

# LONG-TERM EFFECTS OF GH REPLACEMENT THERAPY ON HEMATOPOIESIS IN GH DEFICIENT CHILDREN

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## BACKGROUND

Among their metabolic effects, GH and its mediator IGF-1 have been reported to influence hematopoiesis. Indeed, GH/IGF-1 axis promotes erythropoiesis and GH deficiency (GHD) has been associated with a normochromic and normocytic anemia both in adults and in children.

In contrast, in vivo data on the effects of GH/IGF-1 axis on leukocytes and platelets are scanty.

## OBJECTIVE

To evaluate the effects of four-years GH replacement therapy (GHRT) on hematopoiesis in GHD children.

## METHODS

One hundred GHD children (64M) aged  $9.74 \pm 3.95$  years were enrolled in the study. Anthropometric measures, serum IGF-1 levels and blood count were evaluated at baseline and then annually during the first 4 years of GHRT. One hundred healthy children sex- and age-comparable to the patients were enrolled as controls and evaluated annually.

## RESULTS

Clinical features of GHD patients and controls at baseline are reported in Table.

At the start of the study GHD children showed levels of hemoglobin (Hb) ( $12.5 \pm 1.1$ g/dl), hematocrit (Hct) ( $36.7 \pm 4.0\%$ ) and red cells number (RC) ( $4.62 \pm 0.41 \times 10^6$ /mcl) lower than controls (Hb  $13.0 \pm 1.0$ g/dl,  $p < 0.002$ ; Hct  $38.1 \pm 4.3\%$ ,  $p < 0.02$ ; RC  $4.74 \pm 0.34 \times 10^6$ /mc,  $p < 0.01$ ).

Four years of GHRT were associated with a significant increase in Hb ( $13.2 \pm 1.0$ g/dL,  $p < 0.0001$ ), Hct ( $39.0 \pm 3.4\%$ ,  $p < 0.0001$ ) and RC ( $4.75 \pm 0.41 \times 10^6$ /mcl,  $p < 0.05$ ) which became comparable to controls (Hb  $13.3 \pm 1.3$ g/dl; Hct  $39.7 \pm 7.8\%$ ; RC  $4.78 \pm 0.37 \times 10^6$ /mcl).

Hb levels significantly correlated with IGF-1 serum levels ( $r = 0.32$ ,  $p < 0.0001$ ).

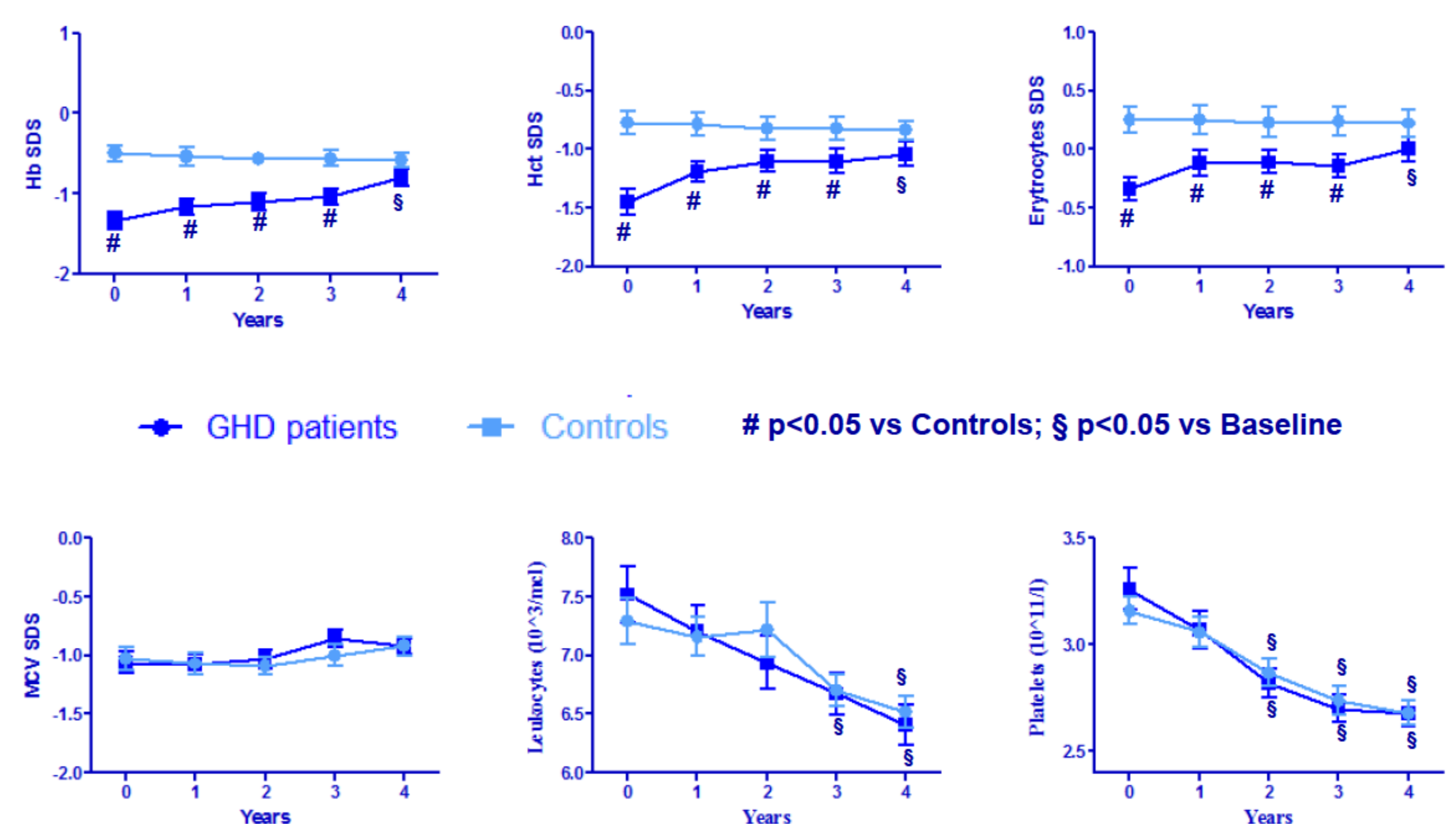
Red cell parameters expressed in SDS in patients and controls throughout the study are showed in Figure.

At baseline seventeen GHD children (17%) showed a normochromic, normocytic anemia while after 4 years of GHRT only 2 patients (2%) were still anemic.

No difference in leukocytes and platelets count was detected between patients and controls neither at baseline nor during the study. A physiological reduction in leukocytes and platelets numbers was observed both in patients and in controls (Figure).

Table – Clinical features of GHD children and controls at baseline

	GHD children	Controls	p
Male /Female	64/36	64/36	NS
Age (years)	$9.73 \pm 3.95$	$9.66 \pm 4.45$	NS
Partial/Severe GHD	68/32	-	
Isolated/Multiple GHD	85/15	-	
Height (SDS)	$-2.32 \pm 0.88$	$-1.00 \pm 1.30$	$< 0.0001$
BMI (SDS)	$-0.41 \pm 1.23$	$-0.11 \pm 1.24$	NS
IGF-1 (SDS)	$-1.91 \pm 0.93$	$0.14 \pm 1.00$	$< 0.0001$



## CONCLUSION

GHD in children is associated with a significant reduction in Hb, Hct and RC. Long-term GHRT improves these anomalies.

Neither GHD nor GHRT have effects on leukocytes and platelets numbers.

