

Supplementary Table 1. Histological characteristics of children with biopsy-confirmed NAFLD.

| n=106 | |
|---------------------------------------|-------------------|
| <i>Steatosis grade</i> | |
| 0 | 0 (%) |
| 1 | 26 (24.5%) |
| 2 | 47 (44.3%) |
| 3 | 33 (31.2%) |
| <i>Lobular inflammation</i> | |
| 0 | 11 (10.4%) |
| 1 | 72 (67.9%) |
| 2 | 23 (21.7%) |
| <i>Portal inflammation</i> | |
| 0 | 8 (7.5%) |
| 1 | 76 (71.7%) |
| 2 | 22 (20.8%) |
| <i>Ballooning degeneration</i> | |
| 0 | 17 (16%) |
| 1 | 44 (41.5%) |
| 2 | 45 (42.5%) |
| <i>Fibrosis stage</i> | |
| 0 | 16 (15.2%) |
| 1 | 69 (65%) |
| 2 | 21 (19.8%) |
| 3 | 0 (0%) |
| Definite NASH | 68 (64.1%) |

Supplementary Table 2. Clinical and biochemical characteristics of children with biopsy-proven NAFLD, stratified by tertiles of eGFR_{Bedside Schwartz}.

| Parameters | 1 st tertile (n=36) | 2 nd tertile (n=35) | 3 rd tertile (n= 35) | p value |
|--|-----------------------------------|-----------------------------------|------------------------------------|---------|
| Age (years) | 12.6 ± 3.0 | 11.3 ± 3.0 | 11.5 ± 2.4 | 0.114 |
| Male sex (%) | 47.2 | 65.7 | 51.4 | 0.271 |
| Weight (kg) | 67.5 ± 24.3 | 60.5 ± 19.4 | 64.2 ± 19.9 | 0.382 |
| BMI (kg/m ²) | 28.4 ± 5.2 | 26.1 ± 4.0 | 27.2 ± 5.7 | 0.161 |
| WC (cm) | 88.5 ± 13.3 | 83.8 ± 13.6 | 83.2 ± 15.1 | 0.219 |
| Systolic blood pressure (mmHg), | 116 ± 12 | 108 ± 14 | 114 ± 16 | 0.101 |
| Diastolic blood pressure (mmHg), | 60 ± 10 | 62 ± 11 | 63 ± 11 | 0.662 |
| Total cholesterol (mg/dL) | 161 ± 25.4 | 152 ± 25.9 | 150 ± 33.4 | 0.213 |
| LDL-cholesterol (mg/dL) | 104 ± 22.3 | 99 ± 26.3 | 93 ± 30.0 | 0.282 |
| HDL-cholesterol (mg/dL) | 44 ± 10.8 | 44 ± 7.9 | 44 ± 9.8 | 0.997 |
| Triglycerides (mg/dL) | 99 (64-172) | 88 (64-110) | 70 (52-129) | 0.219 |
| Fasting glucose (mg/dL) | 83.0 ± 8.9 | 82.8 ± 9.5 | 85.2 ± 15.9 | 0.652 |
| Fasting insulin (mIU/L) | 15.9 (11.0-26.9) | 15.0 (9.6-21.3) | 17.0 (10.0-22.2) | 0.608 |
| HOMA-IR score | 3.5 (2.2-5.9) | 3.3 (1.9-4.9) | 3.7 (2.6-5.3) | 0.628 |
| hs-CRP (mg/L) | 0.11 (0.05-0.25) | 0.09 (0.05-0.31) | 0.05 (0.05-0.31) | 0.709 |
| Blood urea nitrogen (mg/dL) | 14.6 ± 3.3 | 14.8 ± 2.8 | 13.6 ± 2.7 | 0.203 |
| Creatinine (mg/dL) | 0.67 ± 0.11 | 0.55 ± 0.07 | 0.46 ± 0.06 | <0.001 |
| eGFR _{Bedside Schwartz} (mL/min/1.73 m ²) | 94 ± 7 | 114 ± 5 | 139 ± 15 | ND |
| Proteinuria (mg/24h) | 50 (33-70) | 48 (38-56) | 50 (37-66) | 0.797 |
| Albuminuria (mg/24h) | 4.7 (2.7-10.3) | 4.1 (2.7-6.8) | 4.2 (3.3-8.9) | 0.723 |
| AST (IU/L) | 26 (24-32) | 30 (24-35) | 27 (23-32) | 0.567 |
| ALT (IU/L) | 26 (20-34) | 32 (21-48) | 25 (19-49) | 0.750 |
| GGT (IU/L) | 15 (11-19) | 16 (14-23) | 14 (11-20) | 0.354 |
| PIIINP (ng/mL) | 8.30 (5.4-12.0) | 5.51 (3.8-8.8) | 4.61 (2.2-7.4) | 0.003 |
| HA (ng/mL) | 73.8 (38.3-179.0) | 37.3 (28.4-76.0) | 28.0 (21.0-36.4) | <0.001 |
| Definite NASH* (%) | 41.7 | 45.7 | 48.6 | 0.857 |
| PNPLA3 rs738409 | | | | 0.030 |
| CC genotype (%) | 11.1 | 40.0 | 37.1 | |
| CG genotype (%) | 44.4 | 28.6 | 40.0 | |
| GG genotype (%) | 44.4 | 31.4 | 22.9 | |

Data are expressed as means ± SD, medians and interquartile ranges (in parenthesis), or percentages.

*Definite NASH was defined as presence of NAFLD Activity Score [NAS] ≥5 on histology.

Abbreviations: BMI, body mass index; WC, waist circumference; LDL, low-density lipoprotein; HDL, high-density lipoprotein; HOMA-IR, homeostasis model assessment-insulin resistance; hs-CRP, high-sensitivity C-reactive protein; eGFR, glomerular filtration rate; AST, aspartate aminotransferase; ALT, alanine aminotransferase; GGT, gamma-glutamyltransferase; PIIINP, N-terminal propeptide of type III procollagen; HA, hyaluronic acid; NASH, non-alcoholic steatohepatitis; ND, not determined.

Supplementary Table 3. Mann-Whitney U-test comparisons between HA, PIIINP, and PNPLA3 genotype and histological features.

| | Steatosis | | Lobular inflammation | | Portal inflammation | | Ballooning | | Fibrosis | | NAS | |
|--------------------------|-----------|----------|----------------------|----------|---------------------|----------|------------|----------|----------|----------|----------|----------|
| | <i>z</i> | <i>p</i> | <i>z</i> | <i>p</i> | <i>z</i> | <i>p</i> | <i>z</i> | <i>p</i> | <i>z</i> | <i>p</i> | <i>z</i> | <i>p</i> |
| HA | 2.88 | 0.01 | 4.70 | 0.01 | 5.5 | 0.012 | 1.91 | 0.11 | 3.11 | 0.02 | -2.78 | 0.002 |
| PIIINP | -5.51 | 0.001 | -2.82 | 0.001 | 5.8 | 0.001 | -1.79 | 0.08 | 2.88 | 0.001 | 2.67 | 0.001 |
| PNPLA3 (CG+GG) | 3.26 | 0.01 | -2.23 | 0.024 | -1.4 | 0.16 | -1.51 | 0.09 | 2.01 | 0.044 | -5.52 | 0.001 |

Abbreviations: HA, hyaluronic acid; PIIINP, N-terminal propeptide of type III procollagen; NAS, NAFLD activity score.