Nexus between glucose variability and healing rate in patients with diabetic foot ulcer

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Introduction
It is difficult to manage a case of non-healing diabetic foot ulcer due to persistent long standing local infection and microvascular defects. Despite the availability of best of the resources in management of diabetes and diabetes foot ulcers, amputations are still performed quite often. To the best of our knowledge there are very few papers assessing the impact of glycemic variability on healing of diabetic foot ulcer.

The author is this case report uses ambulatory glucose monitoring along with intensive blood glucose management for managing 2 complicated cases of non-healing diabetic ulcers.

Case 1
A 50 year old lady, known case of diabetes for 10 years presented to the clinic with non-healing ulcer in the heel of the left foot since 6 months. She had history of tingling and numbness in both feet since 2 years.

History
6 months ago, she had pain and swelling in left foot. Arterial Doppler showed no lesions which were hemodynamically significant. Venous Doppler was negative for deep vein thrombosis. She was diagnosed as left leg cellulitis with diabetic ulcer. She was initiated on premix insulin 40 IU in the morning and 30 IU in the evening along with 2g of Metformin along with oral antibiotics.

4 months ago, she consulted a vascular surgeon. Left heel ulcer debridement was done and vildagliptin was added to the ongoing therapy as her blood glucose levels were uncontrolled (Fasting Blood Sugar – 133 mg/dl, post prandial blood sugar – 241 mg/dl)

3 months ago, her x ray foot showed mild erosion of calcaneum. Her blood glucose levels were grossly uncontrolled (Fasting Blood Sugar – 156 mg/dl, post prandial blood sugar – 286 mg/dl).

The antidiabetics, antibiotics were continued and platelet derived growth factor (PDGF) for local application was initiated.

2 months ago, patient developed high grade fever with chills. Her total leucocyte counts were 21,900 cells/cumm with neutrophilia. Her serum creatinine level was 1.6 mg/dl and Random Blood Sugar (RBS) level was 315 mg/dl. She was managed with IV antibiotics and other supportive measures. Her condition gradually improved after 2 weeks.
On Examination
Ulcer present in left foot (heel) extending to posterior 1/3rd of the lateral border of the foot. Ulcer measured 6cm*3cm. there was minimal slough and minimal mucopurulent discharge. It was grade 2 and 3 as per Wagner’s diabetic ulcer classification. According to University of Texas diabetic wound classification it was Stage B and Grade III. X-RAY Left Foot showed erosions, radiolucent areas in calcaneum suggestive of osteoporosis. Her glycated hemoglobin (HbA1c) level was 11% and RBS was 256 mg/dl. Her hemoglobin (Hb) level was 8.3gm% and total leucocyte count was 8100 cells/cumm. Her renal function tests and liver function tests were within normal limits.
She was diagnosed as non-healing ulcer over left heel with osteomyelitis of left calcaneum bone along with uncontrolled diabetes. Since the patient declined basal bolus therapy as she did not want 4 injections per day, she was initiated on premix50 analogue insulin BD, Tab. Teneligliptin 20 mg OD and Tab. Metformin 1000mg BD. Glucose profile was monitored with using ambulatory glucose monitoring.

AGP during the first 10 days of therapy

Left Foot Ulcer before treatment

X Ray left foot before treatment initiation
Left Foot ulcer after 2 weeks of treatment

After 3 weeks of treatment initiation

4 weeks after treatment initiation

After 7 weeks of therapy
Case 2
A 50 year old gentleman came with left foot swelling since 3 weeks along with blackish discoloration of left 5th toe since 2 weeks and multiple ulcer points with purulent foul smelling discharge since 1 week.

History:
He is a known case of type 2 diabetes since last 20 years. He has history of loss of sensation in both feet since last 5 years.

On examination:
He had swelling and blackish discoloration of left foot and deep black discoloration (TAR) of 5th toe. There were multiple pockets of wound with lots of slough on dorsum aspect of left foot. The 5th flexor tendon was exposed. The wound was not perforating from dorsal to plantar aspect. According to Wagner’s Diabetic Ulcer Classification it was a Grade 4 ulcer. According to the University of Texas Diabetic wound Stage it was a Stage-D and Grade-III ulcer.

FBS was 364 mg/dl, PPBS was 480 mg/dl and A1c was 14.8%. His Hb was 11.4gm%. Total leucocyte count was 7,200cells/cumm. RFT, LFt and serum electrolytes were within normal limits. X ray left foot, electrocardiogram, arterial Doppler and 2D ECHO were within normal limits.

He was diagnosed to have gangrene of left 5th toe with pre-gangrene of left foot and cellulitis of left leg.

The patient was initiated on intensive insulin therapy (continuous subcutaneous insulin infusion) and broad spectrum antibiotics. Patient was put on ambulatory glucose monitoring. He was advised to avoid weight bearing on that foot and advised to come for regular follow ups.
On continuous subcutaneous insulin infusion

On 3 weeks of initiating therapy

On 8 weeks of initiating therapy

On 5 weeks of initiating therapy

On 13 weeks of initiating therapy
Discussion
In both the cases the amputation was advised by the respective consultants prior visiting our center. The blood glucose levels of both the patients were grossly uncontrolled. Strict control of within day and between day using continuous glucose monitoring, trained dressing and intensive insulin therapy resulted in healing of two complicated diabetic foot ulcer. It was cost effective as the patients were managed on out-patient basis.

Conclusion
Ambulatory glucose monitoring along with intensive management of hyperglycemia resulted in healing of non-healing diabetic ulcers. Glycemic variability clearly needs more attention in management of diabetic foot ulcers.

References