CASE SERIES

Treatment Response of Patients with Erythrodermic Psoriasis after Switching to Guselkumab

Cheng-Ying Chiang · Tsen-Fang Tsai

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ABSTRACT

Introduction: Erythrodermic psoriasis (EP) is a rare variant of psoriasis. Due to its rarity, evidence supporting its treatment is limited. Patients with erythrodermic psoriasis often respond less favorably to conventional treatment and have lower biologic drug survival compared with patients with plaque-type psoriasis. Guselkumab is an anti-interleukin-23 (IL-23) monoclonal antibody and has shown excellent and sustained treatment effect in moderate-to-severe plaque-type psoriasis. Until now, there is only one phase III open-label study reporting the effectiveness of guselkumab in this patient group.

Methods: Patients with chronic erythrodermic psoriasis who had been commenced on guselkumab were included in this retrospective study. All except one of our patients had plaque-type psoriasis prior to EP, and they met the criteria for EP before use of guselkumab. Due to inadequate response to previous treatment, including biologics, they were switched to guselkumab treatment. There was no washout period. They received guselkumab 100 mg at weeks 0 and 4, then every 8 weeks until at least week 28. Data were collected for patient demographics, Patient Area and Severity Index (PASI), and previous treatments.

Results: A total of 13 patients (12 male, 1 female) were included. The mean PASI improvement at weeks 4, 12, 20, and 28 was 37.5%, 60.9%, 67.5%, and 64.7%, respectively. The percentage of PASI 75 responders at weeks 4, 12, 20, and 28 was 15.4%, 38.5%, 53.9%, and 46.2%, respectively. At week 12, eight patients (61.5%) could achieve PASI 50 response. These PASI 50 responders at week 12 showed sustained and improved effectiveness during follow-up.

Conclusion: Guselkumab is an effective biologic for treatment of moderate-to-severe plaque-type psoriasis, but the efficacy and drug survival are still compromised in EP patients. Our data describe the real-world experience of guselkumab treatment in this rare group of patients. Despite previous failure of other biologics, PASI response at week 12 is a good predictor of patient response and drug survival of guselkumab.

Keywords: Drug survival; Erythrodermic psoriasis; Guselkumab
Why carry out this study?
Erythrodermic psoriasis (EP) is a rare variant of psoriasis. Due to its rarity, evidence supporting its treatment is limited. To determine the efficacy of guselkumab in EP patients who were previously treated with other biologics or conventional treatment.

What was learned from the study?
The PASI 75 and PASI 90 response rates were lower in our EP case series than in the pivotal phase III studies which included patients with plaque-type psoriasis.

In our patients with EP, biologic naïvety and no history of secondary failure to biologic were good factors predicting guselkumab efficacy.

In our patients with EP, PASI response at week 12 seems to predict sustained drug survival.

DIGITAL FEATURES
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INTRODUCTION
Erythrodermic psoriasis (EP) is a rare and potentially fatal variant of psoriasis. EP was defined as psoriasis involving more than 80% body surface area (BSA). Due to its low prevalence, evidence-based treatment choices are mainly from small case series or case report.
100 mg at weeks 0 and 4, then every 8 weeks until at least week 28. There was no washout period. However, since our patients were switched from prior biologics or conventional systemic treatment, not every patient’s involved skin lesions exceeded 80% BSA when they started guselkumab treatment. PASI response and treatment-emergent adverse events were evaluated by the same dermatologist at each visit. The study was approved by National Taiwan University Hospital Institutional Review Board (201904124RINC) and was performed in accordance with the Declaration of Helsinki of 1964 and its later amendments. Informed consent to participate in the study was obtained from all patients. There is no identifiable information in our paper.

RESULTS

A total of 13 patients [12 male (92.3%), 1 female (7.7%)] were included. Six (46.2%) of the 13 patients had psoriatic arthritis. Mean ± SD age was 50.5 ± 15.3 years. Mean ± SD body weight was 71.0 ± 13.5 kg, and mean ± SD body mass index was 24.6 ± 4.1 kg/m². Mean ± SD baseline Psoriasis Area and Severity Index (PASI) was 23.8 ± 9.7 (Table 1). Mean PASI improvement at weeks 4, 12, 20, and 28 was 37.5%, 60.9%,

Table 1 Summary of 13 cases of erythrodermic psoriasis

| Patient demographics\(a\) (\(n = 13\)) |  |
|------------------------------------------|---|
| Male sex, \(n\) (%)                     | 12 (92.3%) |
| Age (years) ± SD                        | 50.5 ± 15.3 |
| Body weight (kg) ± SD                   | 71.0 ± 13.5 |
| BMI (kg/m²) ± SD                        | 24.6 ± 4.1 |
| History of psoriasis (years) ± SD       | 24.2 ± 13.3 |
| Psoriatic arthritis, \(n\) (%)           | 6 (46.2%) |
| Previous conventional systemic treatment, \(n\) (%) | 13 (100%) |
| Previous biologics use, \(n\) (%)        | 12 (92.3%) |
| Previous inadequate response to biologics, \(n\) (%) | 11 (84.6%) |
| Baseline PASI (absolute PASI before guselkumab) ± SD | 23.8 ± 8.7 |

PASI improvement

| PASI 50 | Week 4 | Week 12 | Week 20\(b\) | Week 28\(b\) |
|---------|--------|---------|-------------|-------------|
|         | 5 (38.5%) | 8 (61.5%) | 8 (61.5%) | 8 (61.5%) |
| PASI 75 | 2 (15.4%) | 5 (38.5%) | 7 (53.8%) | 6 (46.2%) |
| PASI 90 | 0      | 1 (7.7%) | 3 (23.1%) | 4 (30.8%) |
| PASI 100| 0      | 0       | 0          | 2 (15.4%) |

BMI, body mass index; PASI, Psoriasis Area and Severity Index

\(a\) Values are \(n\) (%) or mean ± SD (standard deviation)

\(b\) One patient discontinued guselkumab treatment at week 12 due to unsatisfactory efficacy

\(\triangle\) Adis
67.5%, and 64.7%, respectively. The percentage of PASI 75 response at weeks 4, 12, 20, and 28 was 15.4%, 38.4%, 53.8%, and 46.2%, respectively. The percentage of PASI 90 response at weeks 4, 12, 20, and 28 was 0%, 7.7%, 23.1%, and 30.8%, respectively (Table 1).

The PASI improvement of each patient is shown in Fig. 1a. By week 12, PASI 50 response was seen in eight patients (61.5%) (Fig. 1a, solid line). Most of these patients showed a trend of continuous PASI improvement through week 36, after which the PASI score gradually became stable. One of them had psoriasis aggravation at week 28 (from PASI 89 response at week 20 to PASI 68 response at week 28) but achieved PASI 90 response again at week 36. The other patient had psoriasis aggravation at week 36 (from PASI 76 at week 28 to PASI 63 at week 36).

For PASI 50 nonresponders at week 12 (five patients, 38.5%) (Fig. 1a, dotted line), there was a decreasing trend in PASI improvement through week 28; one of them was prescribed ciclosporine from week 12, and four patients were switched to other biologics (one switched to brodalumab at week 12; the other three switched to adalimumab, brodalumab, and ixekizumab, respectively, at week 28).

No nasopharyngitis, headache, upper respiratory tract infection, or injection-site reaction was reported in our case series through week 36.

DISCUSSION

Patients with EP often have shorter biologic survival period than patients with plaque-type psoriasis. However, two reports have shown excellent and sustained treatment efficacy of guselkumab in EP [7, 8]. In our case series, not all EP patients responded well to guselkumab. The PASI 75 and PASI 90 response rates were lower in our series than in the pivotal phase III studies which included patients with plaque-type psoriasis [6, 7]. Most of our patients also showed a slower response compared with Sano’s series of EP [7]. This lower and slower treatment response may be partly explained by the history of multiple prior biological failure in our patients [3, 9, 10] and unstable disease course due to the lack of a washout period before receiving guselkumab. However, guselkumab still showed better drug survival compared with our prior experience of other biologics in a similar population, despite more prior biologic failures [1, 2].

At week 4, two patients (15.4%) achieved more than PASI 75 improvement, including the only patient who was biologic naïve. The other
early responder was treated with ustekinumab before and had no history of secondary failure. She was switched to guselkumab treatment due to the lack of ustekinumab in our hospital. These two early responders at week 4 achieved PASI 100 at week 28 (Fig. 1a). The other 11 patients (84.6%) were resistant to at least one previous biological agent (etanercept: 5 patients; adalimumab: 6 patients; golimumab: 2 patients; ustekinumab: 8 patients; secukinumab: 9 patients; ixekizumab: 4 patients), and 9 of them (69.2%) were resistant to more than one biological agent. Two, three, one, one, and two patients had failed two, three, four, five, and six prior biologics, respectively.

PASI 50 response at week 12 seems to predict long-term guselkumab drug survival. We divided the EP patients into responders and non-responders according to their PASI response at week 12 (responders: PASI response ≥ 50 at week 12; non-responders: PASI response < 50 at week 12). Responders (eight patients, 61.5%) showed sustained and improved effectiveness during follow-up (Fig. 1b, mean ± SD PASI improvement at weeks 4, 12, 20, 28, and 36 of 46.9 ± 27.1%, 74.7 ± 14.5%, 85.0 ± 12.7%, 85.7 ± 12.8%, and 88.8 ± 12.5%, respectively). Non-responders showed shorter drug survival and quickly deteriorating PASI improvement after their peak response (Fig. 1b, mean ± SD PASI improvement at weeks 4, 12, 20, and 28 of 22.4 ± 31.3%, 37.0 ± 11.5%, 30.5 ± 12.5%, and 19.5 ± 21.7%, respectively).

Human leukocyte antigen (HLA)-Cw6 has been reported to affect the clinical efficacy of guselkumab in patients with moderate-to-severe psoriasis [11]. However, none of our patients had HLA-Cw6 allele. This may partly explain the slower PASI response in our patient series. However, the effect of HLA-Cw6 positivity on the effectiveness of guselkumab for psoriasis in Asian countries remains unidentified.

The limitations of the current study include a small patient population, lack of a control group, and statistical comparison. Despite these limitations, our data still describe an important real-world experience for this rare group of patients. Further studies are needed to identify the best treatment protocol for patients with EP who have failed prior biologics.

CONCLUSION

Guselkumab is an effective biologic for treatment of moderate-to-severe plaque-type psoriasis, but the efficacy and drug survival are still compromised in EP patients. In our patients with EP, biologic naivety and no history of secondary failure to biologic were good factors predicting guselkumab efficacy. For those patients with history of secondary failure to other biologic, guselkumab is still effective in most patients. PASI response at week 12 seems to predict sustained drug survival.

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Compliance with Ethics Guidelines. Compliance with Ethics Guidelines: The study was approved by National Taiwan University Hospital Institutional Review Board (201904124RINC) and was performed in accordance with the Helsinki Declaration of 1964 and its later amendments. Informed consent to participate in the study was obtained from all patients. There was no identifiable information in our paper.

Data Availability. The datasets generated during and/or analysed during the current study are available from the corresponding author on reasonable request.
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