

This is the peer reviewed version of the following article:

Ortiz-Pinto, Maira Alejandra; Ortiz-Marrón, Honorato; Ferriz-Vidal, Isabel; Martínez-Rubio, María V; Esteban-Vasallo, María; Ordobás-Gavin, María; Galan, Iñaki I. **Association between general and central adiposity and development of hypertension in early childhood.** Eur J Prev Cardiol. 2019 Aug;26(12):1326-1334.

which has been published in final form at:

<https://doi.org/10.1177/2047487319839264>

1 **Association between general and central adiposity and development of hypertension**
2 **in early childhood**

3 Authors: Ortiz-Pinto MA^{1,2,3}, Ortiz-Marrón H⁴, Ferriz Vidal I⁵, Martínez-Rubio MA⁶,
4 Esteban-Vasallo M⁷, Ordobás-Gavin MA⁴, Galán I^{1,2}

5

6 ¹ Centro Nacional de Epidemiología. Instituto de Salud Carlos III. Madrid (España)

7 ² Departamento de Medicina Preventiva y Salud Pública. Universidad Autónoma de
8 Madrid/ IdiPAZ. Madrid (España)

9 ³ Departamento de Salud Pública. Universidad Del Norte. Barranquilla (Colombia)

10 ⁴ Servicio de Epidemiología. Dirección General de Salud Pública. Consejería de Sanidad.
11 Madrid. España

12 ⁵ Centro de Salud Valde las Fuentes. Consejería de Sanidad. Alcobendas (Madrid), España

13 ⁶ Centro de Salud los Fresnos. Consejería de Sanidad. Torrejón de Ardoz (Madrid), España

14 ⁷ Servicio de Informes de Salud y Estudios. Dirección General de Salud Pública. Consejería
15 de Sanidad. Madrid, España

16

17

18

19

20

21 **Corresponding author:**

22 Iñaki Galán, MD, PhD.

23 National Centre for Epidemiology

24 Instituto de Salud Carlos III

25 Monforte de Lemos 5,

26 28029 Madrid (Spain)

27 Telephone: +34 918222679

28 Fax: +34 913877815

29 E-mail: igalan@isciii.es

30

31

32

1 **Abstract**

2 **Objectives:** To evaluate the association of general and abdominal obesity with high blood pressure
3 in young children.

4 **Methods:** Longitudinal study including 1796 participants from the Madrid region (Spain) with
5 baseline at age 4 and a follow-up 2 years later. Blood pressure (BP), body mass index (BMI) and
6 waist circumference were measured during a physical examination. We evaluated the association
7 between obesity at baseline and weight changes between the ages of 4 and 6 and high BP. Data were
8 analyzed using linear and logistic regressions adjusted for covariates.

9 **Results:** Obese 4 year-olds (general or abdominal obesity), experienced an average 4-5 mmHg
10 increase in systolic BP and 2.5-3 mmHg increase in diastolic BP by the age of 6. Compared to
11 children maintaining a non-excess weight (based on BMI) during follow-up, incident and persistent
12 cases of excess weight (overweight or obesity) had an OR for high BP=2.49 (95%CI:1.50 to 4.13)
13 and OR=2.54 (95%CI:1.27 to 5.07), respectively. Regarding abdominal obesity we estimated
14 OR=2.81 (95%CI:0.98 to 8.02) for incident cases and OR=3.42 (95%CI:1.38 to 8.49) for persistent
15 ones. Similar estimates for the waist-height ratio were observed. Individuals who experienced
16 remission to non-excess weight did not have an increased risk for high BP.

17 **Conclusions:** We observed an increased risk for high BP among 4 year-olds who presented
18 persistent or incident cases of excess weight (BMI) or abdominal obesity after two years of follow-
19 up. Children with excess weight or obesity at baseline who remitted to non-excess weight did not
20 exhibit an increased risk for high BP.

21 **Key words:** childhood, obesity, abdominal obesity, arterial hypertension.

22

23

24

25

1 **Introduction**

2 In 2010 the childhood obesity epidemic affected 43 million children worldwide. Further,
3 overweight and obesity have increased between 2- and 3-fold in preschool age over the last three
4 decades⁽¹⁾. Similarly to other Southern European countries, Spain has a very high prevalence of
5 childhood obesity⁽²⁾.

6 Children and adolescents with excess body fat present with adverse cardiometabolic parameters that
7 support the onset of cardiovascular complications, even if no clinical symptoms are detected in
8 childhood⁽³⁾. Compared with children of normal weight, children with overweight and obesity show
9 an average increase of 4.5 and 7.5 mmHg of systolic blood pressure (BP), respectively⁽³⁾. In
10 addition, the presence of high BP in childhood increases the risk of adult hypertension⁽⁴⁾. Analyses
11 of trajectories and effects of childhood overweight and obesity on BP conclude that as excess
12 weight drops, so does high BP⁽⁵⁻¹¹⁾. This evidence-based support for adequate weight control from
13 an early age as a form of prevention of future cardiometabolic complications makes for a highly
14 relevant public health message⁽¹²⁾.

15 Because the use of the body mass index (BMI) as the single indicator of general adiposity is
16 controversial, the collection of additional anthropometric measures is recommended. For instance,
17 combining BMI and waist circumference, a better indicator of central adiposity and visceral fat, is
18 likely to increase sensitivity in the detection of overweight and obesity⁽¹³⁾. This in turn, will enable a
19 better assessment of the link between excess weight and cardiometabolic complications⁽¹⁴⁾. The
20 overall objective of this study was to evaluate the association of general overweight, general obesity
21 and abdominal obesity with high BP in a representative population-based sample of 4-year-old
22 children after 2 years of follow-up.

23 **Materials and Methods**

24 **Study Population**

25 The Longitudinal Study of Childhood Obesity (ELOIN for its acronym in Spanish), is a population-
26 based cohort of the Madrid region, Spain (population: 6.5 million). Study methodology and sample
27 characteristics have been previously published⁽¹⁵⁾. A total of 2,627 children were involved in the
28 original cohort. The number of participants who underwent a physical examination at 4 (baseline
29 measurement) and 6 years of age were 1,796. The information was collected in two consecutive
30 stages: the child's standardized physical examination performed at the health center by the

1 pediatricians participating in the study, followed by a computer-assisted telephone interview of the
2 parents based on a structured questionnaire.

3 **Anthropometric measurements**

4 The physical examination included anthropometric measures collected in a standardized manner by
5 the 32 pediatricians in the study. Weight was assessed using a digital scale (SECA® model 220,
6 precision 0.1 kg); height was measured by means of telescopic stadiometer (SECA® model 220,
7 precision 1.0 mm); waist circumference (WC) was measured with a validated inextensible metric
8 tape with a buckle, just above the iliac crests with the tape horizontally and without compression of
9 the tissues. Two measurements were collected for each participant, and the average was used for the
10 analyses.

11 We calculated the children's BMI (kg/m²) standardizing it for age in months and sex according to
12 the WHO-2007 reference tables. Based on BMI z-scores, overweight was defined as z-BMI > +1
13 standard deviation (SD) and ≤ +2 SD whereas obesity was defined as z-BMI > +2 SD⁽¹⁶⁾.

14 The mean WC value was standardized using the age (in months) and sex reference tables proposed
15 by Fernández et al⁽¹⁷⁾. Abdominal obesity was defined using two criteria: 1) ≥90-percentile of WC
16 according to the recommendations of the International Diabetes Federation (IDF)⁽¹⁸⁾; 2) ≥90-
17 percentile of the WC-height ratio (cm/m)⁽¹⁹⁾.

18 Individuals classified as overweight or obese based on their z-BMI were grouped under the
19 variable "excess weight." Both maintenance and variation of excess weight were captured by
20 defining four categories which took into account an individual's classification at baseline and at
21 follow-up: a) Maintenance of non-excess weight-BMI, i.e., presenting non-excess weight status at
22 both time points; b) Maintenance of excess weight-BMI, i.e., presenting in-excess weight at both
23 time points; c) Incidence, i.e., new cases of excess weight-BMI at follow-up; and d) Remission:
24 recuperating a non-excess weight-BMI status by follow-up. The same categories were defined to
25 evaluate changes in abdominal obesity during follow-up.

26 **Blood pressure**

27 Blood pressure (BP) was measured using the auscultatory method in the right arm. Participants
28 remained seated at rest for 5 minutes before the assessment, with the back and feet supported. The
29 right arm was resting on a firm surface in a horizontal position with the palm of the hand facing up
30 so that the antero-cubital fossa was located at heart level. We measured BP 2 times separated by at

1 least 2 minutes. When differences between both measures were greater than 5 mmHg a third
2 measurement was taken. The average of the measurements was used for analytical purposes.

3 BP values were standardized according to height, age in months, and sex, using the reference tables
4 on *The Fourth Report on High Blood Pressure in Children and Adolescents*⁽²⁰⁾. According to the
5 recommendations of the European Society of Hypertension for Children and Adolescents⁽²¹⁾, high
6 BP was defined using the 90th percentile as the cutoff point for systolic and/or diastolic blood
7 pressure.

8 ***Covariables***

9 Baseline sociodemographic variables collected when participants were 4 years old included sex, age
10 in months, mother's educational achievement, and family socioeconomic status (SES) based on the
11 Family Affluence Scale (low, medium and high)⁽²²⁾. Other covariables included time of exclusive
12 breastfeeding, rest (hours of sleep and leisure), weekly physical exercise (hours at school and
13 extracurricular), and weekly sedentary activities (hours spent in front of screens: television, playing
14 with the computer or the console).

15 ***Statistical analysis***

16 From the initial sample composed of 1796 participants with valid values for systolic and diastolic
17 blood pressure, 15 children were excluded who did not report values for WC. An additional 170
18 children were excluded from all elevated BP analyses because of having altered BP already at age
19 4. After removing participants with missing values in the covariates of interest, the final sample
20 sizes used in the different models of association between BP and overweight-general and abdominal
21 obesity ranged between 1612 and 1796 individuals.

22 We applied linear regression modelling to estimate mean changes in systolic and diastolic BP.
23 Second, we used logistic regression models to estimate the odds ratios (OR) of having high BP by
24 age 6. These models also included the different anthropometric categories capturing weight changes
25 during follow-up.

26 Models were adjusted for the covariates previously described, specifically, all analyses included
27 sex, age, family SES, BP baseline values, as well as those covariates with p-values<0.20 at the
28 bivariate level. Finally, we explored potential interactions of the indicators for overweight-general
29 obesity and abdominal obesity with the variables for sex and SES. All the analyses were performed

1 using Stata v.14 and its Survey Data module to take into account the characteristics of the cluster
2 sampling (pediatric consultations).

3

4 ***Ethical considerations***

5 The study protocol was approved by the Ethics Committee of the Ramón y Cajal Hospital in
6 Madrid. Given that all participants were minors, their parents gave their informed consent in
7 writing.

8 **Results**

9 The characteristics of the sample are described in Table 1. Higher systolic BP was observed in
10 children with low SES, excess weight, and incident and persistent abdominal obesity.

11 Table 2 shows the mean systolic and diastolic BP changes by age 6, according to the classification
12 of weight status and abdominal obesity at age 4 (baseline). Children with overweight at baseline
13 were associated with an increase in BP in two years later, by age 6. Models estimate a β coefficient
14 for systolic pressure = 1.79 (95% CI: 0.71 to 2.86) and β for diastolic pressure= 1.36 (95% CI: 0.15
15 to 2.58). Being obese at baseline was associated to a greater increase in BP, observing β coefficients
16 of 4.77 (95%CI: 2.85 to 6.69) and 3.08 (95%CI: 0.63 to 5.52) for systolic and diastolic pressure
17 respectively. Abdominal obesity according to WC was associated with an increase of 4.43 mmHg
18 (95%CI: 3.01 to 5.85) and 2.48 mmHg (95%CI: 0.56 to 4.40) of the systolic and diastolic pressure
19 respectively. The BP increase associated to baseline waist-height ratio was smaller, being the
20 increase in systolic pressure=2.46 (95%CI: 0.97 to 3.95).

21 Table 3 shows the longitudinal analysis of the variation in excess weight and abdominal obesity,
22 and the mean BP by age 6. Compared to children who maintained a non-excess weight based on
23 BMI or abdominal obesity at either assessment, incident and persistent cases of excess weight or
24 abdominal obesity presented a greater increase in mean BP. The greatest increases in systolic
25 pressure were associated to WC for incident cases (β =7.21; 95%CI: 4.18 to 10.24) as well as for
26 persistent cases (β =7.15; 95%CI: 5.12 to 9.18). In contrast, the greatest increases in diastolic BP
27 were associated with the waist-height ratio, estimating a β =4.03 (95%CI: 2.10 to 5.95) for incident
28 cases and β =4.59 (95%CI: 1.78 to 7.41) for persistent cases. Except regarding WC-based abdominal
29 obesity, individuals experiencing obesity reversal showed similar values to those of participants
30 who maintained a non-excess weight based on BMI or abdominal obesity, or slightly higher but
31 without reaching statistical significance.

1 Table 4 shows crude and adjusted risks of having high BP at age 6, taking into account weight
2 status and abdominal obesity at age 4. In the adjusted model, general obesity (BMI) and abdominal
3 obesity (WC) were associated with high BP (OR=2.55; 95% CI: 1.24 to 5.24; OR=1.99; 95% CI:
4 1.00 to 3.95, respectively).

5 Table 5 shows the results of the longitudinal risk analysis of having high BP at age 6, with respect
6 to the persistence or variation of excess weight and abdominal obesity. The association with the
7 greatest magnitude was observed among persistent cases of WC-based abdominal obesity
8 (OR=3.42; 95%CI: 1.38 to 8.49), followed closely by individuals with persistent obesity according
9 to the waist-height quotient (OR=3.17; 95%CI: 1.12 to 8.91) and excess weight (BMI), OR=2.54
10 (95%CI: 1.27 to 5.07).

11 Incidence of both excess weight (according to BMI) and abdominal obesity (according to waist-
12 height ratio) during the follow-up period were associated to increased risk of high BP presentation:
13 OR=2.49 (95%CI: 1.50 to 4.13) and OR=3.01 (95%CI: 1.44 to 6.28), respectively. When compared
14 to children who maintained non-excess weight, those who remitted excess weight did not present
15 statistically significant differences in risk. We observed the same result regarding abdominal
16 obesity. Finally, no statistically significant interactions with sex or family purchasing level were
17 identified.

18 Finally, we examined how both indices contributed to increases in BP by introducing them
19 simultaneously in the linear regression model (Supplementary Table 1). We observed independent
20 effects on systolic BP increase despite the high correlation between the indices. We tested a model
21 with the four combinations of obesity at 4 years of age and the increase in systolic BP was
22 statistically significant in all of them. I.e., obese patients classified only by BMI had an increase of
23 3.98 mmHg of systolic BP; the obese classified by waist circumference showed an increase of 3.62
24 mmHg; and the obese classified by both measurements had an increase of 5.03 mmHg
25 (Supplementary Table 2). These results support the combined use of both general and abdominal
26 obesity measures as a determinant factor influencing BP.

27

28 **Discussion**

29 After two years of follow-up, children diagnosed with either general or abdominal obesity at the age
30 of 4 showed an average increase of 4-5 mmHg systolic and 2.5-3 mmHg diastolic blood pressure.
31 Based on the resulting odds ratios, the probability of having high blood pressure was 2-2.5 times
32 higher in the obese children. Both the persistent and incident cases of excess weight, general and

1 abdominal obesity, showed an increased risk of blood pressure elevation, whereas cases
2 experiencing remission presented values similar to children with no-excess weight.

3 There is consistent evidence from numerous longitudinal studies supporting the relationship
4 between excess weight and the increase in systolic and diastolic BP in children and adolescents<sup>(5-
5 11,23-27)</sup>. Although comparisons with this literature are problematic due to substantial variations in
6 age groups, lengths of follow-up, and indicators included, the average increase in BP observed in
7 our study is quite similar to those reported by others.

8 Considering that an individual's overweight or obese status can change throughout childhood and
9 adolescence, the different categories of the anthropometric indicators and their effect on
10 cardiometabolic risk, and more specifically, on BP have been analyzed^(5-11,23,25,26). Both persistent
11 and incident cases of excess weight were associated with an increase in BP. In addition, when
12 excess weight remitted during the follow-up, BP decreased⁽⁵⁻¹¹⁾. Here we observed the same
13 associations already at very early ages and with only two years of follow-up, underscoring the
14 importance of controlling excess weight early in life to maintain healthy BP values.

15 Although in their systematic review Friedman et al. reported a greater effect of obesity both on
16 systolic and diastolic pressure in girls than boys⁽³⁾, our data pointed to similar effect in both sexes
17 supporting findings reported in previous longitudinal studies^(5,8,23,24,27). It is possible that the
18 detection of those sex differences by Friedman et al.,⁽³⁾ could be the result of analyzing both
19 longitudinal and cross-sectional studies together.

20 Nevertheless, an understudied question was to know if socioeconomic status could modify the
21 association between adiposity and BP. Consistent with our results, Howe et al.,⁽²⁸⁾ reported that
22 adiposity did not mediate the socioeconomic status differentials in BP.

23 What is also under discussion is the role the different anthropometric indicators play on
24 cardiovascular risk. Recent work proposed waist circumference as a better marker of visceral
25 adiposity than BMI, based on the former's stronger association with obesity-related
26 complications⁽¹⁴⁾. Other authors have proposed the waist circumference corrected for height as the
27 better predictor of cardiometabolic risk. This measure has the obvious advantage of its applicability
28 to different population subgroups without standardization according to reference values⁽²⁹⁾.

29 Studies examining both indicators of general and abdominal obesity failed to find significant
30 differences in their associations with BP^(5,9,24). Similarly, we did not find substantial differences
31 between these two indicators, except that the associations between waist circumference corrected by
32 the height and high BP were, in general, of smaller magnitude. In addition, the area under the

1 receiver operating characteristic (ROC) curves which, in this case, evaluated the indicators'
2 capacity to discriminate between children with and without elevated blood pressure, were similar
3 across our three different adiposity measures at baseline and blood pressure values two years later:
4 BMI (0.7093); WC (0.7059); and WC-height ratio (0.7033). Regardless, it is important to keep in
5 mind that obesity, estimated based on BMI and waist circumference separately, has a sensitivity of
6 80-85%⁽³⁰⁾. That is, either measurement separately fails to detect one of every 5-6 cases of obesity
7 and, thus, the recommendation in pediatrics is the simultaneous use of both measures⁽¹³⁾.

8 ***Limitations and strengths of the study***

9 When interpreting our results, some limitations should be taken into consideration. First, children of
10 foreign or low educational level mothers had a lower response rate at baseline introducing a
11 moderate bias in the selection of the sample which may affect its representativeness at the
12 population level. Second, most of the children in our study are Caucasians of Spanish origin, so our
13 findings cannot be generalized to other ethnicities. Third, the 2-year follow-up does not allow the
14 evaluation of how the association between BP and body weight changes in the mid- to long-term.
15 Finally, given the low prevalence of obese 4-year olds at baseline (5%) and follow-up (9.7%) we
16 combined the two categories, overweight and obese, into an “excess weight” category to maintain
17 statistical power in the longitudinal analyses.

18 It should also be noted that the average systolic and diastolic BP data used to defined high BP came
19 from a single visit to the pediatrician and not from 3 separate measurements, as recommended by
20 the European Society of Hypertension. Finally, taking into account the model's goodness-of-fit,
21 adiposity would explain only 5-8% of variability (depending on the indicators) in unadjusted
22 models, only improving to 7-10% in the adjusted ones. However, as this modest goodness-of-fit
23 mirrors that of those reported in other studies⁽²⁶⁾, our results support previous evidence pointing to
24 the existence of other factors, not included in the analysis, that may explain the residual variance
25 and confound the results.

26 Among the study's strengths, it is worth highlighting its longitudinal design allowing causal
27 associations to be suggested. Another important characteristic is that the sample is representative of
28 the 4-year-old population of the Madrid region, despite of the aforementioned selection bias.
29 Finally, the anthropometric measures, which include various indices and BP, were based on
30 objective tests performed in a standardized manner with baseline measurements at 4 years of age
31 and a follow-up 2 years later. Individuals with high BP at baseline were excluded from the logistic

1 regression analyses estimating incident cases during follow-up. Finally, all models were adjusted
2 for baseline BP values.

3 Pediatricians play a key role in the prevention of adult cardiovascular diseases since the onset of
4 many of the related risk factors occur in childhood. In fact, there is an ongoing heated debate about
5 the pertinence of universal screening for hypertension in children. However, although
6 recommended by numerous clinical guides, evidence supporting the implementation of such a
7 program is still very limited⁽¹²⁾. Notwithstanding, the proposal of a screening program for
8 hypertension, targeted at at-risk groups such as children with excess weight or with general or
9 abdominal obesity, should be evaluated to be integrated into clinical practice.

10 In conclusion, 4 year-olds presenting excess weight based on BMI or abdominal obesity at the
11 beginning of the study, displayed a higher risk of BP increase by the time they turned 6. Incident
12 and persistent cases of excess weight and abdominal obesity were associated with an increased risk
13 of high BP, whereas those who reverted to a normal weight decreased their BP to levels comparable
14 to those with normal anthropometric values at the study's baseline and throughout the follow-up.
15 Therefore, early detection and control of childhood obesity may reduce the risk of high blood
16 pressure.

17 **Acknowledgements**

18 The authors thank all participating families, the pediatricians from the Network of Sentinel
19 Doctors of the Community of Madrid and the two companies interviewing the families:
20 Demométrica SL and Sondaxe SL.

21 **Author Contributions**

22 Ortiz-Pinto MA and Galán I, conceptualized and designed the study, drafted the initial
23 manuscript, and reviewed and revised the manuscript.

24 Ortiz-Pinto MA, Ortiz-Marrón H and Galán I, designed the data collection instruments,
25 collected data, carried out the initial analyses, Ortiz-Pinto MA, Ortiz-Marrón H, Esteban-
26 Vasallo M, Ordobás-Gavin MA, Ferriz Vidal I, Martínez-Rubio MA and Galán I revised the
27 manuscript.

28 Ortiz-Marrón H, Ordobás-Gavin MA, Ferriz Vidal I, Martínez-Rubio MA, collected data,
29 and critically reviewed the manuscript for important intellectual content.

1 All authors approved the final manuscript as submitted and agree to be accountable for all
2 aspects of the work.

3 **Bibliography**

- 4 1. De-Onis M, Blössner M. Prevalence and trends of overweight among preschool children in
5 developing countries. *Am J Clin Nutr* 2000;72(4):1032-9
- 6 2. Ahrens W, Pigeot I, Pohlabeln H, De Henauw S, Lissner L, Molnár D, et al. Prevalence of
7 overweight and obesity in European children below the age of 10. *Int J Obes (Lond)* 2014;38
8 (Suppl 2):S99-107
- 9 3. Friedemann C, Heneghan C, Mahtani K, Thompson M, Perera R, Ward AM. Cardiovascular
10 disease risk in healthy children and its association with body mass index: systematic review and
11 meta-analysis. *BMJ* 2012;345(sep25 2):e4759-e4759
- 12 4. Chen X, Wang Y. Tracking of blood pressure from childhood to adulthood: a systematic review
13 and meta-regression analysis. *Circulation* 2008;117(25):3171-3180
- 14 5. Lawlor DA, Benfield L, Logue J, Tilling K, Howe LD, Fraser A, et al. Association between
15 general and central adiposity in childhood, and change in these, with cardiovascular risk factors
16 in adolescence: prospective cohort study. *BMJ* 2010;341(nov25 1):c6224-c6224
- 17 6. Huang R-C, Burrows S, Mori TA, Oddy WH, Beilin LJ. Lifecourse Adiposity and Blood
18 Pressure Between Birth and 17 Years Old. *Am J Hypertens* 2015;28(8):1056-1063
- 19 7. Mamun AA, Lawlor DA, O'Callaghan MJ, Williams GM, Najman JM. Effect of body mass
20 index changes between ages 5 and 14 on blood pressure at age 14: findings from a birth cohort
21 study. *Hypertension* 2005;45(6):1083-1087
- 22 8. Berentzen NE, van Rossem L, Gehring U, Koppelman GH, Postma DS, de Jongste JC, et al.
23 Overweight patterns throughout childhood and cardiometabolic markers in early adolescence.
24 *Int J Obes (Lond)* 2016;40(1):58-64
- 25 9. Bekkers MBM, Brunekreef B, Koppelman GH, Kerkhof M, de Jongste JC, Smit HA, et al. BMI
26 and waist circumference; cross-sectional and prospective associations with blood pressure and
27 cholesterol in 12-year-olds. *PLoS ONE* 2012;7(12):e51801

- 1 10. Howe LD, Chaturvedi N, Lawlor DA, Ferreira DLS, Fraser A, Davey Smith G, et al. Rapid
2 increases in infant adiposity and overweight/obesity in childhood are associated with higher
3 central and brachial blood pressure in early adulthood. *J Hypertens* 2014;32(9):1789-1796
- 4 11. Juonala M, Magnussen CG, Berenson GS, Venn A, Burns TL, Sabin MA, et al. Childhood
5 adiposity, adult adiposity, and cardiovascular risk factors. *N Engl J Med* 2011;365(20):1876-
6 1885
- 7 12. Bloetzer C, Bovet P, Suris J-C, Simeoni U, Paradis G, Chiolerio A. Screening for cardiovascular
8 disease risk factors beginning in childhood. *Public Health Rev* 2015;36:9. doi: 10.1186/s40985-
9 015-0011-2. eCollection. 2015
- 10 13. Schröder H, Ribas L, Koebnick C, Funtikova A, Gomez SF, Fíto M, et al. Prevalence of
11 abdominal obesity in Spanish children and adolescents. Do we need waist circumference
12 measurements in pediatric practice? *PLoS One* 2014;9(1):e87549
- 13 14. Dobashi K. Evaluation of Obesity in School-Age Children. *J Atheroscler Thromb* 2016;
14 23(1):32-38
- 15 15. Ortiz-Marrón H, Cuadrado-Gamarra JI, Esteban-Vasallo M, Cortés-Rico O, Sánchez-Díaz J,
16 Galán-Labaca I. Estudio Longitudinal de Obesidad Infantil (ELOIN): diseño, participación y
17 características de la muestra. *Rev Española de Cardiol* 2016;69(5):521-523.
- 18 16. Onis M de, Adelheid W Onyango, Borghi E, Siyam A, Nishida C, Siekmann J. Development
19 of a WHO growth reference for school-aged children and adolescents. *Bull World Health*
20 *Organ* 2007; 85(Suppl 9): 660-667
- 21 17. Fernández JR, Redden DT, Pietrobelli A, Allison DB. Waist circumference percentiles in
22 nationally representative samples of African-American, European-American, and Mexican-
23 American children and adolescents. *J Pediatr* 2004; 145(4):439-444
- 24 18. Alberti G, Zimmet P, Kaufman F, Tajima N, Silink M, Arslanian S et al. The IDF consensus
25 definition of the metabolic syndrome in children and adolescents. *Pediatr Diabetes*
26 2007;8(5):299-306

- 1 19. Nambiar S, Hughes I, Davies PS. Developing waist-to-height ratio cut-offs to define
2 overweight and obesity in children and adolescents. *Public Health Nutr* 2010; 13(10):1566-
3 1574
- 4 20. National High Blood Pressure Education Program Working Group on High Blood Pressure in
5 Children and Adolescents B. The fourth report on the diagnosis, evaluation, and treatment of
6 high blood pressure in children and adolescents. *Pediatrics* 2004; 114(2 Suppl 4th Report):555-
7 576
- 8 21. Lurbea E, Agabiti-Rosei E, Cruickshank JK, Dominiczak A, Erdine S, Hirth A et al. 2016
9 European Society of Hypertension guidelines for the management of high blood pressure in
10 children and adolescents. *J Hypertens* 2016;34(1):1887-1920
- 11 22. Boyce W, Torsheim T, Currie C, Zambon A. The family affluence scale as a measure of
12 national wealth: Validation of an adolescent self-report measure. *Soc Indic Res* 2006;78(Suppl
13 3):473-487
- 14 23. Marcus MD, Foster GD, El Ghormli L, Baranowski T, Goldberg L, Jago R, et al. Shifts in BMI
15 category and associated cardiometabolic risk: prospective results from HEALTHY study.
16 *Pediatrics* 2012;129(4):e983-991
- 17 24. Maximova K, O'Loughlin J, Paradis G, Hanley JA, Lynch J. Changes in anthropometric
18 characteristics and blood pressure during adolescence. *Epidemiology* 2010;21(3):324-331
- 19 25. Munthali RJ, Kagura J, Lombard Z, Norris SA. Childhood adiposity trajectories are associated
20 with late adolescent blood pressure: birth to twenty cohort. *BMC Public Health* 2016;16: 665
- 21 26. Jones A, Charakida M, Falaschetti E, Hingorani AD, Finer N, Masi S, et al. Adipose and height
22 growth through childhood and blood pressure status in a large prospective cohort study.
23 *Hypertension* 2012;59(5):919-925
- 24 27. Tu W, Eckert GJ, DiMeglio LA, Yu Z, Jung J, Pratt JH. Intensified effect of adiposity on blood
25 pressure in overweight and obese children. *Hypertension* 2011;58(5):818-824
- 26 28. Howe LD, Galobardes B, Sattar N, Hingorani AD, Deanfield J, Ness AR, et al. Are there
27 socioeconomic inequalities in cardiovascular risk factors in childhood and are they mediated by
28 adiposity? Findings from a prospective cohort study. *Int J Obes (Lond)*. 2010;34(7):1149-1159

- 1 29. Browning LM, Hsieh SD, Ashwell M. A systematic review of waist-to-height ratio as a
2 screening tool for the prediction of cardiovascular disease and diabetes: 0.5 could be a suitable
3 global boundary value. *Nutr Res* 2010;23(2):247-269
- 4 30. Simmonds M, Llewellyn A, Owen CG, Woolacott N. Simple tests for the diagnosis of
5 childhood obesity: a systematic review and meta-analysis. *Obes Rev* 2016;17(12):1301-1315
- 6

- 1 Table 1. Systolic and Diastolic blood pressure at follow-up (age 6), according to the characteristics
- 2 of the sample.

	N	Systolic blood pressure (mmHg)		Diastolic blood pressure (mmHg)	
		Mean (SD)	p-value	Mean (SD)	p-value
Sex			<0.001		0.023
Males	908	93.4(8.4)		55.1(7.7)	
Females	888	91.8(8.5)		54.3(8.6)	
Maternal Educational level ^a			0.579		0.054
Primary or below	61	92.3(8.8)		54.4(8.7)	
First level secondary	330	93.0(8.8)		55.3(8.2)	
Second level secondary	608	92.9(8.5)		55.3(7.9)	
Some College/technical	249	92.1(8.2)		53.6(7.7)	
University level	542	92.2(8.3)		54.3(7.6)	
Family purchasing power			0.062		0.012
Low	817	93.1(8.7)		55.3(7.9)	
Medium	687	92.3(8.2)		54.2(7.9)	
High	291	91.9(8.4)		54.2(8.0)	
Number of months of breastfeeding ^a			0.553		0.310
No breastfeeding	181	92.3(8.5)		54.1(7.7)	
1 to 2 months	274	92.9(8.5)		54.7(8.0)	
3 to 5 months	712	92.7(8.5)		54.8(7.9)	
≥6 months	278	91.9(7.9)		54.2(7.5)	
TV and video games (hours/week) ^a			0.199		0.736
Under 6 hours	704	92.3(8.2)		54.6(7.6)	
6 to 9 hours	605	92.9(8.7)		54.9(8.0)	
> 9 hours	485	92.6(8.5)		54.7(8.2)	
Physical Activity (hours /week) ^a			0.343		0.372
< 2 hours	861	92.9(8.5)		55.0(7.9)	
2 to 4 hours	389	92.2(8.4)		54.2(7.8)	
> 4 hours	492	92.2(8.5)		54.6(7.9)	
Change in excess weight (BMI) ^b between ages 4 and 6			<0.001		<0.001
Persistent non-excess weight	1205	91.2(7.9)		53.8(7.5)	
Persistent excess weight	290	96.9(9.3)		57.5(8.6)	
Incident	201	95.9(8.0)		56.7(7.7)	
Remission	100	90.7(7.8)		53.8(8.3)	
Change in abdominal obesity between ages 4 and 6			<0.001		<0.001

according to waist circumference ^{a,c}				
Persistent non-abdominal obesity	1584	91.8(8.0)		54.2(7.6)
Persistent abdominal obesity	77	99.4(9.7)		58.3(9.8)
Incident	80	99.6(8.6)		59.1(7.4)
Remission	40	95.5(10.1)		57.3(10.3)
Change in abdominal obesity between ages 4 and 6			<0.001	<0.001
according to waist/height ratio ^{a,d}				
Persistent non-abdominal obesity	1523	91.9(8.1)		54.2(7.7)
Persistent abdominal obesity	96	99.1(9.3)		59.1(8.5)
Incident	84	98.3(8.3)		58.5(7.3)
Remission	78	91.6(9.3)		53.9(8.0)

1 SD: Standard Deviation

2 ^a Variables with missing values

3 ^b Excess weight (Body Mass Index) >+1 standard deviation according to WHO-2007 reference tables

4 ^c Abdominal obesity: $\geq 90^{\text{th}}$ percentile according to reference tables by Fernández et al.[17]

5 ^d Abdominal obesity: $\geq 90^{\text{th}}$ percentile of waist(cm)/height (cm) ratio

6

7

1 Table 2. Association between general overweight-obesity and abdominal obesity at 4 years of age,
 2 and systolic and diastolic blood pressure at 6 years of age

	Systolic blood pressure (mmHg)			Diastolic blood pressure (mmHg)		
	Coef. β^e	95%CI	p-value	Coef. β^e	95%CI	p-value
Body Mass Index (N=1796)						
Non-excess weight	(ref)			(ref)		
Excess weight ^a	1.79	(0.71 to 2.86)	0.002	1.36	(0.15 to 2.58)	0.029
Obesity ^b	4.77	(2.85 to 6.69)	<0.001	3.08	(0.63 to 5.52)	0.015
Waist Circumference (N=1788)						
Not obese	(ref)			(ref)		
Obese ^c	4.43	(3.01 to 5.85)	<0.001	2.48	(0.56 to 4.40)	0.013
Waist-height ratio (N=1788)						
Not obese	(ref)			(ref)		
Obese ^d	2.46	(0.97 to 3.95)	0.002	1.61	(0.01 to 3.21)	0.048

3

4 ^a Excess weight (Body Mass Index) >+1 standard deviation (SD) and \leq +2 SD according to WHO-2007 reference tables

5 ^b Obesity (Body Mass Index) >+2 SD according to WHO-2007 reference tables

6 ^c Abdominal Obesity: \geq 90th percentile according to reference tables by Fernández et al. [17]

7 ^d Abdominal Obesity: \geq 90th percentile of waist(cm)/height(cm) ratio

8 ^e β coefficients: estimated using linear regression, adjusted for sex, age, maternal educational level, family purchasing power, hours of
 9 TV screen or video games per week, hours of physical activity per week, breastfeeding, and baseline blood pressure (age 4)

Table 3. Association between persistence and variation of general overweight-obesity and abdominal obesity between the ages of 4 and 6, and systolic and diastolic blood pressure at age 6

	Systolic blood pressure (mmHg)			Diastolic blood pressure (mmHg)		
	Coef. β^d	95%CI	p-value	Coef. β^d	95%CI	p-value
Body Mass Index						
(N=1632) ^a						
Persistent non-excess weight	(ref)			(ref)		
Persistent excess weight	5.06	(3.56 to 6.56)	<0.001	3.18	(1.49 to 4.87)	0.001
Incident	4.85	(3.49 to 6.20)	<0.001	2.77	(1.69 to 3.86)	<0.001
Remission	-0.81	(-2.62 to 0.99)	0.368	0.04	(-2.10 to 2.20)	0.963
Waist Circumference						
(N=1612) ^b						
Persistent non-abdominal obesity	(ref)			(ref)		
Persistent abdominal obesity	7.15	(5.12 to 9.18)	<0.001	3.59	(1.13 to 6.06)	0.006
Incident	7.21	(4.18 to 10.24)	<0.001	3.80	(1.57 to 6.04)	0.002
Remission	2.78	(0.18 to 5.39)	0.037	2.43	(-0.52 to 5.38)	0.103
Waist/height ratio						
(N=1612) ^c						
Persistent non-abdominal obesity	(ref)			(ref)		
Persistent abdominal obesity	7.07	(4.77 to 9.36)	<0.001	4.59	(1.78 to 7.41)	0.002
Incident	6.43	(3.97 to 8.90)	<0.001	4.03	(2.10 to 5.95)	<0.001
Remission	0.30	(-1.69 to 2.31)	0.757	0.26	(-1.21 to 1.75)	0.713

^a Excess weight (Body Mass Index) >+1 standard deviation (SD) according to WHO-2007 reference tables

^b Abdominal Obesity: $\geq 90^{\text{th}}$ percentile according to reference tables by Fernández et al. [17]

^c Abdominal Obesity: $\geq 90^{\text{th}}$ percentile of waist(cm)/height(cm) ratio

^d β coefficients: estimated using linear regression, adjusted for sex, age, maternal educational level, family purchasing power, hours of TV screen or video games per week, hours of physical activity per week, breastfeeding, and baseline blood pressure (age 4)

Table 4. Association between general overweight-obesity and abdominal obesity at baseline (age 4), and high blood pressure at follow-up (age 6)

	High Blood Pressure					
	OR ^e	95%CI	p-value	OR ^f	95%CI	p-value
Body Mass Index (N=1796)						
Non-excess weight	1 (ref)			1 (ref)		
Overweight ^a	1.65	(1.16 to 2.34)	0.007	1.35	(0.93 to 1.96)	0.102
Obesity ^b	3.72	(1.85 to 7.47)	0.001	2.55	(1.24 to 5.24)	0.012
Waist Circumference (N=1788)						
Non obese	1 (ref)			1 (ref)		
Obese ^c	2.92	(1.37 to 6.22)	0.007	1.99	(1.00 to 3.95)	0.048
Waist/height ratio (N=1788)						
Non obese	1 (ref)			1 (ref)		
Obese ^d	2.00	(0.91 to 4.38)	0.079	1.33	(0.61 to 2.90)	0.475

^a Overweight (Body Mass Index) >+1 standard deviation (SD) and ≤ +2 SD according to WHO-2007 reference tables

^b Obesity (Body Mass Index) >+2 SD according to WHO-2007 reference tables

^c Abdominal Obesity: ≥90th percentile according to reference tables by Fernández et al. [17]

^d Abdominal Obesity: ≥90th percentile of waist(cm)/height(cm) ratio

^e Odds ratios: estimated using unadjusted logistic regression

^f Odds ratios: estimated using logistic regression adjusted for sex, age, maternal educational level, family purchasing power, hours of TV screen or video games per week, hours of physical activity per week, breastfeeding, and baseline blood pressure (age 4)

Table 5. Association between persistence and variation of general overweight-obesity and abdominal obesity between the ages of 4 and 6, and high blood pressure at age 6

High Blood Pressure						
	OR ^d	95%CI	p-value	OR ^e	95%CI	p-value
Body Mass Index (N=1632) ^a						
Persistent non-excess weight	1 (ref)			1 (ref)		
Persistent excess weight	2.77	(1.37 to 5.58)	0.006	2.54	(1.27 to 5.07)	0.010
Incident	2.55	(1.46 to 4.44)	0.002	2.49	(1.50 to 4.13)	0.001
Remission	1.32	(0.54 to 3.24)	0.523	1.31	(0.50 to 3.44)	0.562
Waist Circumference (N=1612) ^b						
Persistent non-abdominal obesity	1 (ref)			1 (ref)		
Persistent abdominal obesity	4.13	(1.58 to 10.75)	0.005	3.42	(1.38 to 8.49)	0.009
Incident	3.22	(1.19 to 8.68)	0.022	2.81	(0.98 to 8.02)	0.052
Remission	1.67	(0.41 to 6.70)	0.456	1.57	(0.41 to 5.94)	0.491
Waist/height ratio (N=1612) ^c						
Persistent non-abdominal obesity	1 (ref)			1 (ref)		
Persistent abdominal obesity	3.80	(1.36 to 10.59)	0.012	3.17	(1.12 to 8.91)	0.030
Incident	3.27	(1.50 to 7.10)	0.004	3.01	(1.44 to 6.28)	0.004
Remission	0.83	(0.23 to 3.03)	0.782	0.76	(0.21 to 2.69)	0.668

^a Excess weight (Body Mass Index) >+1 standard deviation (SD) according to WHO-2007 reference tables

^b Abdominal Obesity: ≥90th percentile according to reference tables by Fernández et al. [17]

^c Abdominal Obesity: ≥90th percentile of waist(cm)/height(cm) ratio

^d Odds ratios: estimated using unadjusted logistic regression

^e Odds ratios: estimated using logistic regression adjusted for sex, age, maternal educational level, family purchasing power, hours of TV screen or video games per week, hours of physical activity per week, breastfeeding, and baseline blood pressure (age 4)

Supplementary Table 1. Association between general overweight-obesity and abdominal obesity at 4 years of age adjusted simultaneously, and systolic and diastolic blood pressure at 6 years of age.

(N=1788)	Systolic blood pressure (mmHg)			Diastolic blood pressure (mmHg)		
	Coef. β^c	95%CI	p-value	Coef. β^c	95%CI	p-value
Body Mass Index						
Non-excess weight	(ref)			(ref)		
Overweight ^a	1.49	(0.39 to 2.59)	0.009	1.21	(0.02 to 2.39)	0.045
Obesity ^a	2.96	(1.13 to 4.79)	0.002	2.32	(-0.25 to 4.89)	0.076
Waist circumference						
Non-obese	(ref)			(ref)		
Obese ^b	2.75	(1.40 to 4.10)	<0.001	1.14	(-0.75 to 3.04)	0.228

^a Overweight (Body Mass Index): >+1 standard deviation (SD); obesity: >+2 standard deviation (SD) according to WHO-2007 reference tables

^b Abdominal Obesity: $\geq 90^{\text{th}}$ percentile according to reference tables by Fernández et al. [17]

^c β coefficients: estimated using linear regression, adjusted for sex, age, maternal educational level, family purchasing power, hours of TV screen or video games per week, hours of physical activity per week, breastfeeding, and baseline blood pressure (age 4). Body Mass Index and waist circumference are introduced simultaneously

Supplementary Table 2. Association between general obesity and abdominal obesity at 4 years and systolic and diastolic blood pressure at age 6

(N=1788)	Systolic blood pressure (mmHg)			Diastolic blood pressure (mmHg)		
	Coef. β^c	95%CI	p-value	Coef. β^c	95%CI	p-value
Body Mass Index (BMI) and waist circumference (WC)						
Non-BMI or WC-obesity	(ref)			(ref)		
BMI-obesity ^a	3.62	(0.07 to 7.17)	0.046	1.38	(-1.70 to 4.48)	0.367
WC-obesity ^b	3.98	(1.98 to 5.97)	<0.001	1.40	(-1.21 to 4.03)	0.283
BMI & WC-obesity	5.02	(2.98 to 7.07)	<0.001	3.60	(1.00 to 6.21)	0.017

^a Obesity (Body Mass Index) >+2 standard deviation (SD) according to WHO-2007 reference tables

^b Abdominal Obesity: $\geq 90^{\text{th}}$ percentile according to reference tables by Fernández et al. [17]

^c β coefficients: estimated using linear regression, adjusted for sex, age, maternal educational level, family purchasing power, hours of TV screen or video games per week, hours of physical activity per week, breastfeeding, and baseline blood pressure (age 4)