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Toxic stress, health and nutrition among Brazilian children in shelters



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Abstract

Background: Living in a shelter is an adverse experience that generates toxic stress. This situation can cause the dysregulation of the hypothalamic-pituitary-adrenal axis and exert a negative impact on health. The aim of the present study was to determine the association between toxic stress and social, clinical and nutritional characteristics in children at welfare institutions in a city of northeastern of Brazil.

Methods: An analytical, cross-sectional study was conducted with male and female children up to 60 months of age who live in shelters. Hair cortisol was used for the assessment of stress (immunoassay). The anthropometric data collected were height for age, body mass index for age, arm circumference for age, and head circumference for age (expressed in z-scores). We also evaluated food intake using markers proposed by the Brazilian Dietary and Nutritional Vigilance Surveillance System as well as the occurrence of dental caries and anemia.

Results: Sixty-three children one to 60 months of age participated in the present study. Asthma was the most frequent disease (11.1%). The prevalence of short stature, anemia and dental caries in the sample was 22.2, 22.2 and 9.4%, respectively. Cortisol levels ranged from 0.93 pg/mg to 391.29 pg/mg (median: 6.17 pg/mg). Higher cortisol levels were found in children with illnesses (p = 0.012) and those who had been hospitalized after being admitted to the institutions (p = 0.001).

Conclusions: The majority of children had unhealthy eating behavior. The cortisol concentrations found in the present study were suggestive of dysregulation of the hypothalamic-pituitary-adrenal axis. Hypercortisolism was associated with illness and hospitalization.

Keywords: Anemia, Child, Hair cortisol, Institutionalized, Stressful life events

Background

Although a protection measure, being sheltered in an institution involves the breaking of family ties and affective deprivation and is therefore considered a *stressful life* event or adverse experience [1, 2]. Living in a shelter can lead to emotional problems and some children arrive at the shelter with health problems [1, 3]. Results of the Bucharest Early Intervention Project in Romania revealed

that children in shelters had more symptoms, disorders and disabilities than children who had never be exposed to institutionalization [3, 4].

Intense or prolonged stress is denominated *toxic stress*. When this occurs early in life, it can exert a negative impact on brain structure due to the toxicity of the substances released, such as cortisol, which compromises mental health and can cause irreversible harm to neurodevelopment [5–9]. These adverse experiences in childhood are associated with learning difficulties and behavioral problems, which may be related to sex, genetic factors and extrinsic factors [6, 8, 10]. A positive

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association has been found between stress in early life and aggressivity in childhood [11]. Studies have also shown an association between stress and obesity [12, 13].

Stress can be assessed by the concentration of cortisol in different biological matrices. Cortisol measured in hair is a biomarker of chronic or toxic stress and its concentration corresponds to the activation of the stress response system — the hypothalamic-pituitary-adrenal (HPA) axis [14], which is the main mediator of the consequences of stress to the structure of the brain [15]. A significant increase in hair cortisol has been found in children after beginning school [16], in victims of abuse [17] and in situations of food insecurity [18].

Methods

As children in shelters are exposed to toxic stress, the aim of the present study was to evaluate the association between this stress and social, clinical, and nutritional characteristics in children living in shelters in the city of Fortaleza, Brazil.

An analytical, cross-sectional study with a quantitative approach was conducted involving children up to 60 months of age living in welfare institutions in a city of the northeastern of Brazil, between May and August 2017. All 78 children in this age group sheltered in the city (according to data from the National Registry of Sheltered Children) were selected for the study. The exclusion criteria were a lack of information on the date of birth, physical disabilities and genetic syndromes.

The data were collected with the aid of structured questionnaires through interviews with the staff at the shelters and transcripts of individual files. Data collection was performed by a healthcare team composed of a nutritionist, nurse, dentist and undergraduate student of nutrition. The following variables were collected: sex, age, reason for being sheltered, length of time at the institution, history of being sheltered, dental caries, illnesses, hospitalizations after admission to the shelter, the use of corticoids, anemia, hair cortisol, weight, height, arm circumference (for individuals older than 2 years of age), head circumference (for individuals younger than 2 years of age), markers of food intake, siblings at the same shelter, contact with the mother, maternal mental disease and drug use by the mother.

The reasons for being sheltered at the institution were classified according to the categorization of the Management Department of the Social Assistance System of the Ministry of Social Development: intra-family violence (physical or psychological), sexual abuse, sexual exploitation, negligence or abandonment [19]. For the present study, all forms of violence (physical, psychological and sexual) were grouped into a single category. The main reason that led to being placed into a shelter was used for the purposes of analysis.

The dental examination was performed by a dentist and involved the cleaning of the teeth, the use of a tongue depressor and a light source, with the assistance of another researcher. The children were examined in the supine position. Dental caries experience was recorded using the dmft index, which indicates decayed teeth (d), missing teeth with an indication of extraction (m) and filled teeth (f).

The anthropometric data were those recommended by the Brazilian Dietary and Nutritional Vigilance System [20]. All children were weighed prior to the second meal of the day (just before 9 AM). Due to the routine of the institutions, most children received meals every 3 hours, which impeded weighing after fasting. The anthropometric measures were determined twice by the same evaluator and the mean was recorded. When the two weight readings differed by more than 100 g and height differed by more than 1 cm, the measurements were repeated.

Children younger than 2 years of age were weighed unclothed on a digital pediatric scale (Mobile Baby ELP-25BB, Balmak) with a capacity of 25 kg and precision of 1 g. Children older than 2 years of age were weighed in undergarments on a digital scale (New BK-F/FA, Balmak) with a capacity of 150 kg and precision of 100 g (platform: 40×40 cm; non-slip surface).

The stature of children younger than 2 years of age was determined with the child lying on a flat surface and was measured with the aid of 100-cm wooden pediatric ruler with a precision of 1 cm. The stature of children older than 2 years of age was measured with the child in the standing position with the aid of a retractable aluminum ruled installed in the column of the scale with a capacity of 2 m and precision of 0.5 cm. Arm and head circumference were measured using a plastic, non-flexible metric tape graduated in millimeters.

The indices used for the anthropometric evaluation were stature for age (S/A), body mass index for age (BMI/A), head circumference for age (HC/A) and arm circumference for age (AC/A) according to sex and expressed in z-scores. The Anthro v3. 2.2 program [21] was used for the classification of nutritional status using the cutoff points recommended by the World Health Organization (WHO) and adopted by the Dietary and Nutritional Vigilance System: for the BMI/A index, a zscore < - 2 indicates undernourished/underweight, z $score \ge -2$ and $\le +1$ is the ideal range, z-score > +1and \leq + 2 indicates a risk of overweight and z-score > + 2 indicates excess weight (overweight and obesity); for the S/A index, a z-score – 2 indicates short stature and z-score ≥ - 2 indicates adequate stature. WHO values were used for the classification of birthweight: < 2500 g = low weight; 2500 to 2999 g = insufficientweight; 3000 to 3999 g = adequate weight; and > 4000g = macrosomia [22].

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The "Food Intake Markers" proposed by the Dietary and Nutritional Vigilance System was used for the evaluation of food intake. The classification and analysis of the food intake data were based on the "Guidelines for the Assessment of Food Intake Markers in Primary Care" of the Brazilian Health Ministry, which recommends the division by age group: up to 5 months and 29 days; 6 to 23 months and 29 days; and \geq 2 years [23]. The markers enable defining healthy and unhealthy eating behavior. The consumption of fruit, vegetables and beans is considered a marker of healthy eating, whereas the consumption of ultra-processed foods (processed meats, sweetened beverages, instant pasta, crackers, sweets and cookies with filling) is considered a marker of unhealthy eating. At the institutions, the employees responsible for the direct care of the children worked in shifts; thus, the food consumption form was used to evaluate the frequency of consumption.

Anemia was determined by the concentration of capillary hemoglobin (Hgb) using the *HemoCue* portable photometer for direct reading in $10\,\mu l$ of blood obtained using disposable lancets. In children younger than 1 year of age, the drop of blood was collected from the side of the heel with the puncture performed vertically after disinfection with 70% alcohol and drying of the skin surface with cotton. In children older than 1 year of age, the puncture was performed on the side of the tip of one of the fingers after washing with soap and water, disinfection with 70% alcohol and complete drying. Hgb < 9.5 g/dL in children less than 6 months of age and Hgb < 11 g/dL in children six or more months of age were considered indicative of anemia [24].

Stress was evaluated by the determination of hair cortisol following the Catherine Herba protocol and the criteria of the Society of Hair Testing [25]. Strands of hair measuring at least 3 cm were collected from the posterior vertex of the scalp, with 20 mg of strands used from each child. Cortisol extraction and analysis were performed at the Endocrine and Metabolism Physiology Lab of the Institute of Biological Sciences of the State University of Ceará following the method adopted by other authors [26, 27]. The analysis was performed using an enzyme-linked immunosorbent assay (ELISA) kit (Quality Control Sheet – Cayman Chemical) following the manufacturer's instructions. Cortisol concentrations were expressed as pg/mg.

The statistical analysis was performed with the aid of the SPSS program, version 22.0 (SPSS Inc., Chicago, IL, USA). The Kolmogorov-Smirnov test was used to determine the distribution (normal or non-normal) of the continuous variables. Data with normal distribution were expressed as mean and standard deviation values and those with non-normal distribution were expressed as median and interquartile range. The comparison of means was performed using the Student's t-test (two means) and ANOVA (two or more means). Either the chi-square test (χ^2) or Fisher's exact test was used for the comparison of frequencies. Analysis of covariance (ANCOVA) was used for the analysis of factors associated with mean concentrations of hair cortisol (dependent variable). The independent variables (associated factors) selected for the multivariate analysis were those with a p-value < 0.20 in the bivariate analysis. Variables with a p-value < 0.05 in the final model were considered significantly associated with the dependent variable.

This study was conducted in accordance with the ethical principles stipulated in the Declaration of Helsinki and the norms for research involving human subjects stipulated in Resolution 466/12 of the Brazilian National Board of Health. The study received approval from the Human Research Ethics Committee of the Federal University of Pernambuco (protocol number: 2.019.560, 18 April 2017; certificate number: 64680116.4.00005208). The statement of informed consent and letter of authorization were signed by the coordinating judge of the Children's District Court of Fortaleza.

Results

One shelter declined to participate in the study and three children were excluded (one with microencephaly and two with cerebral palsy). Thus, the data on 63 children from one to 60 months of age were analyzed. However, only 49 children were included in the study of hair cortisol, as the other 11 children had an insufficient quantity of hair or strands less than 3 cm. Girls accounted for 52.4% of the samples and age ranged from one to 60 months (mean: 30.0 ± 16.4 months). The main reasons for being admitted to a shelter were negligence (47.6%) and abandonment (36.5%). Ten children (15.9%) had been living at the institutions for more than 2 years. Age of the children at the time of being admitted to the shelter ranged from 0 to 56 months; four (6.6%) were newborns and 35 (55.5%) were children between one and 24 months of age. A total of 36% of the children had previously been at another shelter. Only 27% of the children had contact with their mothers and 44.4% had siblings at the same institution. Seven (11.1%) of the mothers had a mental disease. A total of 50.8% of mothers were users of psychoactive drugs. However, this figure could be higher, as 20 charts (31.7%) had no information regarding drug use by the mother.

Regarding health status, 27% of the children had some disease. The use of oral or inhaled corticoids (alone or combined) was found in 46% of the sample. The prevalence of dental caries was 9.43%, with a dmft index of 0.28 in the population. Dental caries was not associated with any of the other variables studied. Three children

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refused to undergo the dental examination. Seven had no teeth and the age of these children ranged from one to 11 months. Anemia was associated with the duration of time in the shelter; the frequency was higher among children with up to 12 months at the shelter (p = 0.024). The data on the health status of the children are displayed in Table 1.

The frequency of short stature was 22.2%. Thirteen children (20.6%) were at risk of excess weight and eight (12.7%) had excess weight according to the BMI/A index. AC/A was considered normal in all 33 children evaluated. HC/A was below the -2 z-score in seven (26.9%) among the 26 children evaluated. Regarding food intake, 38 (61.3%) of the children had unhealthy eating behavior.

Cortisol concentration exhibited non-normal distribution and ranged from 0.93 to 391.29 pg/mg (median: 6.17 pg/mg; interquartile range: 7.26). For the statistical analyses, the highest value was excluded because it was very distant from the median. Thus, in the analysis performed with 48 children, cortisol ranged from 0.93 to 54.78 pg/mg. The quartile values were Q1 = 3.2, Q2 = 6.0

Table 1 Health conditions of children residing at welfare shelters in the city of Fortaleza, Brazil, 2017

Health conditions	N	%
Hospitalization after admission to shelte	er (n = 63)	
Yes	15	23. 8
No	48	76.2
Health status $(n = 63)$		
Healthy	46	73.0
III	17	27.0
Asthma	7	11.1
Cow's milk protein allergy	2	3.2
Lactose intolerance	2	3.2
Chronic intestinal constipation	1	1.6
Wheezing infant syndrome	1	1.6
OS ASD ^a with left-to-right flow	1	1.6
Congenital syphilis	1	1.6
Hydrocele	1	1.6
Gastroesophageal reflux disease	1	1.6
Dental caries b ($n = 53$)		
Yes	5	9.43
No	48	90.57
Anemia (n = 62)		
Yes	14	22.6
No	48	77.4

^aOstium secundum atrial septum defect (acyanotic congenital heart defect) without hemodynamic repercussions in this child; ^bnly children with dentition included in calculation;

Source: Author

and Q3 = 10.4. In the ANCOVA controlled for the use of corticoids, no statistically significant differences were found among age groups (Table 2).

However, positive associations were found between hypercortisolism and the occurrence of illness and hospitalization (Table 3).

Discussion

The present results enable us to infer that stress had a negative impact on the health of the sample studied. The most prevalent disease was asthma, which is the main cause of hospitalization among Brazilian children [28] and is positively associated with stress [9, 29]. Contrasting the findings of this study, KAMPS et al. (2014) [30] conducted a case-control study evaluating the effect of asthma and its treatment on the HPA axis and found that the hair cortisol levels of the children with asthma were significantly lower than those of the healthy controls. However, children living in shelters are exposed to numerous stressors related to their past situations as well as those inherent to institutionalization.

Toxic shock may be the trigger for the manifestation of genetically programmed diseases [31]. As hospitalization is an adverse experience [32], becoming ill can initiate a cycle of stressful experiences in children living in a situation of welfare institutionalization. Few children had protective factors, such as contact with their mothers and sibling at the same institution. Not all institutions have psychologists on staff and none had the support of religious groups on a permanent basis.

The illnesses found in the population studied are highly prevalent in this age group. Therefore, disease prevention and control and a reduction in the stay at a shelter could avoid or reduce stress and its negative consequences, which impede full, healthy development. The placement of these children in a foster family (modality that offers care in a family environment and ensures individualized attention) is an alternative to institutionalization, but, unfortunately, is a rare occurrence in Brazil, where the

Table 2 Concentration of hair cortisol per age group of children residing at welfare shelters, Brazil, 2017

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Age group	N Cortisol (pg/mg)			p-	Adjusted	
		Mean	Standard deviation	value	<i>p</i> -value [†]	
0	8	15.82	± 13.86	0.080	0.100	
12 — 24 months	11	9.48	± 5.83			
24 — 36 months	13	11.74	± 15.36			
36 4 8 months	10	3.49	± 2.05			
48 — 60 months	6	3.55	± 1.51			
Total	48	9.16	± 10.83			

[†] p-value adjusted for use of corticoids using ANCOVA. Cortisol as dependent variable Source: Author

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Table 3 Analysis of covariance of hair cortisol and characteristics of children residing at welfare shelters, Brazil, 2017

VARIABLES	Categories	N	Cortisol (pg/mg)		р-	Adjusted
			Mean	Standard deviation	value	<i>p</i> -value [†]
Reason for admission to shelter	Abandonment	18	10.10	10.77	0.707	
	Negligence	21	9.51	12.62		
	Violence	9	6.46	5.87		
Stay at shelter	≤ 12 months	33	10.38	12.52	0.251	
	> 12 months	15	6.47	4.96		
Past history of shelter	Yes	35	9.24	9.58	0.936	
	No	13	8.95	14.12		
Age upon admission	≤12 months	20	10.57	9.81	0.361	
	> 12 months	24	7.48	12.02		
Siblings in shelter	Yes	24	8.64	11.45	0.742	
	No	24	9.68	10.40		
Contact with mother	Yes	12	8.85	10.17	0.910	
	No	36	9.26	11.18		
Illness	Yes	11	15.83	18.40	0.155	0.012*
	No	37	7.18	6.45		
Hospitalization	Yes	11	19.16	18.09	0.039*	0.001*
	No	37	6.18	4.76		
Caries	Yes	2	4.72	1.82	0.559	
	No	46	9.35	11.03		
Anemia	Yes	10	9.92	13.34	0.831	
	No	37	9.08	10.39		
Eating behavior	Healthy	19	8.63	12.20	0.788	
	Unhealthy	29	9.51	10.05		
Stature for age	Low	10	10.36	13.45	0.698	
	Adequate	38	8.84	10.23		

[†] p-value adjusted for use of corticoids using ANCOVA. Cortisol as dependent variable

Source: Author

shelter remains the main modality. Foster care is ensured in Article 34 of the Child and Adolescent Statute and, according to this law, should be a priority measure for the protection of these children [33].

The Bucharest Early Intervention Project in Romania investigated the impact of psychosocial deprivation associated with institutional child care in early life and the findings suggest that early psychosocial negligence can result in cognitive deficits, whereas children under the care of a foster family develop better [34]. Therefore, the institutionalization of children can lead to diseases and neurodevelopmental disorders that can exert a long-term impact on the lives of these children and even influence future generations.

The cortisol data did not exhibit normal distribution, which is similar to findings described in previous studies [14, 35, 36]. Some children had a concentration lower than that found in the literature and, in the analysis of

quartiles, some had concentrations lower than the 1st quartile (3.2), suggesting hypocortisolism. As toxic stress leads to the dysregulation of the HPA axis, it can cause either hypercortisolism or hypocortisolism [37, 38]. Although the mechanism by which hypocortisolism occurs in toxic stress has not yet been clarified, hypotheses suggest that the hyperresponsivity of the HPA makes it hyporesponsive or there may be hypersensitivity of the target tissue [9, 10, 39]. Moreover, exposure to excessive maternal stress in the prenatal period has been associated with a reduction in cortisol in the hair of newborns [40].

There is evidence of hypocortisolism related to toxic stress in children deprived of their mother's care, individuals having suffered trauma [10] and individuals with anxiety disorders [41]. Differences in the activation response of the HPA axis and, consequently, hair cortisol levels may be explained by individual differences in

^{*}Only significant variables ($p \le 0.20$) included in analysis

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perceptions and psychological responses to stress due to diverse factors, such as a genetic predisposition and past traumas [6, 31].

The results regarding nutritional status are similar to those described in previous studies involving institutionalized children independently of the indicator used for the evaluation [42–44]. Among the nutritional disorders found, short stature is a common occurrence and was three times more frequent in the present investigation compared to the findings of a large population-based survey conducted in Brazil with non-institutionalized children [45]. This reveals that factors related to living in a shelter may exert an effect on aspects that are indispensable to normal growth, such as nutrition and health care.

Although the prevalence has been declining, short stature remains one of the major nutritional disorders in Brazil [46]. The illnesses found in the present study do not have an aggressive impact on nutritional status. However, we found unhealthy eating behavior in the majority of the sample, which reveals the low quality of the menu offered.

Approximately half of the children made use of corticoids. The prolonged use of these medications can have side effects, such as linear growth deficit [47]. The use of corticosteroids is associated with a reduction in the functioning of executive cognition as well as a greater likelihood of mood disorders and anxiety in the general adult population [48]. Therefore, the prescription of such medications to populations exposed to toxic stress should be evaluated carefully.

A positive association was found between anemia and a shorter length of stay in the shelter. This finding reveals that anemia may be minimized or corrected at such institutions, as hemoglobin can take months to normalize. Although food intake in the majority of children at the institutions was not considered adequate, it may have been better than that prior to institutionalization, enabling an improvement in hemoglobin levels.

This is the first study on toxic stress in Brazilian children who live in shelters to evaluate hair cortisol as a marker of stress. The age group of the present sample is the same as the majority of children who live in shelters in the city of Fortaleza. Although some data were secondary, this information was collected from official documents that are valid for the entire country, such as the declaration of a live birth.

A control group of non-institutionalized and a larger sample size may have brought other significant associations to light. As chronic stress can exert a considerable biopsychosocial impact, adequately addressing the physical and mental health of institutionalized children is of the utmost importance.

Conclusions

Some of the children in the present study had cortisol concentrations suggestive of the dysregulation of the HPA axis. The occurrence of illness and a history of hospitalization were associated with higher concentrations of hair cortisol. High frequencies of short stature and anemia were found in the population studied and the analysis of dietary habits revealed that the majority of children had unhealthy eating behavior. The present results demonstrate an urgent need to improve the nutrition and health (physical and mental) of the children studied.

Abbreviations

HPA: Hypothalamic-pituitary-adrenal; BMI/A: Mass index for age; HC/A: Head circumference for age; AC/A: Arm circumference for age; S/A: Stature for age; WHO: World Health Organization; Hgb: Hemoglobin; ELISA: Enzyme-linked immunosorbent assay; ANCOVA: Analysis of covariance; dmf-t: decayed teeth (d), missing teeth with an indication of extraction (m) and filled teeth (f)

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Authors' contributions

ACS, responsible for the design of the research project; data collection, analysis and interpretation; and writing of the article. AJML, responsible for the design of the research project; analysis and interpretation; and writing of the article. PCC, responsible for the analysis and interpretation; and writing of the article. ACO, responsible for the data collection, analysis and interpretation; and writing of the article. KAO, responsible for the data collection, analysis and interpretation; and writing of the article. PICL, responsible for the design of the research project; analysis and interpretation; and writing of the article. The author(s) read and approved the final manuscript.

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Availability of data and materials

The datasets used and analyzed in this study are available from the corresponding author upon request.

Declarations

Ethics approval and consent to participate

This study was conducted in accordance with the ethical principles stipulated in the Declaration of Helsinki and the norms for research involving human subjects stipulated in Resolution 466/12 of the Brazilian National Board of Health. This study received approval from the Human Research Ethics Committee of the Federal University of Pernambuco (protocol number: 2.019.560, 18 April 2017; certificate number: 64680116.4.00005208). The statement of informed consent and letter of authorization were signed by the coordinating judge of the Children's District Court of Fortaleza.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

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References

- Álvares AM, Lobato GR. Um estudo exploratório da incidência de sintomas depressivos em crianças e adolescentes em acolhimento institucional. Temas Psicol. 2013;21(1):151–64. https://doi.org/10.9788/TP2013.1-11.
- Berens AC, Nelson CA. The science of early adversity: is there a role for large institutions in the care of vulnerable children? Lancet. 2015;11(15):1–11. https://doi.org/10.1016/S0140-6736(14)61131-4.
- Humphreys KL, Gleason MM, Drury SS, Miron D, Nelson CA, Fox NA, Zeanah CH. Effects of institutional rearing and foster care on psychopathology at age 12 years in Romania: follow-up of an open, randomised controlled trial. Lancet Psychiatry. 2015;2(7):625–34. https://doi.org/10.1016/S2215-0366(1 5)00095-4
- Zeanah CH, Egger HL, Smyke AT, Nelson CA, Fox NA, Marshall PJ, Guthrie D. Institutional rearing and psychiatric disorders in Romanian preschool children. Am J Psychiatry. 2009;166(7):777–85. https://doi.org/10.1176/appi.a jp.2009.08091438.
- Sociedade Brasileira de Pediatria. Avaliação nutricional da criança e do adolescente. Manual de Orientação – Departamento de Nutrologia; 2009. p. 116. [acesso em: 24 Out 2017]. Disponível em: http://www.sbp.com.br/fileadmin/user_upload/pdfs/MANUAL-AVAL-NUTR2009.pdf.
- Franke HA. Toxic stress: effects, prevention and treatment. Children. 2014;1: 390–402. https://doi.org/10.3390/children1030390.
- American Academy of Pediatrics (EUA). Toxic stress toolkit for primary care providers caring for young children. 2016. Disponível em: http://health.ri.gov/publications/toolkits/ToxicStress.pdf. Acesso em: 14 Dez 2018.
- Novais A, Monteiro S, Roque S, Correia-Neves M, Sousa N. How age, sex and genotype shape the stress response. NeurobiolStress. 2016;6:44–56. https:// doi.org/10.1016/j.ynstr.2016.11.004.
- Mendes MA, Cairo S. Estresse e asma na infância e adolescência. Pulmão. 2013;22(3):43–7 Disponível em: http://www.sopterj.com.br/wp-content/ themes/_sopterj_redesign_2017/_revista/2013/n_03/10.pdf>. Acesso em: 2 Dez 2017.
- Lupien SJ, Juster RP, Raymond C, Marin MF. The effects of chronic stress on the human brain: from neurotoxicity, to vulnerability, to opportunity. Front Neuroendocrinol. 2018;41:95–101. https://doi.org/10.1016/j.yfrne.2018.02.001.
- Winiarski DA, Engel ML, Karnik NS, Brennan PA. Early life stress and childhood aggression: mediating and moderating effects of child callousness and stress reactivity. Child Psychiatry Hum Dev. 2018;49:730–9. https://doi.org/10.1007/s10578-018-0785-9.
- Papafotiou C, Christaki E, Van Den Akker EL, Wester VL, Apostolakou F, Papassotiriou I, Chrousos GP, Pervanidou P. Hair cortisol concentrations exhibit a positive association with salivary cortisol profiles and are increased in obese prepubertal girls. Stress. 2017;20(2):1–20. https://doi.org/10.1080/1 0253890.2017.1303830.
- Estrela DC, Mendes BO. Associação entre obesidade e estresse crônico: uma revisão sobre aspectos comportamentais, bioquímicos e hematológicos. Multi Sci J. 2017;1(9):41–50. https://doi.org/10.33837/msj.v1i9.609.
- Vanaelst B, Huybrechts I, Bammann K, Michels N, De Vriendt T, Vyncke K, Sioen I, Iacoviello L, Günther K, Molnar D, Lissner L, Rivet N, Rausl JS, De Henauw S. Intercorrelations between serum, salivary, and hair cortisol and child-reported estimates of stress in elementary school girls. Psychophysiology, ago. 2012a;49(8):1072–81. https://doi.org/10.1111/j.1469-8986.2012.01396.x.
- Russo SJ, Murrough JSW, Han M, Charney DS, Nestler EJ. Neurobiology of resilience. Nat Neurosci. 2012;15(11):1475–84. Disponível em: https://doi.org/10.1038/nn.3234. Acesso em: 05 Fev 2017. https://doi.org/10.1038/nn.3234.
- Rickmeyer C, Lebiger-Vogel J, Leuzinger-Bohleber M. Transition to kindergarten: negative associations between the emotional availability in mother-child relationships and elevated cortisol levels in children with an immigrant background. Front Psychol 2017, 8. https://doi.org/10.1590/1413-81232017222.26252015
- Daruy FL. Aspectos imunoendocrinológicos em adolescentes vítimas de maus-tratos na infância. 111 f. Porto Alegre: Tese(Doutorado em Psicologia) – Pontifícia Universidade Católica do Rio Grande do Sul; 2015.
- Ling J, Robbins LB, Xu D. Food security status and hair cortisol among lowincome mother- child dyads. West J Nurs Res. 2019;41(12):1813–28. https:// doi.org/10.1177/0193945919867112.

- Brasil Ministério do Desenvolvimento Social e Combate à Fome. Manual de instruções para o registro das informações especificadas na Resolução N°04/ 2011 alterada pela Resolução N° 20/2013 da Comissão Intergestores Tripartite − CIT. Brasília: DF; 2014. p. 23. Disponível em: https://aplicacoes.mds.gov.br/sagi/atendimento/doc/Manual de InstrucoesCREAS.pdf>. Acesso em: 22 Jan 2017
- Brasil Ministério da Saúde. Orientações para a coleta e análise de dados antropométricos em serviços de saúde: Norma Técnica do Sistema de Vigilância Alimentar e Nutricional – SISVAN. Série G. Brasília: DF; 2011. Disponível em: http://189.28.128.100/nutricao/docs/geral/orientacoes_coleta_analise_dados_antropometricos.pdf>. Acesso em: 10 Maio 2016
- 21. World Health Organization. Who Anthro for personal computers. In: Software for assessing growth and development of the world's children. Version 2; 2006. [acesso em: 20 Out 2015]. Disponível em: http://www.who.int/childgrowth/software/en/.
- Organización Mundial de la Salud. El estado físico: uso e interpretación de la antropometría. Ginebra: OMS – Serie de Informes Técnicos – 854; 1995. Disponível em: https://www.who.int/childgrowth/publications/physical_status/es/. Acesso em: 14 Maio 2018
- 23. Brasil Ministério da Saúde, Secretaria de Atenção à Saúde, Departamento de Atenção Básica. Orientações para avaliação de marcadores de consumo alimentar na atenção básica [recurso eletrônico]. Brasília: Ministério da Saúde, Secretaria de Atenção à Saúde, Departamento de Atenção Básica, Ministério da Saúde; 2015. p. 33. Disponível em: http://bvsms.saude.gov.br/bvs/publicacoes/marcadores_consumo_alimentar_atencao_basica.pdf> Acesso em: 02 de Junho de 2018
- World Health Organization. The clinical use of blood in medicine, obstetrics, paediatrics, surgery & anaesthesia, trauma & aburns, 2001. [acesso em: 21 Out 2015]. Disponível em: http://apps.who.int/iris/bitstream/10665/423
 97/1/a72894.pdf>.
- Cooper GA, Kronstrand R, Kintz P. Society of Hair Testing guidelines for drug testing in hair. Forensic Sci Int. 2012;218(1–3):20–4. https://doi.org/10.1016/j. forsciint.2011.10.024.
- Dowlati Y, Herrmann N, Swardfager W, Thomson S, Oh PI, Van Uum S, Koren G, Lanctôt KL. Relationship between hair cortisol concentrations and depressive symptoms in patients with coronary artery disease. Neuropsychiatr Dis Treat. 2010;6(1):393–400. https://doi.org/10.2147/NDT.S1 0353
- Noppe G, Van Rossum EF, Koper JW, Manenschijn L, Bruining GJ, De Rijke YB, Van Den Akker EL. Validation and reference ranges of hair cortisol measurement in healthy children. Hormone Res Paediatr. 2014;82(2):97–102. https://doi.org/10.1159/000362519.
- Pedraza DF, Silva FA, Melo NLS, Araujo EMN, Sousa CPC. Estado nutricional e hábitos alimentares de escolares de Campina Grande, Paraíba, Brasil. Cien Saude Colet. 2017;22(2):469–77. https://doi.org/10.1590/1413-81232017222.2 6252015.
- Mendes MA, Sant'anna CC, March MFBP. O estresse em crianças e adolescentes com asma. J Hum Growth Dev. 2013;23(1):80–6. https://doi. org/10.7322/jhqd.50395.
- Kamps AW, Molenmaker M, Kemperman R, Van Der Veen BS, Bocca G, Veeger NJ. Children with asthma have significantly lower long-term cortisol levels in their scalp hair than healthy children. Acta Paediatr. 2014;103(9): 957–61. https://doi.org/10.1111/apa.12685.
- Lipp MEN. Teoria de temas de vida do stress recorrente e crônico. Boletim Academia Paulista de Psicologia. 2006;3:82–93 Acessado: 9 de set de 2018. Disponível em: http://www.redalyc.org/articulo.oa?id=94626311>. Acesso em: 02 Dez. 2017.
- 32. Santos PM, Silva LF, Depianti JRB, Cursino EG, Ribeiro CA. Os cuidados de enfermagem na percepção da criança hospitalizada. Revista Brasileira de Enfermagem. 2016;69(4):646–53. https://doi.org/10.1590/0034-7167.20166904
- Brasil. Lei N° 8.069, de 13 de julho de 1990. Dispões sobre o Estatuto da Criança e do Adolescente. Brasília: Legislação.; 1990. Disponível em: http://www.planalto.gov.br/ccivil_03/Leis/L8069.htm. Acesso em: 24 Maio 2018
- Stamoulis C, Vanderwert RE, Zeanah CH, Fox NA, Nelson CA. Early
 psychosocial neglect adversely impacts developmental trajectories of brain
 oscillations and their interactions. J Cogn Neurosci. 2015;27(12):2512–28.
 https://doi.org/10.1162/jocna00877.
- Karlén J, Frostell A, Theodorsson E, Faresjö T, Ludvigsson J. Maternal influence on child HPA Axis: a prospective study of cortisol levels in hair. Pediatrics. 2013;132(5):1333–e1340. https://doi.org/10.1542/peds.2013-1178.

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- Manenschijn L, Van Kruysbergen RG, De Jong FH, Koper JW, Van Rossum EF. Shift work at young age is associated with elevated long-term cortisol levels and body mass index. Clin Endocrinol Metab. 2011;96(11):E1862–5. https://doi.org/10.1210/jc.2011-1551.
- Slopen N, Mclaughlin KA, Shonkoff JP. Interventions to improve cortisol regulation in children: a systematic review. Pediatrics. 2014;133(2):321–6. https://doi.org/10.1542/peds.2013-1632.
- 38. Vanaelst B, Vriendt T, Huybrechts I, Rinaldi S, Henauw S. Epidemiological approaches to measure childhood stress. Paediatr Perinat Epidemiol. 2012b; 26(3):280–97. https://doi.org/10.1111/j.1365-3016.2012.01258.x.
- Maripuu M, Wikgren M, Karling P, Adolfsson A, Norrback K. Hyper- and hypocortisolism in bipolar disorder - a beneficial influence of lithium on the HPA-axis? J Affect Disord. 2017;213:161–7. https://doi.org/10.1016/j.jad.2017. 02.026
- Van Der Voorn B, Hollanders JJ, Kieviet N, Dolman KM, Rijke YB, Van Rossum EF, Rotteveel J, Honig A, Finken MJ. Maternal stress during pregnancy is associated with decreased cortisol and cortisone levels in neonatal hair. Horm Res Paediatr. 2018. https://doi.org/10.1159/000495007.
- 41. Wester VL, Van Rossum EF. Clinical applications of cortisol measurements in hair. Eur J Endocrinol. 2015;173(4):m1–m10 Disponível em: https://www.ncbi.nlm.nih.gov/pubmed/25924811>. Acesso em: 19 Jun. 2017. https://doi.org/10.1530/EJE-15-0313.
- Kamath SM, Venkatappa KG, Sparshadeep EM. Impact of nutritional status on cognition in institutionalized orphans: a pilot study. J Clin Diagn Res. 2017;11(3):CC01–4. https://doi.org/10.7860/JCDR/2017/22181.9383.
- EL-Kassas GE, Ziade F. The dual burden of malnutrition and associated dietary and lifestyle habits among Lebanese school age children living in orphanages in North Lebanon. J Nutr Metabol 2017;2017: 1–12. https://doi. org/10.1155/2017/4863431
- Silveira AC, Costa SC, Pinho CP, Santana SC. Condição nutricional de crianças admitidas em uma instituição de acolhimento. Rev Bras Promç Saúde. 2016;29(1):5–13. https://doi.org/10.5020/18061230.2016.p5.
- 45. Brasil Ministério da Saúde. Pesquisa Nacional de Demografia e Saúde da Criança e da Mulher – PNDS 2006: dimensões do processo reprodutivo e da saúde da criança. Série G. Brasília: Estatística e Informação em Saúde; 2009. p. 300. [acesso em: 20 jun. de 2015] Disponível em: http://bvsms.saude.gov.br/bvs/publicacoes/pnds_crianca_mulher.pdf
- Gonçalves H, Horta BL, Barros AJ, et al. Infant nutrition and growth: trends and inequalities in four population-based birth cohorts in Pelotas, Brazil, 1982-2015. Int J Epidemiol. 2019;48(Suppl1):i80–8. https://doi.org/10.1093/ ije/dyy233.
- Zhang L, Lasmar LB, CastroRodriguez JA. The impact of asthma and its treatment on growth: an evidence based review. J Pediatr. 2018;95:1–13. https://doi.org/10.1016/j.jped.2018.10.005.
- Savas M, Vinkers CH, Rosmalen JGM, Hartman CA, Wester VL, Van Den Akker ELT, Iyer AM, Mcewen BS, Van Rossum EFC. Systemic and local corticosteroid use is associated with reduced executive cognition, and mood and anxiety disorders. Neuroendocrinology. 2020;110(3–4):282–91. https://doi.org/10.1159/000501617.

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