

**Supporting information for the manuscript:**

**Elevated transaminases potentiate the risk for emerging dysglycemia in children with overweight and obesity**

Florian Koutny, MD<sup>a,b,#</sup>, Robert Stein, MD<sup>c,d,#</sup>, Wieland Kiess, MD<sup>c,e</sup>, Daniel Weghuber, MD<sup>a,b</sup>, Antje Körner, MD<sup>c,e</sup>

#authors contributed equally

**Affiliations:**

<sup>a</sup>Department of Pediatrics, Paracelsus Medical University, Salzburg, Austria

<sup>b</sup>Obesity Research Unit, Paracelsus Medical University, Salzburg, Austria

<sup>c</sup>University of Leipzig, Medical Faculty, University Hospital for Children and Adolescents, Center for Pediatric Research, Leipzig, Germany

<sup>d</sup>Helmholtz Institute for Metabolic, Obesity and Vascular Research (HI-MAG) of the Helmholtz Zentrum München at the University of Leipzig and University Hospital Leipzig, Leipzig, Germany

<sup>e</sup>Leipzig Research Center for Civilization Diseases (LIFE Child), Leipzig, Germany

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**Address correspondence to:**

Antje Körner, MD

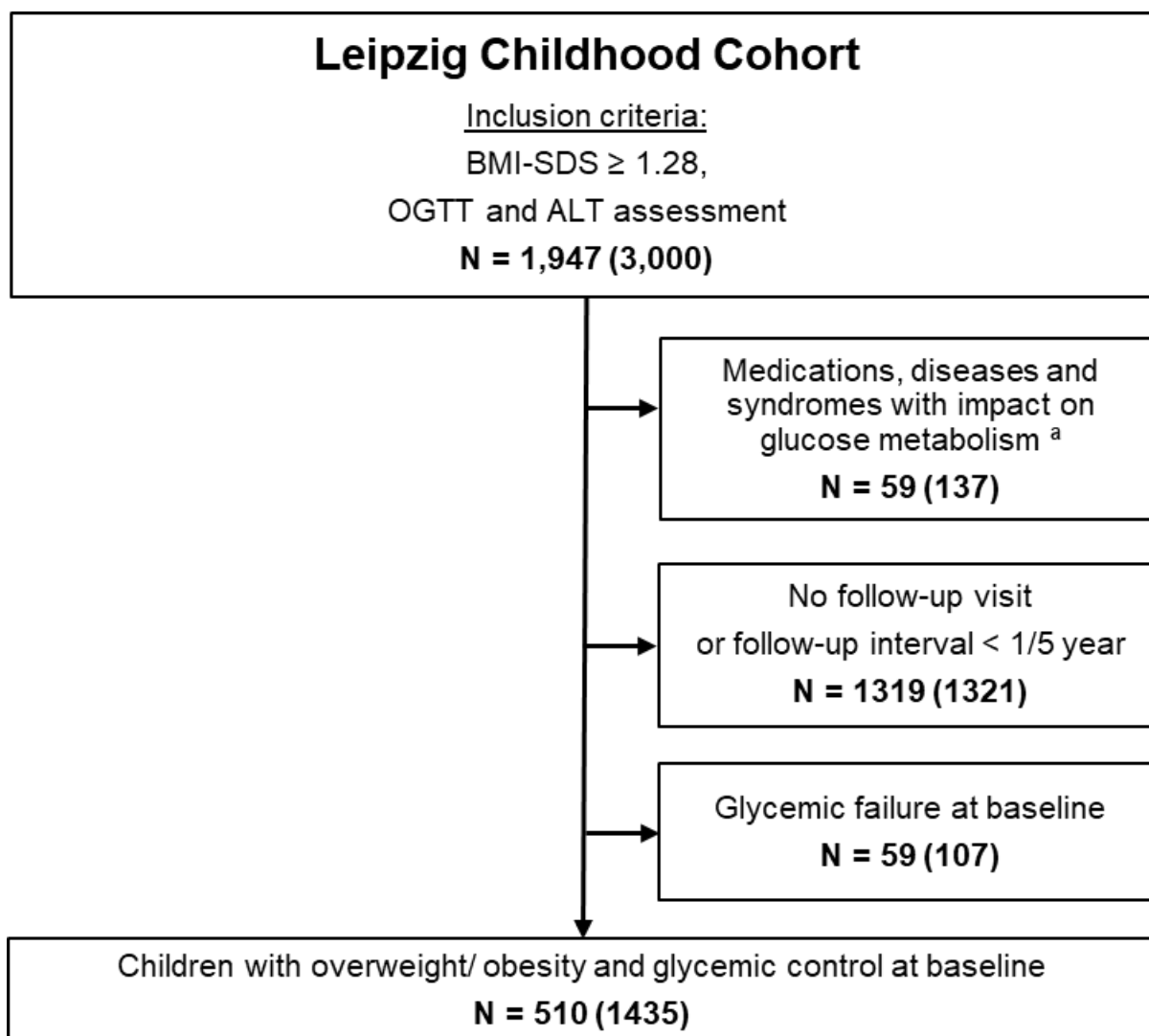
Center for Pediatric Research Leipzig

University Hospital for Children & Adolescents

Liebigstr. 20a, 04103 Leipzig, Germany

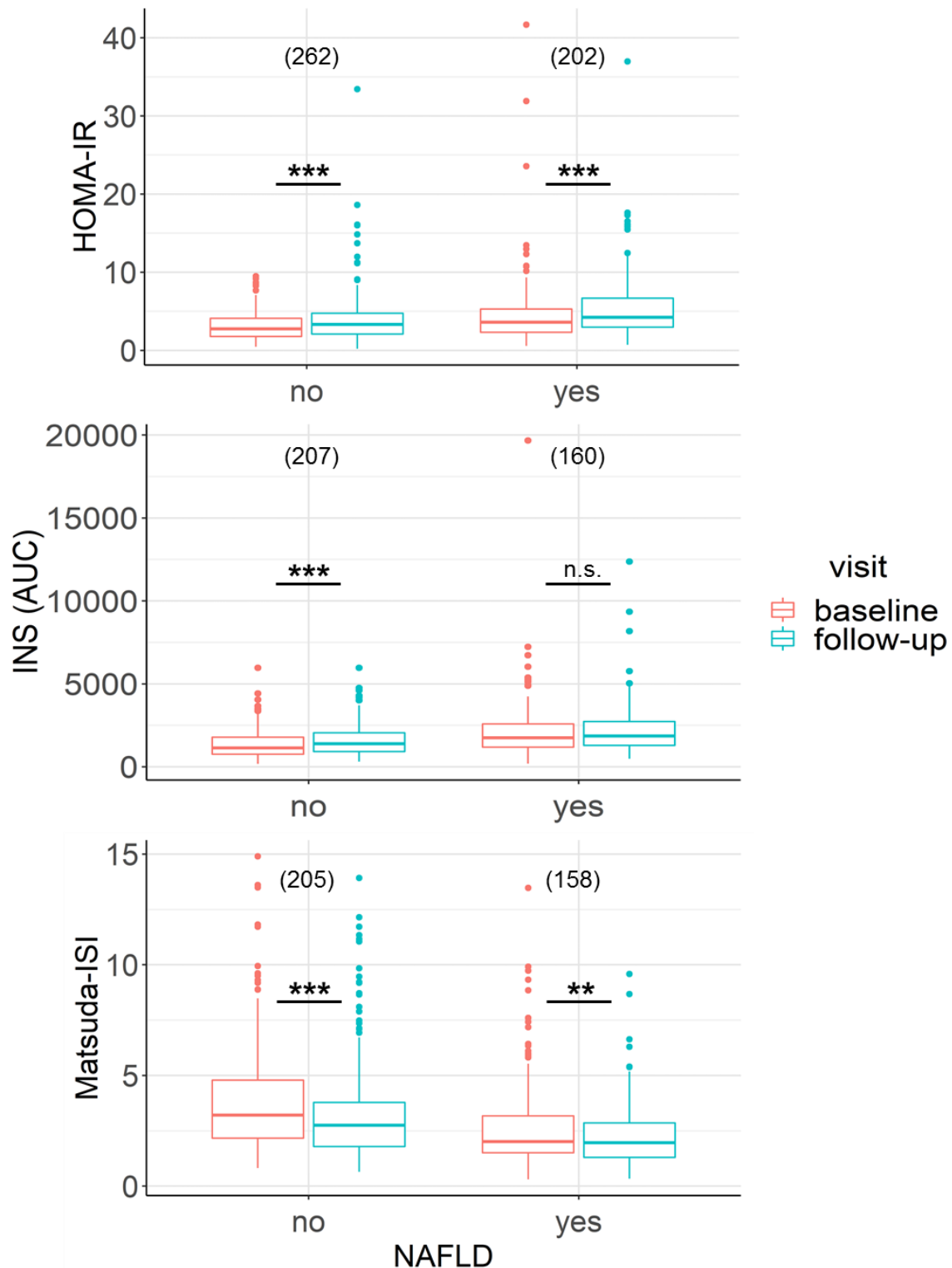
Email: Antje.Koerner@medizin.uni-leipzig.de

Phone: +49-341-9726500



**Figure S1: Selection of study population.** We included all children and adolescents with overweight or obesity from the Leipzig Childhood Cohort, who underwent OGTT and ALT assessment. <sup>a</sup> Participants with intake of medications affecting glucose metabolism (insulin, antipsychotics, retinoids, systemic glucocorticoids, growth hormone, immunosuppressives), as well as diseases affecting glucose metabolism (type 1 diabetes, rheumatic diseases, malabsorption, pancreatitis, spinal muscular atrophy, congenital adrenal hyperplasia, congenital metabolic diseases, nephrotic syndrome, cancer) and syndromes (Prader-Willi, Bardet-Biedl, Trisomy 16, Trisomy 21, Beckwith-Wiedemann, Ehlers-Danlos, Trichorhinophalangeal, Kabuki, Poland, Noonan, Klinefelter, Microdeletion 15q13.3) were excluded from the analyses.

Abbreviations: *ALT*, alanine aminotransferase; *BMI-SDS*, standard deviation score of body mass index; *N*, number of participants (number of observations); *OGTT*, oral glucose tolerance testing.



**Figure S2: Insulin resistance over time.** We compared measures of insulin resistance and sensitivity among participants with elevated transaminases (NAFLD=yes) and with normal range transaminases (NAFLD=no) between the baseline visit and the last follow-up visit. Only participants without metformin intake and data available at both visits were selected prior to this analysis, the respective number of participants is indicated in brackets above each plot. Hepatic insulin resistance (HOMA-IR) diminishes over time among children with overweight/obesity and elevated ALT, whereas peripheral insulin resistance ( $INS_{AUC}$ ) and peripheral insulin sensitivity (Matsuda-ISI) remains stable or only worsens slightly within the elevated ALT-group. Statistical analysis was performed with a two-sided paired t-test after logarithmic transformation. \*\*  $p < 0.01$ ; \*\*\*  $p < 0.001$ ; n.s., not significant; HOMA-IR, homeostasis model assessment for insulin resistance,  $INS_{AUC}$ , area under the insulin curve during OGTT, Matsuda-ISI, insulin sensitivity index according to Matsuda during OGTT.

**Table S1: Subgroups of dysglycemia**

<b>Outcome</b>	<b>N</b>	<b>Portion of total (%)</b>
Dysglycemia (total)	62	100
Metformin intake	29	46.8
IFG + IGT	29	46.8
IFG + DGT	2	3.2
DFG + IGT	1	1.6
DFG + DGT	1	1.6

Abbreviations: *DFG*, diabetic fasting glucose, *DGT*, diabetic glucose tolerance, *IFG*, impaired fasting glucose, *IGT*, impaired glucose tolerance