# The correlation between the increase in insulin-like growth factor-I 24 hours after the first injection of growth hormone and the improved growth

Kenichi Miyako, Yuichi Mushimoto, Atsuko Kawano Department of Endocrinology and Metabolism, Fukuoka Children's Hospital, Fukuoka, Japan

Disclosure: Nothing to Disclose.

#### Introduction

Insulin-like growth factor-I (IGF-I) is a biomarker of growth hormone (GH), and is often used to titrate the dose of GH therapy. However, IGF-I production is regulated by not only GH but also other factors. We investigated whether the increase in IGF-I at several time points after the commencement of GH therapy could be a predictive factor for the improved growth.

## Subjects and Methods

We studied 45 pre-pubertal patients with growth hormone deficiency (GHD) that had continued with GH treatment for more than 2 years (29 boys and 16 girls). GHD was mild in 20 patients, moderate in 18, and severe in seven.

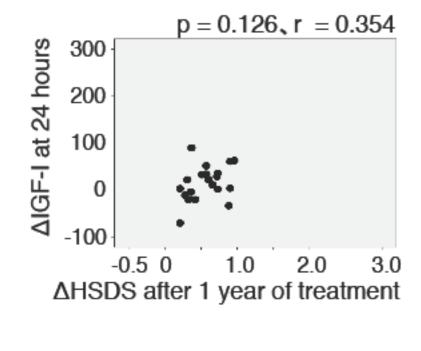
We statistically analyzed the correlation between the following values:

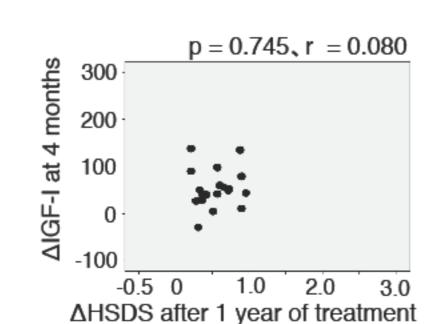
- The increase in IGF-I (ΔIGF-I) at 24 hours, 4 months,
   year, or 2 years after GH therapy was started.
- 2)The increase in height SD score (ΔHSDS) at 1 year or 2 years.

Table 1. Patients characteristics (n=45)

GHD	Mild	Moderate	Severe
	(n=20)	(n=18)	(n=7)
Age at the	$7.4 \pm 1.5$	$7.3 \pm 1.3$	$5.8 \pm 2.0$
commencement	(5.5 <b>~</b> 11.3)	(5.5~9.8)	(3.1~8.1)
of GH therapy			
HSDS at the	$-2.9 \pm 0.6$	$-3.3 \pm 1.3$	$-3.2 \pm 0.9$
commencement	(-4.2~-2.1)	(-7.7 <b>~</b> -2.4)	(-4.3 <b>~</b> -2.2)
of GH therapy			

Fig 1A. Mild GHD (ΔHSDS after 1 year of treatment)





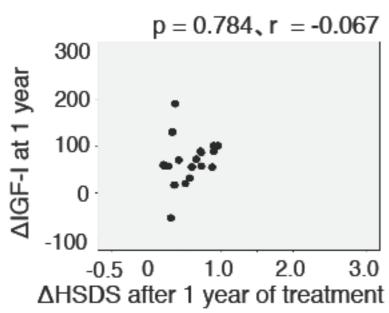
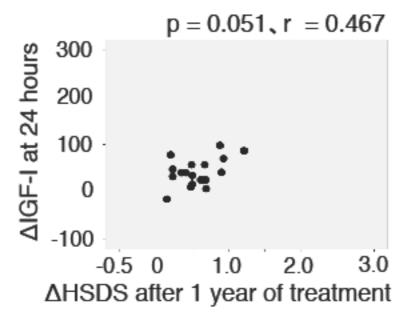
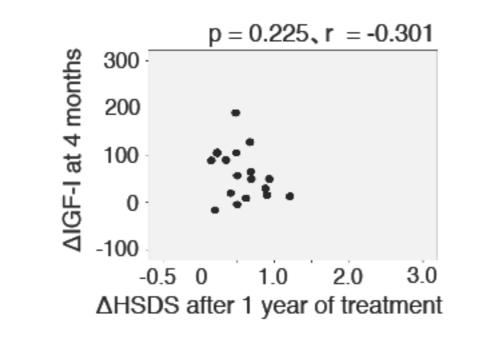


Fig 1B. Moderate GHD (ΔHSDS after 1 year of treatment)





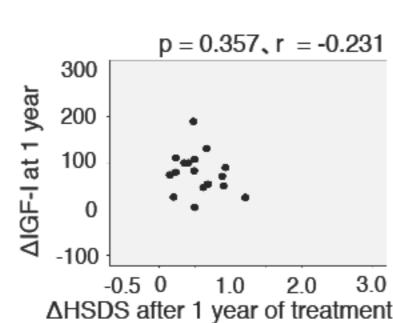
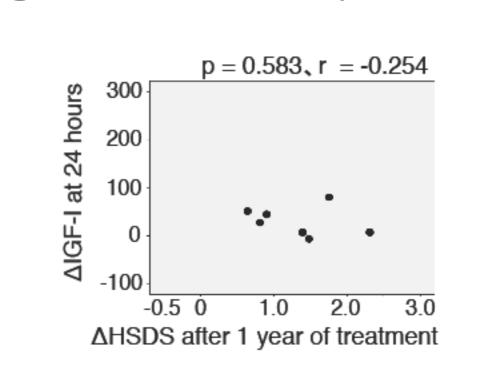
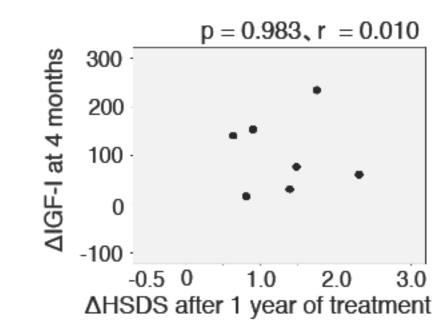


Fig 1C. Severe GHD (ΔHSDS after 1 year of treatment)





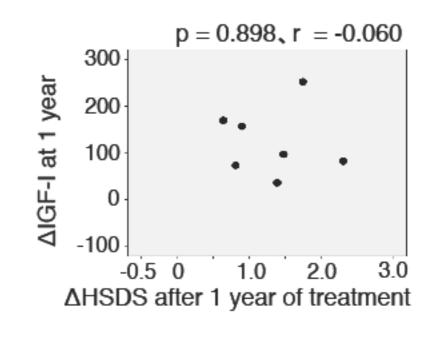
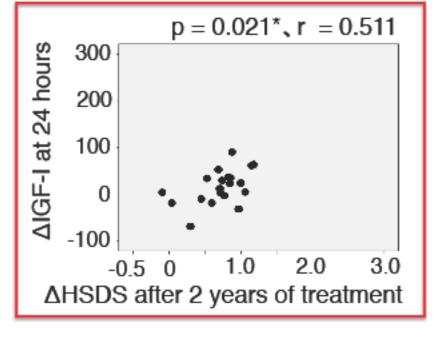
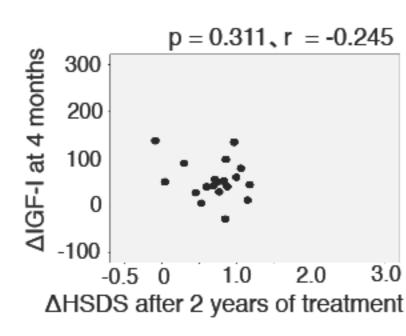
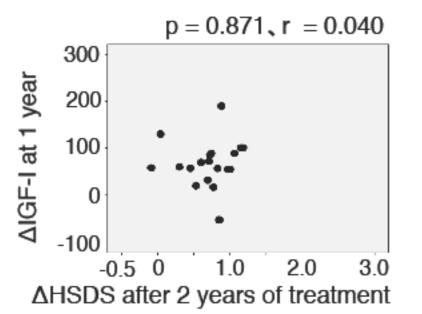


Fig 2A. Mild GHD (ΔHSDS after 2 years of treatment)







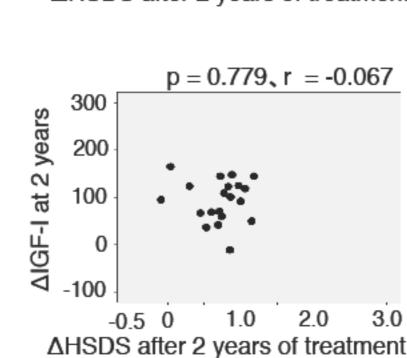
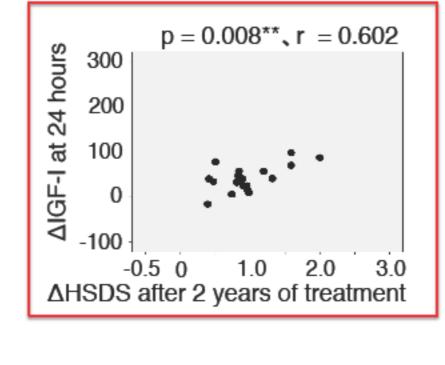
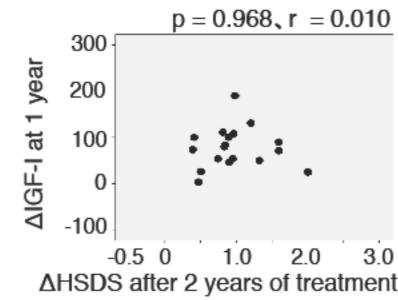
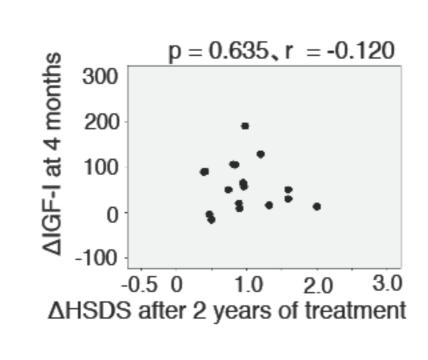


Fig 2B. Moderate GHD (ΔHSDS after 2 years of treatment)







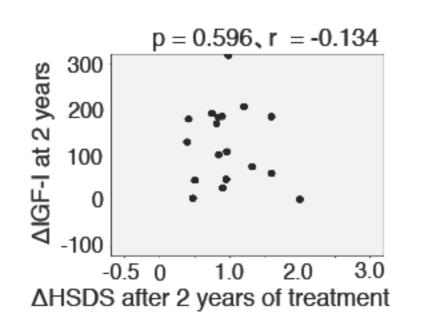
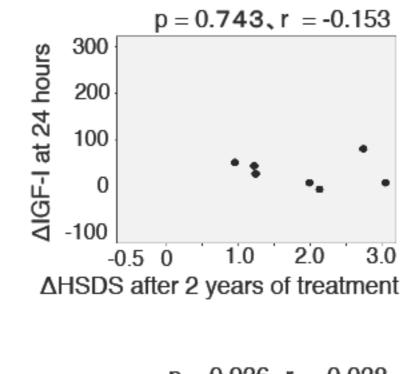
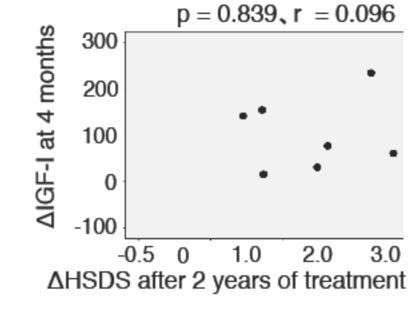
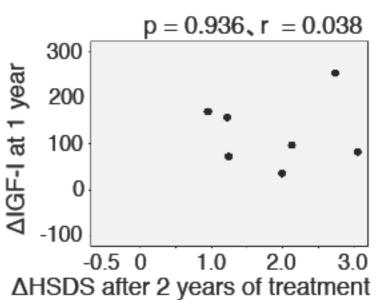
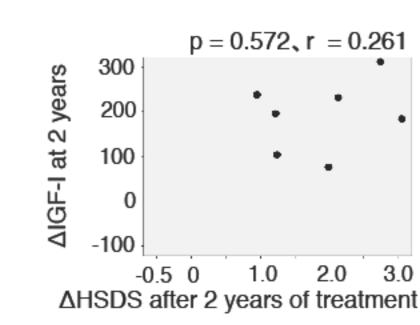


Fig 2C. Severe GHD ( $\Delta$ HSDS after 2 years of treatment)









### Results

The  $\Delta$ IGF-I 24 hours after GH therapy was started was significantly correlated with  $\Delta$ HSDS at 2 years in mild and moderate GHD patients (Figs 2A and 2B: p=0.021, r=0.511 and p=0.008, r=0.602, respectively). The  $\Delta$ IGF-I at 4 months, 1 year, or 2 years in the mild and moderate GHD patients and at any time points in the severe GHD patients was not correlated with  $\Delta$ HSDS at 2 years. When we analyzed both mild and moderate GHD patients together, the 24 hours  $\Delta$ IGF-I was significantly correlated with  $\Delta$ HSDS at 1 year as well (p=0.016, r = 0.389, data not shown).

### Discussion

This study clarified that the  $\Delta$ IGF-I 24 hours after the initiation of GH therapy was significantly correlated with the improved growth of GHD patients.

Since IGF-I is affected by various factors, i.e. nutrition and pubertal stage, its value immediately after the initiation of GH therapy is important to reflect the efficacy of GH.

### Conclusion

IGF-1 24 hours after the first injection of GH is an early and useful predictive factor for the efficacy of GH in GHD patients.

### References

- Pinchas Cohen, et al. Dose-sparing and safety-enhancing effects of an IGF-I-based dosing regimen in short children treated with growth hormone
  in a 2-year randomized controlled trial: therapeutic and pharmacoeconomic considerations. Clinical Endocrinology 81, 71–76, 2014.
- Toshiaki Tanaka, et al. Efficacy and Safety of Up to 8 Years of Long-term Growth Hormone Treatment in Short Children Born Small for Gestational Age in Japan: Analysis of the Subpopulation According to the Japanese Guideline. Clin Pediatr Endocrinol 21, 57-68, 2012.







