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Thesis Title	Effect of Sitagliptin on chronic plaque psoriasis of diabetic patients in Iraq	
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Abstract	Psoriasis represents a complex chronic systemic T cell immunemediated inflammatory disease characterized by erythematous, scaly plaques of skin and joints. Inflammatory mediators such as IL-17, IL-6, TNF- <i>a</i> and possibly oxidative stress process have stimulated abnormal proliferation of keratinocytes and differentiation resulting in characteristic appearance of psoriasis. Psoriatic patients with plaque psoriasis particularly those with high body mass index have increasing risk of developing a diabetes mellitus type 2 (DM2) and hyperlipidemia. Advance treatment of psoriasis with biological agents such as TNF antagonists and IL-17 antagonist like ustekinumab have been tried with a better result than traditional treatment. However their use have been limited in some patients, because of severe side effects. Sitagliptin is a dipeptidyl peptidase- IV (DPP-IV) inhibitor exerts anti- inflammatory effect when used in patients with type 2 diabetes without any reported severe side effects. Therefore the objective of current study was to assess the effect of Sitagliptin on psoriasis area and severity index (PASI) score in psoriatic patients with diabetes via interfering with metabolic syndrome parameters, inflammatory mediators and oxidative stress markers. Patients and Methods	

The study was conducted on 48 diabetic patients with
moderate to
severe plaque psoriasis who were divided into two groups:
Placebo group
(n=24) patients were administered placebo once daily plus
dietary control
and exercise for 12 weeks ; Sitagliptin group (<i>n</i> =24) patients
were
administered Sitagliptin tablet 100mg once a day plus dietary
control and
XIX
exercise for 12 weeks. PASI score for all patients was assessed
before
and after 12 weeks of treatment. The fasting blood samples
were obtained
from the patients in both groups at baseline and after 12 week
of therapy
used to measure the concentration of serum fasting blood
sugar (FBS),
HbA1c, triglyceride(TG), cholesterol, low density lipoprotein
(LDL),
very low density lipoprotein (VLDL), high density lipoprotein
(HDL),
tumor necrosis factor alpha (TNF-α), intreleukin-17 (IL-17),
intreleukin-6
(IL-6), intreleukin-10 (IL-10), malondialdehyde (MDA) and
reduced
glutathione (GSH), and to determine their correlation with
PASI score
after 12 weeks of treatment. Punch biopsies with size of 5mm
diameter
were performed for both groups at baseline and after 12 week
treatment and sent for histopathological examination and
psoriasis
histopathological score (PHS) was measured for patients in
both groups at
baseline and after 12 week of treatment.
Results: Command with baseling in Site clintin group and placeba
Compared with baseline in Sitagnpun group and placebo
12 week, the level of DASI score was significantly reduced (D <
12 week, the level of rAS1 score was significantly reduced (r < 0.05)
vive) after 12 weeks of Sitaglintin treatment. The level of FRS
HhA1e TC
cholesterol I DI. VI DI TNF-a II -17 II -6 and MDA ware
significantly reduced ($P < 0.05$) after 12 wooks of Situalistin
treatment
when compared with baseline in Sitaglintin group and
nlacebo-treated

group after 12 week and positively correlated ($P < 0.05$) with
PASI score.
In contrast the level of HDL, IL10 and GSH were
significantly increased
(P < 0.05) after 12 weeks of Sitagliptin treatment in
comparison to
baseline in sitagliptin group and with that of placebo-treated group after
$\frac{1}{2}$
12 week and negatively correlated with PASI score ($P < 0.05$). PHS was
significantly reduced ($D < 0.05$) often 12 weeks in comparison
to baseline $(P < 0.05)$ after 12 weeks in comparison
in Sitagliptin-treated group and with that of placebo-treated
group after
XX
12 week. Histopathological examination also revealed a
significant
improvement ($P < 0.05$) in epidermal histological features and
dermal
lymphocytic infiltration with no effect on dermal blood
vessels.
Conclusion
The present study revealed that Sitagliptin reduce PASI score
without
complete absence of psoriatic plaques via an improvement in
hyperglycemia, hyperlipidemia, inflammatory mediators or
cytokines and
oxidative stress markers with confirmation of our clinical and
immunological results by significant improvement in PHS.