

CLINICAL RESEARCH

e-ISSN 1643-3750 © Med Sci Monit, 2021; 27: e930214 DOI: 10.12659/MSM.930214

.....

Received Accepted Available online Published	d: 2020.12. d: 2021.02. e: 2021.03. d: 2021.05.	.02 .08 .03 .14	Recognizing Early Regu Pediatric Care: The For	lation Disorders in Healthy Offspring Project				
Authors' Contribution: Study Design A Data Collection B Statistical Analysis C Data Interpretation D Manuscript Preparation E Literature Search F Funds Collection G Corresponding Author: Source of support: Background:		ABCDEFG 1 ABCDEF 2 BDEF 1 BDEF 3 BDEF 4 BCDEF 1 ABCDEF 1,5	Noémi Scheuring Ildikó Danis Eszter Papp Pálma Benedek Tünde Németh Ágnes Gulácsi László Szabó	 Department of Pediatrics, Heim Pál National Pediatric Institute, Budapest, Hungary Institute of Mental Health, Semmelweis University, Budapest, Hungary Department of Otolaryngology, Heim Pál National Pediatric Institute, Budapest, Hungary Department of Pediatrics, Dr. Halász Géza Medical Center, Dabas, Hungary Department of Family Care Methodology, Faculty of Health Sciences, Semmelweis University, Budapest, Hungary 				
		ding Author: e of support:	László Szabó, e-mail: szabo.laszlo.md@gmail.com Funding was provided by Budapest City Council, Committee for Health and Social Politics (Support contract: 127/2010), Hungary; Szundi Foundation for Children with Sleep Disorders, Budapest, Hungary; Foundation for Supporting the Development of Heim Pál Children's Hospital, Budapest, Hungary; and Electrooxygen Ltd., Budapest, Hungary Regulation disorders are already apparent in infancy. The For Healthy Offspring Project was the first Hungarian study aimed at building an effective model for screening and examining the prevalence and complex (medi- cal and psychosocial) background of classic behavior regulation disorders (excessive crying, feeding, and sleep problems) in infancy.					
		ackground:						
	Materia	l/Methods:	Data were collected from families of 0- to 3-year-old children in a pediatric hospital and its neighboring areas					
Results: Conclusions: Keywords: Full-text PDF:		Results: onclusions:	In the questionnaires, medical examinations, and In the questionnaire study about their children's be ing, 16% reported feeding problems, and 10% report ical examinations were also conducted, the prevaler 15.2% for sleep disorders, 10.3% for breastfeeding of were referred to the screening program (n=183) had other children in our study. Regulation disorders we some cases. We developed a complex model to screen for regulation dence (5-15%) of early childhood regulation disorders In order to effectively recognize early regulation disor tional field should be adapted in general Hungarian	whavior (n=1133), 15% of mothers reported excessive cry- ted sleep problems. In a subsample (n=619) in which med- nce of medical diagnoses was 15.0% for excessive crying, difficulties, and 14.8% for feeding disorders. Children who I significantly more behavior regulation disorders than the ere found to be comorbid with other health conditions in tory problems in early childhood. This study adds more in- n problems and other health conditions. The general inci- in other countries is likely similar to that found in Hungary. orders, diagnostic instruments widely used in the interna- pediatric care.				
		Keywords:	Crying • Feeding Behavior • Infant Behavior • Sleep Disorders					
		ll-text PDF:	https://www.medscimonit.com/abstract/index/idArt/930214					
			2 4460 H 2 6 H 2 1	2 52				



Background

Emotional and behavior regulation disorders in infancy and toddlerhood are guite frequent, with an occurrence of 5-20% in a normal population [1-5]. Primary or classic regulation disorders ("the classic triad"), such as excessive and persistent crying and sleep and feeding disorders, are already seen in early infancy and can also be recognized and diagnosed in primary care [1,6-10]. According to the framework of developmental psychopathology [11], in most cases, the cumulative combination of somatic, interactional, and psychosocial environmental risk factors and a lack of significant protective factors leads to problematic behaviors associated with several types of early mental health problems [10]. Similar to other early childhood mental health problems, the background of regulation disorders is also assumed to be influenced by complex mechanisms, whereby the individual physical and psychological characteristics of the parents and the children, their common early history (pregnancy, birth, early care), the actual parentchild interactions, and the developing relationships serve as key proximal mediators of more complex distal factors, such as the sociodemographic situation, family structure, stressful life events, and social support. Naturally, the manifestation of certain problems can depend on other moderating factors, such as the age, sex, and temperament of the child [1,12].

This clinical area is specifically located at the juncture of medicine, psychology, and education, and it therefore requires an interdisciplinary approach and handling. The risk factors that compromise childhood development, parent-child interactions, and family functioning and the protective factors that support resilient development are important issues to be borne in mind in both research and clinical practice [13]. Since most of the relevant literature comes from small-sample studies, continued research is needed with larger samples to further explore the clinical significance of regulation disorders [14]. In the meantime, the findings as a whole highlight the paucity of evidence about this group of infants and the need to prioritize them for research and clinical work [15].

The study, screening, and treatment of early childhood mental health problems have a decades-long history in international practice, but the investigation of such disorders, as well as the related prevention and intervention activities, remains an understudied area in Hungary. To date, only a few national private and public initiatives have expressed interest in developing this area. The For Healthy Offspring Project, initiated by the Heim Pál National Pediatric Institute in Budapest, was the first Hungarian research to establish an effective hospital model for screening and to examine the prevalence of emotional and behavior regulation disorders in early childhood (0-3 years) and the significance of different risks and protective factors behind them. We hypothesized that the prevalence of regulation disorders in Hungary is similar to that in other countries. We also hypothesized that the association between excessive crying and sleep disorders is strong. We developed a complex model to screen for regulatory problems in early childhood.

In this article, our aims were to (1) introduce the model of our screening program and our large-sample hospital research, (2) report the occurrence of major regulation disorders (excessive crying, sleep and feeding problems) in our sample, and (3) report associations between regulation disorders and other examined medical diseases.

We hypothesized that (1) the prevalence of regulation disorders in our Hungarian sample is similar to that of other countries, (2) the associations between excessive crying, sleep and feeding disorders are significant, and (3) these regulation problems may also be moderately associated with other diseases.

Material and Methods

Families of 0- to 3-year-old children with eating or sleeping problems or extreme crying from 3 departments of the Heim Pál National Pediatric Institute in Budapest and neighboring areas were included in this study. Data were collected from July 2010 to June 2011 in a cross-sectional design study.

We obtained information about early childhood regulation disorders from 4 sources, including questionnaires, medical examinations, and individual and small group consultations.

The model of the screening process, the data collection, and the administration of the For Healthy Offspring project are shown in **Figure 1**.

During the research period, we recruited from among all families with children under 3 years of age (n=1855) within 3 departments of the hospital, and 580 families volunteered to participate. This represents a response rate of 31.4%. During the same period, we also collected data (n=584) with the help of health visitor nurses in neighboring areas. The nurses mainly administered the questionnaires to the families with whom they visited spontaneously during their work or who visited them during their consulting hours. Thus, both subsamples were specific and selective in terms of willingness to participate and motivation to share concerns. We hypothesized that hospital rates underrepresent the real incidence of regulation disorders, while area rates overrepresent them.

In summary, in our sample, the inclusion criterion was age under 3 years among those children who visited the 3 hospital departments or lived in neighboring areas. There were



Figure 1. The model of the data collection and the analyses in the For Healthy Offspring project.

no exclusion criteria. Data collection could be biased by some methodological issues such as willingness to participate and motivation to share concerns.

Although our sample was not representative, it was nevertheless adequately heterogeneous in all relevant sociodemographic characteristics (**Table 1**).

Questionnaires were given to parents (n=1164) by doctors and nurses working in 3 departments (Pediatric, Sleep, and Neurology) of the Heim Pál National Pediatric Institute (n=580) and also by health nurses and general practitioners in local areas (n=584). Mothers responded in 1133 cases.

Medical examinations and/or diagnostic evaluations were performed in 619 cases.

When completing the questionnaire, the parents were offered a complex screening program (a longer medical consultation) in our hospital if any of the most common behavior regulation disorders were present in the child. A total of 183 families took part in this complex diagnostic evaluation.

Afterwards, 35 parent-infant dyads also took part in smallgroup consultations. For some families, individual consultations and psychotherapy were recommended with the support volunteer hospital or other institutional professional. The process and professional content of the Screening Program are described below.

The Screening Program: Diagnostic Evaluation

Medical Consultation

Medical consultation (performed by NS and EP) consisted of a focused, detailed history taking that was followed by a physical examination. During the physical examination, healthy somatic, motor, and psychosocial developmental signs were carefully considered. In most cases, both parents were present when the child was examined; however, in a few cases, only the mother was present. The consultation normally lasted 1 h.

History taking started with the discussion of the symptom(s) that the parents had concerns about, and it was followed by general pediatric questions. To obtain broader knowledge of circumstances, certain aspects of the child's psychiatric history (birth circumstances, pregnancy, perinatal period, early sensory-motor and mental development, family sociodemographic characteristics) were also included. In all cases, the primary consideration was to determine the organic causes

Table 1. Demographic variables*.

Variable	Sample of mothers who filled in the questionnaires (N=1133)	Subsample where medical examinations were also conducted (n=619)		
	Percent/mean (SD)	Percent/mean (SD)		
Child's gender				
Воу	52.7%	55.2%		
Girl	47.3%	44.8%		
Child's age (months)	15.3 (10.8)	15.02 (11.07)		
0-2	9.7%	12.1%		
3-5	14.9%	15.9%		
6-8	10.1%	8.5%		
9-11	10.6%	10.3%		
12-17	16.0%	15.4%		
18-23	13.2%	12.0%		
24-36	25.4%	25.7%		
Number of children in the family	1.7 (0.9)	1.7 (0.9)		
No siblings	52.7%	52.4%		
1 sibling	35.5%	35.8%		
≥2 siblings	11.8%	11.8%		
Oldest	58.3%	57.6%		
Second	31.6%	31.6%		
Multiple	10.1%	10.8%		
Age of Mother (years)	32.0 (5.1)	31.7 (5.3)		
≤25	10.5%	11.9%		
26-30	25.6%	25.1%		
31-35	40.0%	40.3%		
36-40	20.2%	19.4%		
≥40	3.7%	3.3%		
Married/in common-law marriage	93%	90.7%		
Maternal education				
Maximum elementary or skilled worker	17.7%	20.6%		
High school	34.9%	34.0%		
Postgraduate/College degree or more	47.3%	45.4%		
Employed	66.2%	65.2%		

* Frequencies for valid answers. The proportion of missing values is 0-5%.

of the symptoms. If no physical abnormalities were found, we assumed the causative effect of psychosocial factors.

Diagnostic Evaluation of Organic Causes, Differential Diagnosis

The results of the first consultation determined the subsequent diagnostic steps. Diagnostic evaluation of the organic causes was performed depending on the presence of abnormal findings. Examinations were performed at the Pediatric Department as an outpatient service and included laboratory tests and radiological imaging tests. In some cases, specific examinations were performed by a gastroenterologist, neurologist, otorhinolaryngologist, ophthalmologist, or cardiologist. An obstructive sleep apnea symptoms test and, in cases of indeterminate symptoms of the infant or the presence of apnea, polysomnographic monitoring combined with esophageal pH monitoring was performed (by PB) at the Sleep Ambulance by Somnoscreen Plus (SOMNOmedics, GmbH, Randersacker, Germany). The following symptoms of unspecified/unexplained origin, usually with no underlying medical condition, were found: periodic breath-holding spells or strange breathing sounds, change of skin tone, loss of consciousness, loss of postural tone, and seizure. A complex motor evaluation of the infant was completed by a physiotherapist.

In cases of medical illness, our work was based on general pediatric diagnostic steps. We had more difficulty diagnosing early behavior regulation disorders. The International Classification of Diseases, 10th Revision (ICD-10) [16], which is used in Hungary, does not include clear directions about early childhood regulation disorders. In the differential diagnostics of regulation disorders, we relied on the principles of the German system, as described by Hédervári-Heller [17] in Hungarian and German, although it is not widely used in Hungary. Moreover, we greatly profited from reviewing the classification criteria of the Diagnostic Classification of Mental Health and Developmental Disorders of Infancy and Early Childhood – Revised (DC: 0-3R) [18], which was not used in Hungary at that time.

Small Group Consultation

As long as no organic abnormality was found, part of the sample was offered a small-group consultation. These consultations took place in the presence of the parents (or only the mothers) and the infants on 2 occasions, each lasting 90 min. The consultations were conducted by a pediatric psychotherapist (TN) with the participation of psychologists and psychiatrists from the hospital. During the small-group consultations, the following aspects were spontaneously observed: (a) infant's developmental level and the quality of his or her playing activity; (b) quality of the parent-infant interactions; (c) emotional reactions between the parents and their moods while together; (d) parent-infant attachment patterns based on the balance between exploration and attachment behaviors; (e) infant's attitude toward the parents and the professionals in attendance; and (f) ambience of the consultation meeting. At the end of the second meeting, the experiences were discussed with the families in a private meeting, and individual therapy for the parents or parent-infant consultations were offered if necessary. As professionals worked together in a team, they decided by consensus if regulation problems were present and whether additional care was needed due to psychosocial, relationship, or interaction difficulties. In many cases, the 2 small-group consultations were sufficient to resolve mild regulation difficulties.

Measuring Instruments

Questionnaires

A questionnaire package was specifically designed for this study. In our edited basic questionnaire, the parents were asked in detail about their family background, housing and job circumstances, their financial status, their health, healthrelated and psychological characteristics of the pregnancy and the birth of the examined infant, the newborn period, breastfeeding and early care, the infant's physical and mental condition, and his or her behavioral characteristics. We then focused on the 3 main areas of regulation disorders. Detailed questions were asked about each topic to determine if the infant was affected by intense crying and restlessness, feeding and weight gain difficulties, or sleep disorders. Most of the questions were closed-ended, discrete questions with 2 or more possible answers, or Likert-type scale items, usually with 5 levels. Only the "other" questions were open ended. A pilot study was conducted to ensure that the questions were well understood by the respondents and could be answered easily and reliably. We were not able to measure internal consistency because these questions were single items and did not form scales. In this article, we used only the socio-demographic information and some Likert-type questions on early regulation (crying, sleep, and feeding).

Medical Diagnoses

The diagnoses determined in course of medical examinations and during individual and small-group consultations (see the detailed description of the Screening Program above) were included in our database.

Statistical Analysis

Quantitative analyses were performed using IBM SPSS Statistics 20.0 software package (IBM, Armonk, NY, USA).

	Not diagnosed	Diagnosed	z-statistics in
	n; Mean (SD)	n; Mean (SD)	Mann-Whitney tests
Medical diagnoses: Excessive crying, restlessness			
How can you characterize your child by strong prolonged crying when he/she was a little baby? (5-point Likert scale)	n=507	n=92	z=-5.205;
	2.16 (1.10)	2.90 (1.28)	p<0.001
According to your observations, how much was your child crying or how fussy was your child in the previous two weeks?	n=486	n=90	<i>z</i> =-3.227;
	2.83 (1.12)	3.31 (1.28)	p=0.001
How could you soothe him/her when he/she was crying?	n=505	n=92	<i>z</i> =-3.641;
(5-point Likert scale)	4.32 (0.77)	3.97 (0.90)	p<0.001
How much distress does it cause you when your child is crying? (5-point Likert scale)	n=501	n=91	<i>z</i> =-3.208;
	3.59 (1.22)	4.05 (0.95)	p=0.001
Medical diagnoses: Loss of appetite – normal weight gain			
Is feeding your child a challenge for you?	n=534	n=28	z=-2.38;
(5-point Likert scale)	1.72 (1.03)	2.39 (1.47)	p=0.018
Medical diagnoses: No weight gain or weight loss – alimentary			
Is feeding your child a challenge for you?	n=534	n=28	z=-3.71;
(5-point Likert scale)	1.72 (1.03)	2.39 (1.47)	p<0.001
Medical diagnoses: Sleep disorders (sleep awakenings, sleep-onset problems)			
Generally, how many times does your child wake up during the night?	n=543;	n=19;	z=-6.618;
	1.71 (1.04)	2.79 (1.40)	p<0.001
How much distress does it cause you when your child wakes up? (5- point Likert scale)	n=467;	n=91;	<i>z</i> =-7.51
	2.28 (1.26)	3.48 (1.29)	p<0.001
How much distress does it cause you to put your child to sleep? (5-point Likert scale)	n=499;	n=92;	z=-5.75;
	1.76 (1.00)	2.55 (1.32)	p<0.001

Table 2. Diagnoses of regulation disorders in medical examinations and differences between maternal answers in questionnaires.

Table 6 shows the significant differences between means and standard deviations of answers to questions in parental questionnaire in subgroups where different regulation disorders were diagnosed or not diagnosed.

For the descriptive statistics in this article, we calculated prevalence distributions and for examining associations between regulation problems and other health conditions, we ran crosstabs (χ^2 -tests) in the cases of diagnoses (yes/no; 1/0 categorical variables) and Mann-Whitney tests in the cases of questionnaire data (Likert-type items, see **Table 2**).

Ethical Approval

The Institutional Ethics Committee of Heim Pál Children's Hospital approved the study (authorization number: 11/04.2010).

Informed Consent

Informed consent was obtained from the parents of all individual participants included in the study.

Results

Regulation Disorders in Medical Examinations and Screening Program

A total of 1133 mothers (**Table 1**) answered the questionnaires. In a subsample, 619 children had also medical examinations. The sex distribution was almost half and half, male and female, in both groups. The children's ages and other sociodemographic factors were almost the same or very similar in each subgroup. In the whole sample, the average age of children was 15.3 ± 10.8 months. The average number of children in the families was 1.7 ± 0.9 . The average age of the mothers was 32 ± 5.1 years, and more than 90% were married. In addition to medical diagnoses, the main early childhood behavior regulation disorders are also categorized in **Table 3**. In the subsample in which medical examinations were conducted, excessive crying and/or restlessness was present in 15.0%, sleep disorders in Table 3. Frequency of early behavior regulation problems and other frequent medical diagnoses*.

Diagnostic categories in the medical examinations	Subsample where medical examinations were conducted (n=619)	Not referred to the screening program after medical examinations (n=436)	Referred to the screening program after medical examinations (n=183)
Excessive crying, restlessness	15.0%	5.7%	37.2%
Sleeping problems			
Sleep disorders (sleep awakenings, sleep-onset problems)	15.2%	0.7%	49.7%
Snoring	2.9%	1.6%	6.0%
Feeding and weight gain problems			
Breastfeeding difficulty	10.3%	4.1%	25.1%
Loss of appetite – normal weight gain	5.0%	1.8%	12.6%
No weight gain or weight loss – alimentary	3.2%	0.9%	8.7%
No weight gain or weight loss – organic	6.6%	8.3%	2.7%
Breathing problems			
Irregular breathing	4.5%	4.8%	3.8%
Affective apnea	1.9%	1.6%	2.7%
Breathing stops	6.6%	9.2%	0.5%
Indeterminate symptoms of the infant	10.5%	12.8%	4.9%
Symptoms of possible neurological conditions			
Uncertain sickness, bizarre movements	4.8%	5.5%	3.3%
Activity or concentration difficulties	4.4%	1.4%	11.5%
Pulmonological conditions or recurrent upper airway infec	tions		
Recurrent upper airway infections	24.2%	25.5%	21.3%
Recurrent wheezing	23.9%	30.7%	7.7%
Subglottic laryngitis	9.9%	11.5%	6.0%
Gastrointestinal complaints			
Abdominal colic	12.3%	8.7%	20.8%
Constipation	4.0%	1.6%	9.8%

* In n=619 cases, medical examinations were carried out, followed by a deeper screening program for regulation disorders in n=183 cases, while in n=436 cases this was not necessary. Table 2 presents the prevalences of different disease and disorder categories in each subgroup, showing higher prevalences of regulation disorders in the screening subgroup (see bold figures).

15.2%, breastfeeding problems in 10.3%, and collective feeding disorders in 14.8%. The prevalence of constipation was 4.0%, while abdominal colic was present in 12.3%. Medical examinations were followed by a deeper screening program for regulation disorders in 183 cases, while in 436 cases this was not necessary. **Table 3** presents the prevalence of different disease and disorder categories in each subgroup, showing higher prevalence of regulation disorders in the screening subgroup.

Comorbidity Between Different Early Childhood Regulation Disorders

In cases in which excessive crying was present, the comorbidity with sleep disorders was 50% ($\chi^2(1)=106.20$; *P*<0.001); with breastfeeding disorders 22.6% ($\chi^2(1)=17.69$; *P*<0.001); and with loss of appetite 11.8% ($\chi^2(1)=10.70$; *P*<0.001). In those cases in which sleep disorders were present, the comorbidity with

COMORBIDITY Diagnostic categories in the medical examinations	In subgroup where excessive crying, restless- ness is not present	In subgroup where excessive crying, restless-ness is present	Crosstabs (χ²-tests)	In subgroup where sleep disorders (sleep awakenings, sleep-onset problems) is not present	In subgroup where sleep disorders (sleep awakenings, sleep-onset problems) is present	Crosstabs (χ²-tests)	In subgroup where breast- feeding difficulty is not present	In subgroup where breast- feeding difficulty is present	Crosstabs (χ²-tests)
Sleep disorders (sleep awakenings, sleep-onset problems) present in (%)	8.8%	50.0%	χ ² (1)=106.20; <i>p</i> <0.001						
Breastfeeding difficulty present in (%)	8.2%	22.6%	χ ² (1)=17.69; <i>p</i> <0.001	12.4%	39.1%	χ ² (1)=31.60; <i>p</i> <0.001			
Loss of appetite – normal weight gain present in (%)	3.8%	11.8%	χ ² (1)=10.70; p=0.001				4.0%	14.1%	χ ² (1)=12.30; <i>p</i> <0.001
No weight gain or weight loss – alimentary present in (%)							2.3%	10.9%	χ ² (1)=13.56; <i>p</i> <0.001

Table 4. Comorbidity between different early childhood regulation problems*.

* Crosstabs (χ^2 -tests): χ^2 (df); *p*; frequencies of joint regulation disorders. Table 3 shows the prevalence of important regulation disorders in subgroups where a specific disorder is present or not. E.g. If excessive crying is present, sleep disorders are also present in 50% of the cases, but this proportion is only 8.8% in a subgroup where excessive crying is not a significant complaint.

breastfeeding disorders was 39.1% ($\chi^2(1)=31.60$; *P*<0.001). Other data are displayed in **Table 4**.

Comorbidity Between Early Childhood Regulation Disorders and Other Health Conditions

In those cases in which excessive crying was present, the comorbidity with concentration difficulties was 10.8% ($\chi^2(1)=10.71$; P<0.001) and with abdominal colic it was 26.9% ($\chi^2(1)=21.67$; P=0.001). When sleep disorders or breastfeeding difficulties or organic feeding difficulties were diagnosed, abdominal colic was also present in 21.3% ($\chi^2(1)=8.33$; P<0.001), 23.4% ($\chi^2(1)=8.25$; P=0.004), and 34.1% ($\chi^2(1)=19.50$; P<0.001) of the cases, respectively. Interestingly, pulmonological conditions or recurrent upper airway infections were more frequent in the subgroups in which regulation problems were not a concern. The other relationships are shown in **Table 5**.

Regulation Disorders in the Parental Report

According to the questionnaire answers, 14.7% of the mothers' reported low self-confidence when interpreting their infant's signs; it was 22.1% in a subgroup that was referred to the screening program. A total of 15.6% of mothers characterized their children as strong criers (24.4% in a subgroup that was referred to the screening program). A total of 16% of the children had some type of feeding or weight gain disorder (32.6% in the screening subgroup), and 10% awoke 4 or more times during the night (21% in the screening subgroup). The frequency of symptoms of crying, feeding, and sleeping per parent reporting are shown in **Table 6**.

Diagnoses of Regulation Disorders in Medical Examinations and Differences Among Maternal Answers in Questionnaires

According to the z-statistics in Mann-Whitney tests, mothers of children with a diagnosis of excessive crying reported

COMORBIDITY Diagnostic categories in the medical examinations		Activity or concentration difficulties (%)	Recurrent upper airway infections (%)	Recurrent wheezing (%)	Subglottic laryngitis (%)	Abdominal colic (%)	Constipation (%)
Excessive crying, restlessness	not present vs present	3.2 < 10.8	26.0 > 14.0	27.2 > 5.4	11.0 > 3.2	9.7 < 26.9	
	Crosstabs (χ²-tests)	χ ² (1)=10.7; <i>p</i> =0.001	χ ² (1)=6.27; <i>p</i> =0.012	$\chi^2(1)=20.66;$ p<0.001	$\chi^2(1)=5.41;$ p=0.020	χ ² (1)=21.67; <i>p</i> <0.001	
Sleep disorders (sleep awakenings,	not present vs present	3.4 < 9.6		27.4 > 4.3	11.2 > 2.1	10.7 < 21.3	3.2 < 8.5
sleep-onset problems)	Crosstabs (χ²-tests)	$\chi^2(1)=7.22;$ p=0.007		$\chi^{2}(1)=25.53;$ p<0.001	$\chi^2(1)=7.45;$ p=0.006	χ ² (1)=8.33; <i>p</i> =0.004	χ ² (1)=5.72; p=0.017
Breastfeeding difficulty	not present vs present	3.6 < 10.9		25.9 > 6.3		11.0 < 23.4	
	Crosstabs (χ²-tests)	$\chi^2(1)=7.40;$ p=0.007		χ ² (1)=12.24; <i>p</i> <0.001		χ ² (1)=8.25; <i>p</i> =0.004	
Loss of appetite – normal weight	not present vs present		25.0 > 3.2				
gain	Crosstabs (χ²-tests)		$\chi^2(1)=7.67;$ p=0.006				
No weight gain or weight loss – alimentary	not present vs present	4.0 < 15.0		24.5 > 5.0			
	Crosstabs (χ²-tests)	$\chi^2(1)=5.61;$ p=0.018		χ²(1)=4.06; p=0.044			
No weight gain or weight loss –	not present vs present			25.1 > 7.3		10.7 < 34.1	
organic	Crosstabs (χ²-tests)			$\chi^2(1)=6.65;$ p=0.010		$\chi^2(1)=19.50;$ p<0.001	

Table 5. Comorbidity between early childhood regulation problems and other health conditions*.

* Crosstabs (χ^2 -tests): χ^2 (df); P; frequencies of joint diagnoses. Table 4 shows the prevalence of important somatic complaints in subgroups where a specific regulation disorder is present or not. E.g. If excessive crying is present, abdominal colic is also present in 26.9% of the cases, but this proportion is only 9.7% in a subgroup where excessive crying is not a significant complaint.

significantly more problematic crying behavior (long prolonged crying in early infancy: P<0.001, crying and fussiness in the last 2 weeks: P=0.001, soothability: P<0.001, parental distress: P=0.001) compared with those without this diagnosis.

Mothers of children with a diagnosis of loss of appetite or weight loss reported significantly more problematic feeding behavior (feeding as a challenge: P<0.001) compared with mothers of children who did not have this diagnosis.

Children with a diagnosis of sleep disorders had significantly more problematic sleep behavior reported based on the parental questionnaire (nightwakings, sleep onset, parental distress compared with those who did not have this diagnosis; P<0.001 for all differences) using Likert scale in the questionnaire. Detailed results are shown in **Table 2**.

Discussion

Crying is part of the normal development of an infant. It is a form of communication with parents and results from various stimuli, such as hunger, discomfort, or pain. Excessive crying in the early months is a frequent concern. Pediatricians have to understand and adequately manage the problem and offer support to exhausted parents. Excessive crying may interfere with the mother-infant, father-infant, and mother-father interactions and may increase the risk of child abuse [19,20]. Over the last 15 years, other regulatory disorders of early childhood have attracted the increased attention of both researchers and pediatric practitioners [21-23]. Table 6. Frequencies of behavior regulation difficulties and feelings about them reported by mothers in questionnaires.

Questions and emphasized values	Sample of mothers filled in questionnaires (n=1133) n; %	Subsample where medical examinations were also conducted (n=619) n; %	Subsample referred to the screening program (n=183) n; %	Subsample not referred to the screening program (n=981) n; %
How confident are/were you in interpreting your infant's signs? (5-point Likert scale: <i>values 1-3 for low self-confidence</i>)	1121	595	181	940
	14.7%	16.5%	22.1%	13.3%
CRYING				
How can you characterize your child by strong prolonged crying when he/she was a little baby? (5-point Likert scale: <i>values 4-5</i>)	1121 15.6%	599 16.5%	180 24.4%	947 13.9%
According to your observations, how much was your child crying or how fussy was your child in the previous two weeks? (<i>more than 1 hour</i>) (<i>more than 3 hours</i>)	1112 9.3% 4.5%	576 12.3% 8.6%	177 17.5% 5.6%	935 7.7% 3.7%
How could you soothe him/her when he/she was crying? (5-point Likert scale: <i>values 1-3 for low self-confidence</i>)	1118	597	180	938
	12.5%	15.6%	20.0%	11.1%
How much distress does it cause you when your child is crying? (5-point Likert scale: <i>values 4-5</i>)	1113	592	176	937
	54.5%	61.2%	69.3%	51.7%
FEEDING				
Does your child have any feeding or weight gain problems? (<i>yes</i>)	1099	584	175	924
	16.0%	18.2%	32.6%	12.9%
Is feeding your child a problem for you?	1058	562	175	883
(5-point Likert scale: <i>values 4-5</i>)	9.6%	9.4%	20.0%	7.5%
SLEEPING				
Generally, how many times does your child wake up during the night? (<i>4 times or more</i>)	1105	587	176	929
	10.0%	11.2%	21%	7.9%
How much distress does it cause you when your child wakes up? (5-point Likert scale: <i>values 4-5</i>)	1033 23.6%	558 23.0%	167 34.7%	866 21.5%
How can your child fall asleep? (it's difficult even after a long process of putting down)	1105	588	177	928
	8.9%	9.0%	14.7%	7.8%
How much distress does it cause you to put your child to sleep? (5-point Likert scale: <i>values 4-5</i>)	1115	591	178	937
	9.3%	9.0%	14.6%	8.3%

* *n*=valid answers to the question; frequency of emphasized values. Table 5 shows the total number of answers (n) to separate questions and the proportions of answers (%) indicating problems.

Sleeping Disorders

In international pediatrician surveys and cross-cultural comparisons, sleeping disorders (night awakenings and sleep-onset difficulties) are one of the most frequent (10-76%) parental concerns [1,2]. In our large sample (N=1133) questionnaire study, 10% of the children had sleeping disorders. In a subsample in which diagnoses were determined based on medical examinations and consultations, we found an incidence of 15.2% for sleep disorders. This rate was similar to reports from other countries [1,2].

Excessive Crying

The occurrence of excessive crying was 8-30% in previous largesample studies [3,24,25]. In our questionnaire study, 15% of

mothers reported intensive crying in the infants. In a subsample in which diagnoses were determined based on medical examinations and consultations, we found an incidence of 15.0% of excessive crying and/or restlessness. Prevalence rates for excessive crying were lower in some European countries, with 1.5% in the Netherlands [2] and 9.2% in Denmark [26], but comparable to the 16.3% reported in Germany [27].

Eating Disorders

Eating disorders are also common in infancy. Prevalence numbers range from 20% to 25% in the normal population [4,28,29] and from 40% to 80% in infants with disabilities. Mild feeding difficulties occur in approximately 30% of children [4]. The prevalence of clinical feeding disorders is 3-10% [5], and the incidence of more severe failure to thrive is about 3-4% [1].

According to our questionnaire results, 16% of the children had feeding disorders. In a subsample in which diagnoses were determined based on medical examinations and consultations, we found an incidence of 10.3% for breastfeeding disorders and 14.8% for different feeding disorders.

Sleep and feeding disorders are the leading concerns in clinical samples. In a large sample (N=701) in one of the most renowned European clinical programs, the Munich Program [30], occurrence of problems for children were the following: sleep disorders (62.8%), feeding disorders (40.4%), chronic restlessness-motor activity-lack of interest in play (30.1%), excessive crying (29.4%), dysfunctional sleep-wake organization (25.8%), excessive defiance (20.3%), excessive clinging-separation anxiety-social withdrawal (12.3%), and aggression-oppositional behavior (6.8%).

Multiple Regulatory Disorders

Behavioral disorders in infancy can affect an infant's development [21,22,31,32]. Furthermore, infants with behavioral disorders are more likely to have impaired parent-infant relationships [23,33,34].

Approximately 20% of all infants show symptoms of excessive crying, sleeping, or feeding disorders in the first year of life [31]. The prevalence of colic in infants is about 20%, but it depends on parental perception of crying [35]. In our study, the prevalence was 12.3%. In a systematic review and meta-analysis, colic prevalence at 5-6 weeks of age (25.1%) was significantly higher than colic prevalence at 8-9 weeks of age (10.8%) [36]. Most maternal and child health nurses were unaware of the evidence that crying is not associated with gastro-esophageal reflux, but most of them reported that reflux causes pain [37].

Further, a small minority of children (1-2%) will manifest all 3, leading to multiple regulatory disorders [27,38]. Infants with multiple moderate-to-severe regulatory problems experience >10 times the odds of clinically significant mental health concerns during childhood, and these symptoms appear to worsen over time [39].

Comorbidity

Little is known about the association between excessive crying and sleeping or eating disorders in population samples [40]. We found comorbidity among different regulation disorders. Where one type of behavior regulation disorder was present, another type of regulation disorder was more frequently diagnosed (**Table 4**). Crosstabs (χ^2 tests) have proved (*P*<0.05) that infants who were referred to the screening program because of medical considerations had more frequent behavior regulation disorders than other infants in our study. This, in turn, indicates that the differential diagnostic process was successful in our program.

Wolke et al [41] found that 32.7% of parents reported that their infant had a crying, sleeping, or feeding disorder, and a further 14.6% reported their infant as having more than one of these disorders. Specifically, comorbidity was most likely to occur between crying and sleep disorders. Multiple regulatory problems may identify infants with a high burden of comorbidity that extends into childhood [39,42].

In a retrospective study by von Kries et al [27], a higher prevalence of sleep and eating disorders was found in children up to 4 years of age who were reported to have had excessive crying beyond the sixth month. In our study, the connection between sleep disorders and excessive crying was 50%, but some studies have not found a relationship between excessive crying and sleep disorders and other indicators of multiple regulatory disorders [43], while other studies have found either sleeping or other regulatory disorders [41,44].

The few children with excessive crying and either severe sleeping or eating disorders might constitute a group of infants with multiple regulatory disorders outside the continuous spectrum of normal behavior [40].

Moreover, there is growing evidence of the negative implications for infants whose excessive, persistent crying is present with other regulatory disorders, such as feeding and sleeping disorders [41,45].

A meta-analysis of 22 longitudinal studies showed evidence associating excessive crying and other regulatory disorders (sleeping and eating) in the first months of life with adaptive problems at school age, mainly related to attention deficit-hyperactivity disorder symptoms and associated behaviors [31,32,46,47]. In our study, the comorbidity between excessive crying and concentration difficulties was 10.8%.

There was no connection between regulation and breathing disorders. Breathing stops and sleep disorders were successfully differentiated. Among those diagnosed with sleep disorders, there were fewer infants who had breathing stops than among the others. Also, there was no relationship with symptoms of possible neurological conditions and uncertain sickness or abnormal movements.

Infants who had early activity or concentration disorders were significantly more highly represented among those who had behavior regulation disorders as well. The occurrence of abdominal colic and constipation, symptoms that are often found to have a psychosomatic background, also correlated with the occurrence of other regulation disorders. The occurrence of diagnoses of excessive crying, sleep disorders, and breastfeeding and other feeding disorders was significantly lower in children in whom recurrent upper airway infections, recurrent wheezing, or laryngitis were diagnosed (**Table 5**).

Questions about both parental observations and subjective feelings were asked in questionnaires. Some important extreme values of questions are presented for the whole sample, for the subsample in which medical examinations were also conducted, and for the subsample that was referred to the screening program. Mothers of infants who were referred to the screening program because of medical considerations reported more problems in the questionnaires than other mothers in our study (**Table 6**).

The mothers of the infants with diagnoses of excessive crying, sleep, or feeding disorders reported significantly more problematic behavior in questionnaires as well (**Table 2**).

Our screening program included children who showed signs of regulation disorders and who were referred for a detailed diagnostic evaluation. The incidence of diagnoses in this program was the following: sleep disorders (49.7%), excessive crying (37.2%), and functional feeding disorders (no organic background; 21.3%).

In order to offer appropriate medical support, the differential diagnostic process is important in separating acute secondary symptoms (e.g., crying, mild and transient sleep and feeding disorders because of general discomfort, nonspecific complaints, pain) of frequent child illnesses (eg, respiratory diseases) from comorbid chronic behavior regulation disorders. In our study, we investigated the relationship between regulation disorders and other health conditions from medical records for which a screening model enabled careful and thorough differential diagnostics.

Outpatient treatment is sufficient for crying or sleeping disorders in infancy. However, hospitalization may be required for feeding disorders because not all feeding disorders can be treated on an outpatient basis [48].

In a study conducted by Schmid and Wolke [46], excessive infant crying (10.1%) was specifically associated with maternal anxiety disorders, especially in infants of younger and less educated first-time mothers. Feeding disorders (36.4%) were predicted by maternal anxiety (and comorbid depressive) disorders in primiparous mothers and infants with lower birth weight. Infant sleeping disorders (12.2%) were related to maternal depressive (and comorbid anxiety) disorders irrespective of maternal parity [49-52].

In our experience, designating risk groups in pediatric care is a complex problem. On the one hand, many parents may report regulation disorders, while based on a strict diagnostic system, a clinical disorder cannot be determined. On the other hand, we can assume that there are many hidden cases in which infants could have a clinically relevant regulation disorder but their parents do not interpret these behaviors as being problematic and do not report them to pediatricians or health care nurses. Thus, we can assume that some cases could remain hidden, while in others, study is requested without a clinical basis. In the latter circumstance attention from a therapeutic perspective is still necessary because of the parents' concern. Questions about the parents' feelings about the specific problems can contribute to a better understanding of the actual situation.

Limitations

Most of the data in the current study were collected via maternal reports. These are often limited by social desirability and reporter bias. Another limitation of this study was our lack of access to father-child interactions. Thus, future research should include both mothers and fathers. Another limitation there was no ethnic diversity, with only white participants in the study. It is important to note that our study used only a short period of data collection. Finally, our sample is not representative; the results can be regarded as estimations for recognizing early childhood regulation disorders in pediatric care.

Conclusions

The For Healthy Offspring Project was the first study to examine the prevalence and the complex (medical and psychosocial) background of the classic behavior regulation disorders (excessive crying, feeding, and sleep problems) in infancy and toddlerhood in Hungary. In this article, the relationship between regulation disorders and other health conditions were investigated. Crying and sleep and feeding disorders are challenging for most parents, but only a small fraction of these cases can be categorized as clinical disorders. Although our study is not representative, according to our findings, we can hypothesize that the general incidence of early childhood regulation disorders in international research of 5-15% is likely similar to that found in Hungary as well. This study added more information about the associations between regulation problems and other health conditions. Our model for screening enabled the careful differential diagnostic process of separating acute secondary symptoms from comorbid chronic behavior regulation disorders. We highlighted that no other data are currently available on the frequency and types of early childhood regulation problems in Hungary. In order to effectively recognize early behavior regulation disorders in daily practice, diagnostic instruments widely used in international field should be adapted in general Hungarian pediatric care.

References:

- 1. Zeanah CH Jr., ed. Handbook of infant mental health, $3^{\rm rd}$ ed. New York, NY: The Guilford Press; 2009
- Mindell JA, Sadeh A, Wiegand B, et al. Cross-cultural differences in infant and toddler sleep. Sleep Med. 2010;11(3):274-80
- 3. Wurmser H, Laubereau B, Hermann M, et al. Excessive infant crying: Often not confined to the first 3 months of age. Early Hum Dev. 2001;64(1):1-6
- McDermott BM, Mamun AA, Najman JM, et al. Preschool children perceived by mothers as irregular eaters: Physical and psychosocial predictors of a birth cohort study. J Dev Behav Ped. 2008;29:197-205
- Corbett SS, Drewett RF. To what extent is failure to thrive in infancy associated with poorer cognitive development. A review and meta-analysis. J Child Psychol Psyc. 2004;45:641-54
- 6. Herman M, Le A. The crying infant. Emerg Med Clin N Am. 2006;25:1137-59
- 7. Bernard-Bonnin AC. Feeding problems of infants and toddlers. Can Fam Physician. 2006;52:1247-51
- 8. Carter JC, Wrede JE. Overview of sleep and sleep disorders in infancy and childhood. Pediatr Ann. 2017;46(4):e133-38
- 9. Keren M. Eating and feeding disorders in the first five years of life: revising the DC: 0-3R diagnostic classification of mental health and developmental disorders of infancy and early childhood and rationale for the new DC: 0-5 proposed criteria. Infant Ment Health J. 2016;37(5):498-508
- 0-5TM Diagnostic Classification of Mental Health and Developmental Disorders of Infancy and Early Childhood. Washington, DC: Zero to Three Press; 2016
- 11. Hinshaw SP. Developmental psychopathology as a scientific discipline: relevance to behavioral and emotional disorders of childhood and adolescence. In: Beauchaine TP, Hinshaw SP, editors. Child and adolescent psychopathology. Hoboken, NJ: John Wiley & Sons Inc.; 2008;3-26
- Papousek M, Schieche M, Wurmser H, editors. Disorders of behavioral and emotional regulation in the first years of life: Early risks and intervention in the developing parent-infant relationship. Washington, DC: Zero to Three, 2008
- 13. Masten AS, Monn AR. Child and family resilience: A call for integrated science, practice, and professional training. Fam Relat. 2015;64:5-21
- 14. De Gangi GA. Pediatric disorders of regulation in affect and behavior, 2nd ed. London: Academic Press; 2017;38
- Keren, M. Eating and feeding disorders in early childhood. In: Zeanah CH, editor. Handbook of infant mental health, 4th ed. New York-London: Guilford Press; 2018;383-97
- ICD-10 International statistical classification of diseases and related health problems, 10th revision, Volume 2. Geneva: World Health Organization; 2010

Acknowledgments

We would like to thank all the families who participated, the hospital team, and the research partners (especially Judit Gervai et al at the Hungarian Academy of Sciences) who contributed to our work. The program could not have been organized without their generous cooperation and assistance.

Conflicts of Interest

None.

- Hédervári-Heller É. [Parent-infant consultation and therapy. Behavioral disorders in infancy and early childhood]. Budapest: Animula; 2008 [in Hungarian]
- Diagnostic classification of mental health and development disorders of infancy and early childhood: DC: 0-3R. Washington, DC, Zero To Three; 2005.
- Botha E, Joronen K, Kaunonen M. The consequences of having an excessively crying infant in the family: an integrative literature review. Scand J Caring Sci. 2019;33(4): 779-90
- Tedgård E, Tedgård U, Råstam M, et al. Parenting stress and its correlates in an infant mental health unit: a cross-sectional study. Nord J Psychiatry. 2020;74(1):30-39
- 21. Rocha NACF, Silva FPS, Santos MM, et al. Impact of mother-infant interaction on development during the first year of life: A systematic review. J Child Health Care. 2019;24(3):365-85
- 22. Toffol E, Rantalainen V, Lahti-Pulkkinen M, et al. Infant regulatory behavior problems during first month of life and neurobehavioral outcomes in early childhood. Eur Child Adolesc Psychiatry. 2019;28:847-59
- 23. Northrup JB, Iverson JM. The development of mother-infant coordination across the first year of life. Dev Psychol. 2020;56(2):221-36
- 24. Kim JS. Excessive crying: Behavioral and emotional regulation disorder in infancy. Korean J Pediatr. 2011;54:229-33
- Reijneveld SA, Brugman E, Hirasing RA. Excessive infant crying: the impact of varying definitions. Pediatrics. 2001;108:893-97
- 26. Alvarez M. Caregiving and early infant crying in a Danish community. J Dev Behav Pediatr. 2004;25(2):91-98
- von Kries R, Kalies H, Papousek M. Excessive crying beyond 3 months may herald other features of multiple regulatory problems. Arch Pediatr Adolesc Med. 2006;160(5):508-11
- Wright CM, Parkinson KN, Shipton D, et al. How do toddler eating problems relate to their eating behaviour, food preferences, and growth? Pediatrics. 2007;120(4):e1069-75
- 29. Chatoor I. Diagnosis and treatment of feeding disorders in infants, toddlers and young children. Washington, DC: Zero to Three; 2009
- 30. Wurmser H, Papousek M. Facts and figures: database of the Munich interdisciplinary research and intervention program for fussy babies. In: Papousek M, Schieche M, Wurmser H, editors. Disorders of behavioral and emotional regulation in the first years of life: Early risks and intervention in the developing parent-infant relationship. Washington, DC: Zero to Three; 2008;27-52
- Hemmi MH, Wolke D, Schneider S. Associations between problems with crying, sleeping and/or feeding in infancy and long-term behavioral outcomes in childhood: A meta-analysis. Arch Dis Child. 2011;96(7):622-29
- Bilgin A, Baumann N, Jaekel J, et al. Early crying, sleeping, and feeding problems and trajectories of attention problems from childhood to adulthood, child development. Child Dev. 2020;91(1):e77-91

- Papousek M, von Hofacker N. Persistent crying in early infancy: A non-trivial condition of risk for the developing mother infant relationship. Child Care Health Dev. 1998;24(5):395-424
- Martini J, Petzoldt J, Knappe S, et al. Infant, maternal, and familial predictors and correlates of regulatory problems in early infancy: The differential role of infant temperament and maternal anxiety and depression. Early Hum Dev. 2017;115:23-31
- Bellù R, Condò M. Functional gastrointestinal disorders in newborns: Nutritional perspectives. Pediatr Med Chir. 2018;40:198
- 36. Wolke D, Bilgin A, Samara M. Systematic review and meta-analysis: Fussing and crying durations and prevalence of colic in infants, J Pediatr. 2017;185:55-61
- McGann J, Manohar J, Hiscock H, et al. Caring for crying babies: A mixedmethods study to understand factors influencing nurses' and doctors' management of infant colic. J Pediatr Child Health. 2018;54:653-60
- Schmid G, Schreier A, Meyer R, et al. A prospective study on the persistence of infant crying, sleeping and feeding problems and preschool behavior. Acta Pediatr. 2010;99:286-90
- 39. Cook F, Giallo R, Hiscock H, et al. Infant regulation and child mental health concerns: A longitudinal study. Pediatrics. 2019;143(3): e20180977
- 40. Barr RG. Colic and crying syndromes in infants. Pediatrics. 1998;102(Suppl. E):1282-86
- Wolke D, Meyer R, Ohrt B, et al. Co-morbidity of crying and feeding problems in infancy: Concurrent and predictive associations. Early Dev Parent. 1995;4(4):191-207
- 42. Olsen AL, Ammitzbøll J, Olsen EM, et al. Problems of feeding, sleeping and excessive crying in infancy: A general population study. Arch Dis Child. 2019;104:1034-41

- Kirjavainen J, Kirjavainen T, Huhtala V, et al. Infants with colic have a normal sleep structure at 2 and 7 months of age. J Pediatr. 2001;138:218-23
- 44. Canivet C, Jakobsson I, Hagander B. Infantile colic. Follow-up at four years of age: Still more "emotional." Acta Paediatr. 2000;89:13-17
- 45. Wolke DP, Schmid SG, Schreier AP, et al. Crying and feeding problems in infancy and cognitive outcome in preschool children born at risk: A prospective population study. J Dev Behav Pediatr. 2009;30(3):226-38
- 46. Schmid G, Wolke D. Preschool regulatory problems and attention-deficit/ hyperactivity and cognitive deficits at school age in children born at risk: different phenotypes of dysregulation? Early Hum Dev. 2014;90:399-405
- 47. Santos IS, Matijasevich A, Capilheira MF, et al. Excessive crying at 3 months of age and behavioral problems at 4 years age: A prospective cohort study. J Epidemiol Community Health. 2015;69:654-59
- 48. Bolten MI. Infant psychiatric disorders. Eur Child Adolesc Psychiatry. 2013;22(Suppl. 1): S69-S74
- Petzoldt J, Wittchen HU, Einsle F, et al. Maternal anxiety versus depressive disorders: Specific relations to infants' crying, feeding and sleeping problems. Child Care Health Dev. 2016;42(2):231-45
- Samdan G, Kiel N, Petermann F, et al. The relationship between parental behavior and infant regulation: a systematic review. Dev Rev. 2020;57:100923
- Dias CC, Figueiredo B. Mother's prenatal and postpartum depression symptoms and infant's sleep problems at 6 months. Infant Ment Health J. 2020;41(5):614-27
- 52. Væver MS, Pedersen IE, Smith-Nielsen J, et al. Maternal postpartum depression is a risk factor for infant emotional variability at 4 months. Infant Ment Health J. 2020;41(4):477-94