than the LP according to evaluation by nurses. Compared to CT, the feasibility and safety of CAIP were considered similar. Most nurses chose CAIP as the protocol they would like to see adopted as the standard algorithm in their ICU.

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Appendix 1. Questionnaire evaluating nurses' perception of efficacy, complexity, feasibility, safety of three different blood glucose control protocols, as well as which of these they would like to see adopted in their Intensive Care Unit (Translated from Portuguese)

#### Part A: efficacy of glycemic control

Question A1. How do you evaluate the Computer-Assisted Insulin Protocol (CAIP) performance regarding its efficiency to maintain glucose levels in the target range (100-130 mg/dL)?

- $\hfill\square$  Glucose levels were almost all the time under control (>90% of the time).
- Glucose levels were under control most of the time (about 75% of the time).
- Glucose levels were under control half of the time (about 50% of the time).
- Glucose levels were under control only in a minor part of the time (about 25% of all time).
- □ Glucose levels were rarely under control (<10% of the time).

Question A2. How do you evaluate the Leuven Protocol (LP) performance regarding its efficiency to maintain glucose levels in the target range (80-110 mg/dL)?

- □ Glucose levels were almost all the time under control (>90% of the time).
- □ Glucose levels were under control most of the time (about 75% of the time).
- □ Glucose levels were under control half of the time (about 50% of the time).
- Glucose levels were under control only in a minor part of the time (about 25% of all time).
- □ Glucose levels were rarely under control (<10% of the time).

Question A3. How do you evaluate the Conventional treatment (CT) performance regarding its efficiency to maintain glucose levels in the target range (80-150 mg/dl)?

- □ Glucose levels were almost all the time under control (>90% of the time).
- □ Glucose levels were under control most of the time (about 75% of the time).
- □ Glucose levels were under control half of the time (about 50% of the time).
- Glucose levels were under control only in a minor part of the time (about 25% of all time).
- □ Glucose levels were rarely under control (<10% of the time).

Question A4. In your opinion, which protocol was the most efficient to control blood glucose levels?

- Computer-Assisted Insulin Protocol (CAIP).
- Leuven protocol (LP).
- Conventional treatment (CT).
- □ No difference between the protocols.

#### Part B: complexity and feasibility of protocols

Question B1. How do you evaluate the Computer-Assisted Insulin Protocol (CAIP) in relation to its complexity?

- Very easy.
- □ Easy.
- $\hfill\square$  Difficult.
- Very difficult.

#### Appendix 1. Continuation

Question B2. How do you evaluate the Leuven protocol (LP) in relation to its complexity?

- □ Very easy.
- 🗆 Easy.
- Difficult.
- □ Very difficult.

Question B3. How do you evaluate the Conventional treatment (CT) in relation to its complexity?

- □ Very easy.
- 🗆 Easy.
- Difficult.
- □ Very difficult.

Question B4. In your opinion, which protocol was the most feasible?

- Computer-Assisted Insulin Protocol (CAIP).
- Leuven protocol (LP).
- □ Conventional treatment (CT).
- □ No difference between the protocols.

#### Part C: safety

Question C1. In your opinion, more hypoglycemic episodes (blood glucose <40mg/dL) occurred in which protocol?

- Computer-Assisted Insulin Protocol (CAIP).
- □ Leuven protocol (LP).
- □ Conventional treatment (CT).
- $\hfill\square$  No difference between the protocols.

#### Part D: general preferences

Question D1. Among the glycemic control protocols tested in this study, which of them would you like to see adopted as the standard protocol in your institution?

- □ Computer-Assisted Insulin Protocol (CAIP).
- □ Leuven protocol (LP).
- Conventional treatment (CT).
- $\hfill\square$  No difference between the protocols.