

**Table S5.** Effect of rhIGF-1 therapy on BMI SDS (Registry population)

|                            | Change from baseline                      |                                 | Change from baseline                           |                   | P value <sup>a</sup> |
|----------------------------|---|---------------------------------|--|-------------------|----------------------|
|                            | N   | Mean (95% CI)                   | N  | Mean (95% CI)     |                      |
| <b>NPP-LS<br/>(n = 21)</b> |   | <b>NPP-non-LS<br/>(N = 114)</b> |  |                   |                      |
| 1 year                     | 10  | 0.44 (0; 1.23)                  | 77   | 0.11 (0; 0.24)    | 0.744                |
| 2 years                    | 10  | 0.55 (0; 1.53)                  | 68   | 0.21 (0.08; 0.65) | 0.828                |
| 3 years                    | 9   | 0.78 (0; 1.89)                  | 56   | 0.29 (0.11; 0.46) | 0.258                |
| 4 years                    | 8   | 0.99 (0; 2.16)                  | 42   | 0.20 (0; 0.42)    | 0.142                |
| 5 years                    | 8   | 1.16 (0.19; 2.14)               | 25   | 0.30 (0; 0.66)    | 0.062                |
|                            |   |                                 |  |                   |                      |
|                            | <b>NPP-non-LS–responders<br/>(n = 50)</b> |                                 | <b>NPP-non-LS–poor-responders<br/>(n = 38)</b> |                   | P value <sup>b</sup> |
| 1 year                     | 44  | 0.10 (0; 0.31)                  | 33   | 0.12 (0; 0.27)    | 0.930                |
| 2 years                    | 35  | 0.21 (0.01; 0.41)               | 25   | 0.19 (0.01; 0.36) | 0.862                |
| 3 years                    | 30  | 0.21 (0; 0.47)                  | 20   | 0.34 (0.12; 0.56) | 0.467                |
| 4 years                    | 23  | 0.11 (0; 0.44)                  | 11   | 0.36 (0; 0.77)    | 0.357                |
| 5 years                    | 17  | 0.38 (0; 0.88)                  | 5  | 0.29 (0.06; 0.52) | 0.875                |
|                            |   |                                 |  |                   |                      |
|                            | <b>Non-NPP-LS<br/>(n = 17)</b>            |                                 | <b>Non-NPP-non-LS<br/>(n = 86)</b>             |                   |                      |
| 1 year                     | 14  | 0.01 (0.01; 0.32)               | 56   | 0.35 (0.19; 0.52) | -                    |
| 2 years                    | 13  | -0.00 (0; 0.48)                 | 37   | 0.45 (0.23; 0.66) | -                    |
| 3 years                    | 9   | 0.06 (0; 0.49)                  | 24   | 0.51 (0.26; 0.76) | -                    |
| 4 years                    | 8   | 0.35 (0; 1.27)                  | 15   | 0.56(0.25; 0.88)  | -                    |
| 5 years                    | 4   | 0.17 (0; 1.27)                  | 10   | 0.70 (0.08; 1.32) | -                    |

<sup>a</sup>NPP-non-Laron syndrome versus NPP-Laron syndrome mean BMI SDS. <sup>b</sup>NPP-non-Laron syndrome responders versus NPP-non-Laron syndrome-poor responders mean BMI SDS. Responders were defined as patients with change in height SDS in year 1 of  $\geq 0.3$ . Poor-responders were defined as patients with change in height SDS in year 1 of  $< 0.3$ . CI, confidence interval; N, number of patients with available data at each time point. LS, Laron syndrome; non-NPP, not treatment naïve and/or pubertal; NPP, treatment-naïve and prepubertal; SDS, standard deviation score.