

SUPPLEMENTARY DATA

Glucose and insulin concentrations during the hyperinsulinemic-euglycemic clamp studies

There were no significant differences in mean glucose concentration between non-GC users before and after acute prednisolone administration during the steady state period of the low-dose clamp (4.8 ± 0.2 mmol/L vs 4.9 ± 0.1 mmol/L, $p=0.17$) or the high-dose clamp (4.9 ± 0.2 mmol/L vs 4.9 ± 0.2 mmol/L, $p=0.91$) (Supplementary Fig. 1). Glucose concentrations were not significantly different between non-GC users before prednisolone and GC users during the steady state period of the low-dose clamp (4.8 ± 0.2 mmol/L vs 4.8 ± 0.2 mmol/L, $p=0.83$) or the high-dose clamp (4.9 ± 0.2 mmol/L vs 5.0 ± 0.2 mmol/L, $p=0.89$) (Supplementary Fig. 1).

There were no significant differences in mean insulin concentrations between non-GC users before and after acute prednisolone administration during the steady state period of the low-dose clamp (49 ± 12 μ U/mL vs 54 ± 8 μ U/mL, $p=0.051$) or the high-dose clamp (286 ± 46 μ U/mL vs 285 ± 57 μ U/mL, $p=0.77$) (Supplementary Fig. 2). Insulin concentrations were not significantly different between non-GC users before prednisolone and GC users during the steady state period of the low-dose clamp (49 ± 12 μ U/mL vs 51 ± 17 μ U/mL, $p=0.83$) or the high dose clamp (286 ± 46 μ U/mL vs 293 ± 65 μ U/mL, $p=0.83$) (Supplementary Fig. 2).

Glucose enrichment

There was a slight but statistically significant increase in glucose enrichment during the basal phase and low-dose clamp (Supplementary Table 2 and Supplementary Fig. 3). For example, during the basal phase there was a 0.0015% increase in $6,6\text{-}^2\text{H}_2$ glucose enrichment per minute that was statistically significant. There were no group effects ($p \geq 0.16$, ANOVA).

Supplementary Table 1. Non-esterified fatty acids (mmol/L) during each phase of the hyperinsulinemic-euglycemic clamp studies. Data are mean \pm standard deviation, p -value > 0.05 for all between group comparisons.

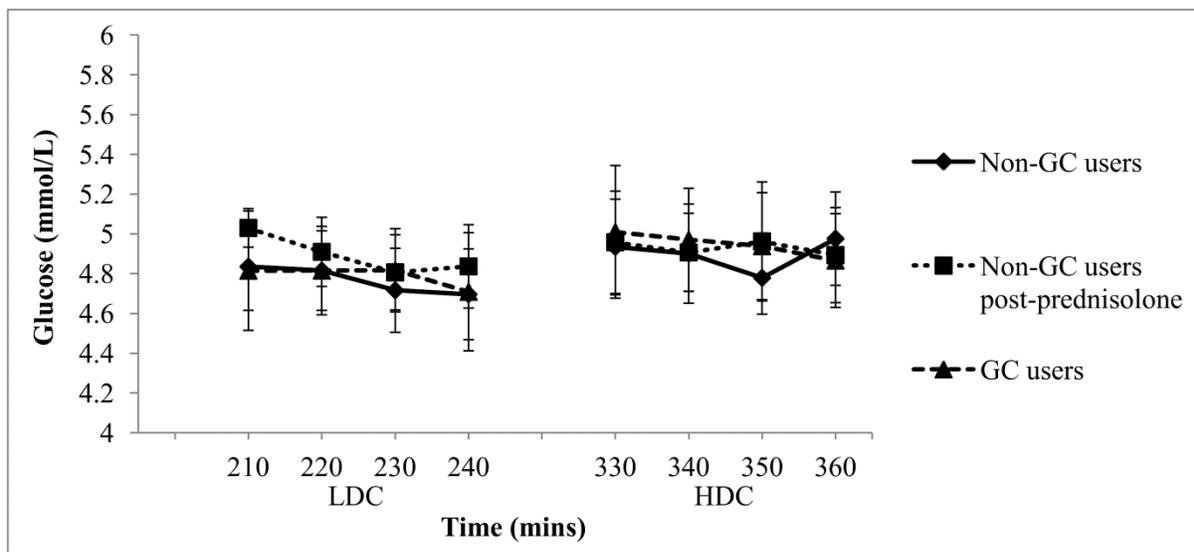
	Non-GC users	Non-GC users post-prednisolone	GC users
Basal	0.42 \pm 0.16	0.36 \pm 0.12	0.36 \pm 0.16
Low-dose clamp	0.05 \pm 0.01	0.06 \pm 0.02	0.06 \pm 0.05
High-dose clamp	0.03 \pm 0.01	0.02 \pm 0.01	0.03 \pm 0.01

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Supplementary Table 2. 6,6-²H₂ glucose enrichment slopes during the steady state period of each phase of the hyperinsulinemic-euglycemic clamp studies in all subjects. Slope represents percent change in 6,6-²H₂ glucose enrichment per minute. P-value is derived from t-test versus zero.

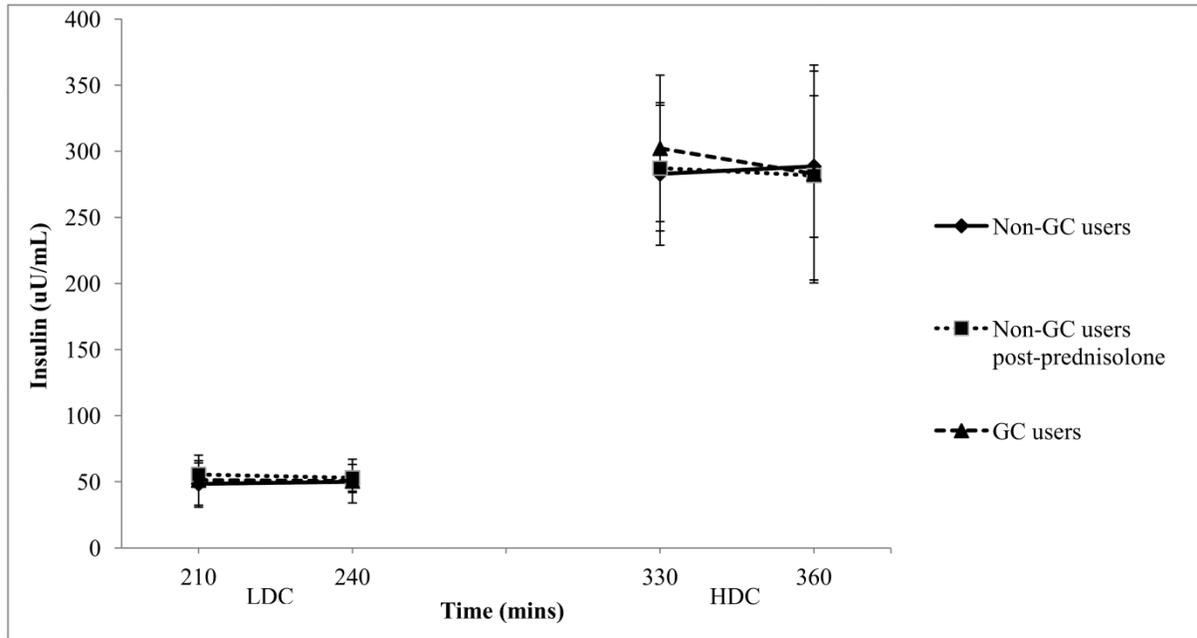
	Slope	SD	p-value
Basal	0.0015	0.0011	0.00000009
Low-dose clamp	0.0005	0.0008	0.0018
High-dose clamp	0.0001	0.0008	0.5280

Supplementary Figure 1. Glucose concentration during steady state period of low-dose and high-dose clamp studies in non-glucocorticoid (GC) users pre- and post- administration of low-dose prednisolone for 7-10 days, and in chronic GC users. Time reflects number of minutes after commencement of estimation of endogenous glucose production. Data are mean ± SD. LDC = low-dose clamp, HDC = high-dose clamp. *p*-value > 0.05 for mean glucose concentration in each steady state period between all groups.



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Supplementary Figure 2. Insulin concentration during steady state period of low-dose and high-dose clamp studies in non-glucocorticoid (GC) users pre- and post- administration of low-dose prednisolone for 7-10 days, and in chronic GC users. Time reflects number of minutes after commencement of estimation of endogenous glucose production. Data are mean \pm SD. LDC = low-dose clamp, HDC = high-dose clamp. p -value > 0.05 for mean insulin concentration in each steady state period between all groups.



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Supplementary Figure 3. 6,6-²H₂ glucose enrichment during steady state period of estimation of endogenous glucose production (EGP), low-dose and high-dose clamp studies in non-glucocorticoid (GC) users pre- and post- administration of low-dose prednisolone for 7-10 days, and in chronic GC users. Time reflects number of minutes after commencement of estimation of endogenous glucose production. Data are mean ± SD. B = basal EGP, LDC = low-dose clamp, HDC = high-dose clamp. *P*-values are as per Supplementary Table 2.

