Granuloma annulare (GA) is a fairly common granulomatous disease characterized by painless, erythematous, annular papules with necrobiosis and granuloma formation in histopathology. There are four clinically distinct subtypes: localized, subcutaneous, generalized and perforating. The first two subtypes primarily occur in childhood, particularly in the first 5–6 years of life [1].

An 8-year-old boy was referred to the paediatric dermatology unit due to multiple unilateral skin lesions associated with a moderate itch. The patient’s parents reported that the first lesion had appeared 5 years before on the patient’s right dorsum of the foot and calf shortly after diabetes mellitus type I diagnosis. A treatment with cryotherapy (1 procedure) and mild topical corticosteroids (3 weeks) was partially effective: lesions improved with treatment but did not resolve completely. Besides the diabetes type I diagnosis (T1DM) (diagnosed according to the internationally accepted criteria [2]), the child was otherwise healthy. On admission, physical examination revealed monomorphic papules in the dorsal area of the right foot and spreading to the calf (Figures 1 A, B). The complete cutaneous inspection did not reveal any other changes on the skin, hair, nail or teeth. Full blood count, routine biochemical analysis and urinalysis were within normal limits, with slightly elevated liver enzymes. Histopathological examination from lesional skin showed...
perivascular lymphocytic infiltrate admixed with numerous histiocytes (Figures 2 A, B).

The clinical presentation of the case described herein is unique in its non-annular morphology and unilateral distribution confined to the right dorsum of the foot and calf. According to the best of our knowledge, this finding of chronic GA located solely on the limb without any additional lesions ever appearing on the torso or any other body part besides the right side of the body has not been reported so far in children. In adults, Borda et al. [3] reported a unilateral GA confined to one leg present for 14 years without generalization to the torso. The patient had concomitant pyoderma gangrenosum and chronic lymphocytic leukaemia [4].

GA has been widely described in adults in association with several systemic diseases, such as viral infections (especially HIV), dyslipidaemia, malignancy (especially hematologic), and thyroid disease [1]. The correlation with diabetes mellitus (DM) in adults is still being discussed [1, 4], however, several cases associating GA with diabetes, especially the onset of the latter disease, in children have been reported [5–8]. Spicuzza et al. [7] presented a rare case of an 8-year-old girl with subcutaneous GA as the first sign of T1DM. The lesions appeared 4 months before the T1DM diagnosis, while Agrawal et al. [9] presented a case of subcutaneous granuloma annulare which appeared nearly 3 months before the T1DM diagnosis. Wong and Verbov [10] reported the case of a 3-year-old boy presenting with subcutaneous GA on the scalp skin area, who subsequently developed T1DM only 2 months after the onset of GA. In that case, the clinical picture worsened with poor glycaemic control or intercurrent illnesses. Grogg and Nascimento [11] reported that two of 34 children (5.9%) with subcutaneous GA were affected by concomitant DM (mean follow-up was 5 years). Kakourou et al. [12] assessed glucose tolerance in a cohort of 15 children with multiple GA and found that these children had higher glycaemic and lower insulimic parameters compared to matched controls, and that 11 of 15 children had a positive family history of DM. It was speculated that prolonged exposure to high glucose levels may contribute to the expression of GA and collagen alterations have been suggested as a primary event in the process of this skin manifestation. In our case, proper control of T1DM did not affect the course of skin lesions that have slowly spread on the skin within 5 years. Dahl in 2007 [13] presented a report with long-term follow-up of Mayo Clinic records of patients with GA. All patients have had follow-up visits of at least 20 years (with the mean follow-up of 35 years). Three out of 32 (9.4%) patients have had or have developed T1DM during the follow-up. On the other hand, Rapelanoro-Rabenja et al. [14] assessed thirty children with a localized variant of GA. No association with diabetes mellitus was found, however, the study’s follow-up period was not specified. A few years later, in 1998, Martínón-Torres et al. [15] retrospectively assessed 42 children with GA with a follow-up of at least 3 years. Again, diabetes and other potentially related disorders were excluded in all cases. In the end, despite a large number of studies published on GA and DM, definitive evidence for the correlation or association has not been found yet. At least in children, a longer follow-up period seems to reveal the possibility of T1DM diagnosis.
Unilateral papular granuloma annulare in a type I diabetic child: a case report and literature review

Treatment options in GA suffer from a lack of large clinical trials. It seems that localized GA may respond to intralesional glucocorticoid, and widespread disease may be initially treated with antimalarial drugs or phototherapy [1]. In our case, 8 weeks of topical mometasone furoate ointment 1 mg/g was effective and a full remission was achieved. At a 6-month follow-up no recurrence was observed.

In conclusion, both our case and the previous literature reports suggest a possible relationship between the altered glucose metabolism and GA presenting within children population. GA in childhood should prompt the assessment of comorbidities. As DM screening is inexpensive, testing and long-term follow-up of all paediatric patients with GA should be considered. Although the morphological presentation of the GA in the described case is not unusual, the unilateral localization in GA has been sporadically reported; that is why we consider this case report to have practical values for everyday clinical practice.

Conflict of interest

The authors declare no conflict of interest.

References