CASE REPORT



Unexpected adverse event of insulin therapy in diabetes mellitus

Matteo Borro^{1,2} · Giuseppe Murdaca¹ · Mario Monachesi³ · Simone Negrini¹

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Background

Type 2 diabetes mellitus is a metabolic disorder whose main finding is chronic hyperglycemia due to a mix of peripheral insulin resistance and secretory insulin deficiency. Duration and severity of abnormal glycemia affect patient outcome with cardiovascular complications being responsible for about 50% of mortality. Even microvascular complications, including neuropathy, retinopathy, and nephropathy, present an increased incidence correlated to duration of diabetes. In particular, the prevalence of neuropathy significantly increases and correlates with the progression from normal glucose tolerance (7, 4%) to overt diabetes (28%) [1]. The symmetric sensorimotor polyneuropathy is the most common form of diabetes-related neuropathy caused by metabolic and micro-vessel alterations as a result of chronic hyperglycemia exposure [1]. Treatment of chronic hyperglycemia should be started as soon as possible comprehending insulin therapy (IT) when indicated. Delaying the introduction of IT may impact on chronic complications [2]. Insulin pens have become the most common injection devices, and the 4-mm needle is now recommended for adults and children regardless of age, sex, ethnicity, or BMI. It has demonstrated to be safer, better tolerated, less painful and equivalently effective than the 8-mm needle that showed an higher risk of intramuscular injection [3]. The correct use

Managed by Antonio Secchi.

Matteo Borro borromatteo@libero.it

- ¹ Department of Internal Medicine (DiMI), Clinical Immunology Unit, University of Genoa and Policlinico San Martino, Viale Benedetto XV 6, 16132 Genova, Italy
- ² Department of Internal Medicine, San Paolo Hospital, Via Genova 30, 17100 Savona, Italy
- ³ SSD Endocrinologia E Diabetologia, ASL2 Savonese, Via Manzoni 14, 17100 Savona, Italy

of insulin devices comprehends the proper knowledge of the specific device, the verification of free and unobstructed flow before injection, the adequate cleaning, spacing and rotation between sites of injection, the appropriate time and the exact direction to push the thumb button, such as vertically, and the duration of the pressure on it until the needle is completely withdrawn from the body [3]. Moreover, after injection, needles should be disposed of with appropriate training. In fact, pens present with two different caps: an inner cap that protects the patient from the needle and the external cap that should be recapped before the safe removal of the needle itself. Unfortunately, compliance to devices is suboptimal and adverse events are relatively frequent [4].

Objective

Here, we report a peculiar case report of an adverse events related to insulin pen use in a type 2 diabetes mellitus patient.

Case report

A 68-year-old male presented to the Emergency Department with complaints of pain, swelling and functional limitation of the second and third fingers of the left hand. Medical history included insulin-treated type 2 diabetes mellitus complicated by severe peripheral polyneuropathy and chronic kidney disease. On physical examination, there were signs of cellulitis extending from the metacarpophalangeal joints to the distal phalanges of the second and third left fingers. Patient denied any trauma or injury to the site. Hand X-rays were performed and revealed three different foreign bodies compatible with metal splinter or needles: two were located in the second finger, close to the last interphalangeal joint and in the distal epiphysis of first phalanx respectively, and



Fig. 1 Enlargement of the oblique image of the left hand showing the three foreign bodies (red arrows) and the evident inflammatory edema of second and third fingers (color figure online)

one was detected in the diaphysis of the second phalanx of the third finger (Fig. 1). After a course of systemic nonsteroidal anti-inflammatory drugs and amoxicillin/clavulanic acid 875/125 mg / 8 h, foreign bodies were surgically removed and have been identified as 4-mm insulin pen needles. On questioning, because of its severe sensitivity loss, patient revealed an extreme difficulty in correctly recapping the needle of insulin pen with the external protection before safely discharge the device. He then confirmed that, holding the pen with his left hand, he sometimes broke the needle while recapping the pen. Sensitivity loss due to diabetic polyneuropathy prevented the patient from feeling pain when needles broke into his fingers and only the signs of inflammation revealed their presence. Accordingly, patient was instructed on the proper technique to safely discharge the needle and recap the device after injections.

Discussion

The recommended insulin injection sites are the abdomen, thigh, buttock and upper arm. In order to avoid infections, beyond the chosen site, an appropriate cleaning of the area is necessary before administration [3]. After adequately priming the device, the recommended 4-mm pen needle should be inserted at 90° in the skin so that insulin is safely released into subcutaneous fat tissue and the thumb button should be kept pressed until the patient slowly count to 10 [3]. Most frequent skin-related adverse event associated with IT is lipohypertrophy due to the lack of skin spacing and rotation of injection site [3, 4].

Recommendations suggest that the injection site should be changed at least every four weeks and a correct spacing consists of 1 fingerbreadth (approximately 1 cm) [3]. The risk of intramuscular injection is another challenge, both related to patient characteristic such as skin and subcutaneous thickness and needle length. Intramuscular administration of insulin leads to poor glycemic control and higher risk of hypoglycemia because of the extreme variability in insulin absorption through muscles (especially into exercising muscles) [3]. A further well-known problem is the failure to remove the inner needle cap that makes the patient not taking the insulin dose and, consequently, glycemic control is not achieved [5]. Our clinical case highlights the possibility of other and various adverse events related both to misuse of insulin devices and to the presence of disabling complication. Specifically, our patient was treated with insulin for two years and received a sufficient education on the correct use of the device, but his severe peripheral neuropathy compromised his ability to safely remove the needle. This clinical case underlines that knowledge, skills of patients and their education and counseling in the correct use of insulin pens as well as the regular check of their handling are of crucial importance and strictly recommended to ensure effectiveness and safety of treatments. Moreover, clinicians should be aware that comorbidities and complications of the disease may impact on compliance to devices and their correct use.

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Code availability Not applicable.

Compliance with ethical standards

Conflict of interest The authors declare that they have no conflict of interest.

Human and animal rights The patient was informed about his inclusion in case report and gave his informed consent. The study was performed in accordance with the ethical standards as laid down in the 1964 Declaration of Helsinki and its later amendments or comparable ethical standards. Anyway anonymity is guaranteed in the case report and images do not allow identification.

Informed consent All individual participants gave their informed consent prior to their inclusion in the case report. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/.

References

1. Stolar M (2010) Glycemic control and complications in type 2 diabetes mellitus. Am J Med 123(3 Suppl):S3-11

- American Diabetes A. 9 (2020) Pharmacologic approaches to glycemic treatment: standards of medical care in diabetes-2020. Diabetes Care 43:S98–S110
- 3. Frid AH, Kreugel G, Grassi G et al (2016) New insulin delivery recommendations. Mayo Clin Proc 91(9):1231–1255
- Pozzuoli GM, Laudato M, Barone M, Crisci F, Pozzuoli B (2018) Errors in insulin treatment management and risk of lipohypertrophy. Acta Diabetol 55(1):67–73
- Wei ET, Koh E, Kelly MS, Wright LA, Tylee TS (2020) Patient errors in use of injectable antidiabetic medications: a need for improved clinic-based education. J Am Pharm Assoc (2003) 60(5):e76–e80

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