

Literature review on chemical pollutants in indoor air in public settings for children and overview of their health effects

with a focus on schools, kindergartens and day-care centres



Supplementary publication to the screening tool for assessment of health risks from combined exposure to multiple chemicals in indoor air in public settings for children

Abstract

This publication summarizes the scientific information on children's health outcomes related to exposure to chemical pollutants in indoor air in public settings; on chemicals that are commonly detected in indoor air in schools, kindergartens and day-care centres; and on the likelihood of co-occurrence of these chemicals. It served as a background document to facilitate the selection of priority adverse effects (health endpoints) and chemicals to be included in the WHO Regional Office for Europe's screening tool for assessment of risks from combined exposure to multiple chemicals in indoor air in public settings for children.

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with a focus on schools, kindergartens and day-care centres

Supplementary publication to the screening tool for assessment of health risks from combined exposure to multiple chemicals in indoor air in public settings for children

CONTENTS

ACKNOWLEDGEMENTS	V
LIST OF ABBREVIATIONS	vi
BACKGROUND	1
References	2
SECTION 1: IMPACT OF INDOOR AIR CHEMICAL POLLUTION ON CHILDREN'S HEALTH	4
Scope	4
Methods	4
Effects on the respiratory system and allergic sensitization	5
Effects on the nervous system	6
Effects on the cardiovascular system	8
Carcinogenic effects	8
Effects on the endocrine system	9
Summary	10
References	11
SECTION 2: POLLUTANTS MOST COMMONLY DETECTED IN INDOOR AIR	17
Scope	17
Methods	17
Results	18
References	21
SECTION 3: CO-OCCURRENCE OF CHEMICAL POLLUTANTS IN INDOOR AIR	25
Scope	25
Methods	25
Summary	28
References	28
ANNEX 1: DATABASE OF CHEMICAL POLLUTANTS IN INDOOR AIR IN SCHOOLS,	
KINDEDCADTENS AND DAY CARE CENTRES	35

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LIST OF ABBREVIATIONS

ASD autism spectrum disorder BDE brominated diphenyl ether

BREATH Brain Development and Air Pollution Ultrafine Particles in School Children

CAS Chemical Abstracts Service

CO carbon monoxide CO₂ carbon dioxide DBP dibutyl phthalate

DEHP di(2-ethylhexyl) phthalate

DEP diethyl phthalate
DiBP diisobutyl phthalate
DnBP di-n-butyl phthalate
DOI digital object identifier

ECA European Collaborative Action EDC endocrine-disrupting chemical

EnVIE Co-ordination Action on Indoor Air Quality and Health Effects

FEV1 forced expiratory volume

FVC forced vital capacity
HCH hexachlorocyclohexane

HESE Health Effects of School Environment

HESEINT Interventions on Health Effects of School Environment

IARC International Agency for Research on Cancer IPCS International Programme on Chemical Safety

IQ intelligence quotient
LOD limit of detection
LOQ limit of quantification

NA not applicable NO nitric oxide

NO₂ nitrogen dioxide NO_x nitrogen oxides

 O_3 ozone

OxyVOC oxygenated volatile organic compound

PAH polycyclic aromatic hydrocarbon PBDE polybrominated diphenyl ether

PCB polychlorinated biphenyl

PM particulate matter

ABBREVIATIONS contd.

PM $_{2.5}$ particulate matter with a diameter of less than 2.5 µm PM $_{10}$ particulate matter with a diameter of less than 10 µm

PPM parts per million

PRISMA Preferred Reporting Items for Systematic Reviews and Meta-Analyses

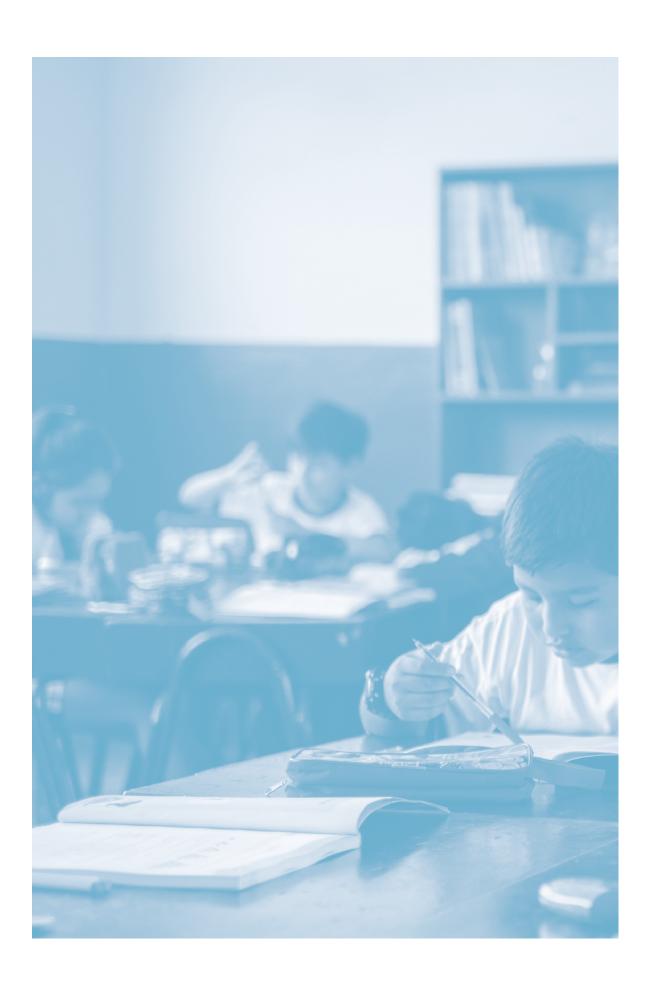
SEARCH School Environment and Respiratory Health of Children

SINPHONIE Schools Indoor Pollution and Health Observatory Network in Europe

SO₂ sulfur dioxide

SVOC semi-volatile organic compound

TRAP traffic-related air pollution VOC volatile organic compound



BACKGROUND

Children are exposed to different environmental factors in places where they learn and live that can have a profound influence on their health, growth and development. Quality of indoor air is one of the determinants of children's health and well-being, and a priority for public health (1,2).

Indoor air pollution in public settings depends on many factors: geographical conditions; penetration of outdoor air pollutants; emissions from building materials and other products used indoors; type of activities; and ventilation rate (3). A broad list of pollutants in air inside buildings has been investigated and reported in studies. Pollutants found in the air in schools and other public settings for children include particulate matter (PM), volatile and semi-volatile organic compounds (VOCs and SVOCs), polycyclic aromatic hydrocarbons (PAHs), aldehydes, musks, phthalates, inorganic chemicals (carbon dioxide (CO₂), nitrogen dioxide (NO₂), ozone (O₃)), and other organic and inorganic compounds, among which are well known allergens, irritants and carcinogens. In studies of air pollution in European schools, kindergartens and day-care centres conducted between 2012 and 2017, around 90 chemical pollutants were identified. Concentrations exceeding national and international reference values were often reported.

Evidence of linkages between health impacts and exposure to both individual chemicals and mixtures of pollutants is reported in many studies. Exposure to indoor air pollution is associated with a variety of health problems in children, including effects on respiratory, nervous and immune systems and impairment of cognitive development. It can also increase risks of health impairments later in life, including cardiovascular diseases or cancer (3–7). Links between poor indoor air quality and adverse health outcomes have been demonstrated in national and international research projects, such as School Environment and Respiratory Health of Children (SEARCH) (8), Interventions on Health Effects of School Environment (HESEINT) (9), and Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE) (3), as well as other studies conducted in the WHO European Region and globally.

Simultaneous exposure to multiple substances is of high relevance in public settings for children, given the co-occurrence of chemicals and the time spent indoors. Additive effects and higher health risks from combined exposures cannot be excluded (10), as a number of studies of combined exposure to so-called chemical families (PAHs, VOCs) and their health effects have demonstrated (11–14).

To advance assessment of risks of chemicals and to move away from a single-chemical-based approach, the WHO–International Programme on Chemical Safety (IPCS) developed a framework for assessment of risks from combined exposure to multiple chemicals (15). This framework served as the basis for developing a WHO screening tool for assessment of health risks from combined exposure to selected chemicals in indoor air in public settings for children (schools, kindergartens and day-care centres) (16).

This publication was prepared in the context of the work on the screening tool, and served as a background document to facilitate the discussions of the expert group on the priority health

outcomes and chemicals of concern (17,18). It consists of three sections, each providing information which facilitated answering the following specific questions related to combined exposure risks assessment.

- What health effects of indoor air pollution in schools and other public settings for children have been reported in epidemiological studies more commonly? A summary of evidence of negative health impacts of indoor air pollution in public settings for children is presented in Section 1.
- What chemicals are most commonly detected in air in schools, kindergartens and daycare centres? A list of chemicals for consideration by the expert group based on a review of scientific publications is presented in Section 2.
- ♦ *Is co-exposure likely given the context and timeframe?* An analysis of co-occurrence of chemicals in air in schools, day-care centres and kindergartens is presented in Section 3.

Resource- and time-efficient approaches were applied to collect information needed to answer these questions and to underpin the development of the screening tool for assessment of risks from combined exposure to multiple chemicals in indoor air in public settings for children.

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SECTION 1: IMPACT OF INDOOR AIR CHEMICAL POLLUTION ON CHILDREN'S HEALTH

The evidence of the negative health impacts of pollutants in indoor air in public settings for children is growing. In 2014 the WHO Regional Office for Europe published an overview of combined exposure to multiple pollutants in indoor air in public settings for children (1). This section summarizes the evidence collected between 2010 and 2019 of children's health effects from such exposures. It served as a scientific background for the selection of priority health endpoints in the process of the developing the screening tool (2,3).

Scope

The starting point for scoping the work and collecting information on the effects of indoor air pollution on children's health were the results of European projects, including Co-ordination Action on Indoor Air Quality and Health Effects (EnVIE) (4). Summarizing the findings of many studies, EnVIE concluded that the following diseases and symptoms are of particular interest in relation to poor indoor air quality: allergic sensitivity and asthma, lung cancer, chronic obstructive pulmonary disease, airborne respiratory infections, and irritation (sick building syndrome) (5). Many other studies also stressed the link between these health disorders and indoor air quality, including in public settings for children.

This overview therefore focuses on respiratory and immune system disorders (acute and chronic) including asthma and allergic sensitivity; cardiovascular, nervous and endocrine system dysfunctions; and carcinogenicity. It focuses on studies related to combined exposure risks of indoor air pollutants in public settings for children, and also considers publications describing effects of individual chemicals and chemical families. Given the health relevance of exposure to PM, this overview also includes evidence of the effects on children's health linked to exposure to PM in public settings.¹

Methods

The authors searched relevant publications in PubMed using the following keywords and their combinations: health risks, chemicals, chemical pollutants, cumulative risks, health effects, indoor air, schools, kindergartens, respiratory system, cardiovascular system, immune system, nervous system, neurological development, endocrine system, cancer.

In addition to evidence on health effects originating from epidemiological studies, information on adverse-effect endpoints of selected chemicals (see Section 2) from animal experiments was used to identify priority adverse-effect endpoints for grouping chemicals to assess risks from combined exposure.

¹ PM is complex mixture of organic and inorganic compounds rather than an individual chemical. As such, PM is not included in the assessment of health risks from combined exposure to multiple chemicals.

Effects on the respiratory system and allergic sensitization

Respiratory system disorders, both acute and chronic, due to indoor air pollution were reported in numerous studies of health effects from indoor air pollutants.

In the SEARCH project, both outdoor and indoor air pollutants (carbon monoxide (CO), CO_2 , particulate matter 10 µm or less in diameter (PM₁₀), formaldehyde, NO_2 and VOCs) as well as temperature and relative humidity were monitored in 247 classrooms of 63 schools in 6 European countries. Increased levels of air pollutants in combination with insufficiency of natural ventilation were associated with higher rates of chronic bronchitis and asthmatic symptoms in children (6).

A review of literature linking indoor air pollution with respiratory illnesses among school-aged children suggests that children living in households or studying in schools in urban areas are more likely to suffer from respiratory illnesses compared with children living or studying in rural areas (7).

In the SINPHONIE project, the link between indoor air pollutants and health outcomes, assessed by a questionnaire and examination of respiratory system functions, was investigated in 114 schools in 23 countries. Strong association was reported between formaldehyde levels and dry throat, nasal allergy and phlegm in children. Exposure to VOCs was linked to allergic and respiratory symptoms (limonene: irritative cough; trichloroethylene and tetrachloroethylene: nasal allergy, wheezing). Exposure to indoor NO_2 , particulate matter 2.5 μ m or less in diameter (PM_{2.5}), and O_3 was associated with irritative cough. Furthermore, the link between O_3 concentration and wheezing (ever), nasal allergy (<12 months), doctor-diagnosed nasal allergy and asthma was observed (8).

The cross-sectional European Union-funded Health Effects of School Environment (HESE) project looked at the effects of indoor air pollution on the respiratory health of schoolchildren in Denmark, France, Italy, Norway and Sweden (9). Significantly lower nasal patency was observed in schoolchildren exposed to levels of PM_{10} higher than 50 $\mu g/m^3$ than in those exposed to lower levels.

A French study (10) related concentrations of air pollution in schools to skin-prick tests, prevalence of exercise-induced and reported asthma, and allergies in 6683 children (9–11 years old) attending 108 randomly selected schools. The results revealed that asthma (exercise-induced, past year and lifetime) was strongly positively associated with indoor concentrations of benzene, sulfur dioxide (SO_2), PM_{10} , the nitrogen oxides most relevant to air pollution (NO_x) and CO, while lifetime allergic rhinitis was linked with PM_{10} , and sensitization to pollens was associated with benzene and PM_{10}

In the study of Annesi-Maesano et al. (11), an increased prevalence of past-year allergic asthma in relation to exposure to high concentrations of $PM_{2.5}$, acrolein and NO_2 in the classrooms was observed. In addition, strong positive correlation between exercise-induced asthma and concentration of $PM_{2.5}$ and acrolein was reported.

Wallner et al. (12) studied air pollution (different chemical classes, biological allergens and thermal comfort) in schools in Austria in relation to its effect on lung function. A negative association was found between airflow in the lungs of schoolchildren and presence of formaldehyde in the air, and benzyl butyl phthalate and the sum of polybrominated diphenyl ethers in dust. Concentrations of ethylbenzene, m,p-xylenes and o-xylene negatively

correlated with forced vital capacity (FVC) and forced expiratory volume (FEV1) in the first second. Chen et al. (13) also observed adverse effects on children's respiratory system, such as decreased lung function tests (FEV1 and FVC), associated with indoor air pollution.

Associations of lifetime asthma with exposure to PM and formaldehyde, rhinitis with total VOCs, and lower respiratory symptoms with formaldehyde levels, as well as potential more significant risks for children's health from combined exposure to these compounds, were also reported (1,11).

In Portugal, VOCs, aldehydes, PM_{2.5}, PM₁₀, bacteria, fungi, temperature and relative humidity levels were measured simultaneously both indoors and outdoors during a five-day period in 73 classrooms of 20 schools. In total, 978 children (8–10 years old) were involved in the study. Lung function (spirometry tests) and exhaled nitric oxide (NO) were investigated in 761 and 318 children, respectively. According to the results, despite no classrooms presenting concentrations of individual VOC pollutants exceeding the WHO indoor air quality guidelines (14) or the INDEX recommendations (15), children exposed to higher total VOC concentrations had a two-fold increased risk of having asthma-related symptoms. These findings were supported by the results of spirometry and exhaled NO tests indicating chronic airway inflammation. PM_{2.5} mass concentration levels inside and outside of day-care centres were associated with airway inflammation in preschool children, measured by fractional exhaled NO (16). Residential proximity to industrial sites associated with high VOC levels represented a significantly higher risk of not attending school due to sore throat, cough and cold compared to non-exposed groups (17).

As reported by European Collaborative Action (ECA), chemicals that might play an important role in triggering asthma symptoms include formaldehyde, aromatic and aliphatic chemical compounds, and phthalates or emissions from plastic materials (18). Indoor chemical products resulting from ozonolysis of terpenes may also play a role, but the evidence is more limited (18). NO_2 , PM, VOCs and formaldehyde in combination with dust mites, bacteria and fungal endotoxins may also increase the risk of development of asthma and wheeze in children (19). Higher $PM_{2.5}$ and PM_{10} mass concentration levels increased the odds of asthma-like symptoms (20). Exposure to PAHs has also been linked to asthma development and exacerbation through promoting oxidative stress and immune responses (21).

In summary, the most frequently reported adverse-effect endpoints of indoor air pollutants in the respiratory system of children include allergic rhinitis, asthma development or exacerbation (expressed as reported past-year or lifetime asthma, exercise-induced asthma, hospital admissions and emergency visits for asthma, asthma-like symptoms, wheezing, increased bronchial responsiveness), chronic airway inflammation (expressed as decreased lung function, decreased index of FEV1 and FVC detected by spirometry, enhanced exhaled NO), chronic bronchitis, and acute respiratory infections (pneumonia hospitalization).

Effects on the nervous system

Development of the nervous system (proliferation, migration, differentiation, myelinization of neurons, synaptogenesis and regulated apoptosis) extends from the embryonic period through adolescence (22). This is a critical developmental window, when exposure to chemicals including from those in indoor air can result in neurodevelopmental disorders.

Known neurotoxic chemicals detected in indoor air are certain VOCs and PAHs. VOCs investigated in schools were found to be risk factors for the development of sick building

syndrome, characterized by headache and fatigue (23). Exposure to PAHs, in particular to benzo[a]pyrene, in the school environment during preadolescent school age was associated with subclinical changes on the caudate nucleus revealed in magnetic resonance imaging. This can play a crucial role in many cognitive and behaviour processes (24).

In the Brain Development and Air Pollution Ultrafine Particles in School Children (BREATH) cohort study in Barcelona, Spain, conducted in 39 schools exposed to high or low traffic-related air pollution (TRAP), 2715 children 7–10 years old were tested four times via computerized tests. Air pollution monitoring (elemental carbon, NO_2 and ultrafine particles with a diameter of 10–700 nm) was conducted outdoors and indoors in two separate one-week campaigns in each school. Cognitive development was assessed through working memory and inattentiveness tests. After adjusting to age, sex, maternal education, socioeconomic status and air pollution exposure at home, the study found that children from the schools with higher levels of air pollution had significantly lower growth of cognitive development during 12 months than children from the schools with lower levels of pollution (25). This negative association persisted during a three-and-a-half-year period (26). In another study based on the BREATH cohort, annual average concentrations of PAHs, elemental carbon and NO_2 at the children's schools were associated with smaller caudate nuclei volumes and higher scores of behavioural disorders.

The observed effects depend on other characteristics (genetic) of children: lower improvement in inattentiveness over time in children carrying the *APOE* ε4 allele than in other children suggests that ε4 carriers may be more vulnerable to adverse neurobiological effects of TRAP exposure than non-carriers. That means that genetic factors could modify cognitive and other brain functions in response to exposure to indoor air pollutants (27).

Suades-Gonzalez et al. (28) reviewed 31 epidemiological studies of exposure to air pollutants and neuropsychological development outcomes (delay of cognitive and psychomotor development, intelligence quotient (IQ), learning disabilities, reading comprehension, memory functions, speed of reactions, attention, coordination, perceptual coding, span length) and behaviour outcomes (attention disturbance, autistic traits, autism spectrum disorder (ASD)). Sufficient evidence of relations between pre- or postnatal exposure to PAHs and decreased IQ and between exposures to $PM_{2.5}$ and ASD was reported, whereas limited evidence of linkages between NO $_{\circ}$ and ASD were observed.

A study conducted in China looked for linkages between indoor air concentrations of NO_2 and PM_{10} in a school located in an area with severe TRAP and a school with predicted low traffic density based on ambient air quality monitoring data and neurodevelopment of children (282 pupils were examined). In multiple ordinal logistic regression models after adjusting for other covariates (household pollution, demographics, birth weight, delivery method, breastfeeding, vision, familiarity with computer games) indoor TRAP exposure was significantly associated with poorer performance on neurobehavioural tests (29).

According to several epidemiological and animal studies, early-life exposure to phthalates, bisphenol A, polychlorinated biphenyls (PCBs), polybrominated diphenyl ethers (PBDEs) and PAHs (contaminants that are often present in indoor air) may result in neurodevelopmental deficits, either via oxidative stress and neuroinflammation or interference with hormones during neural development (30–33).

In summary, exposure to indoor air pollutants can negatively affect neuropsychological development measured as cognitive and psychomotor development, IQ, learning abilities,

reading comprehension, memory functions, reading and math scores, reaction speed, attention, coordination, perceptual coding and span length.

Effects on the cardiovascular system

The evidence on the relationship between long-term exposure to air pollution, in particular to PM and cardiovascular mortality, is well established based on studies in adult populations. Much less is known about the association between air pollution and changes in cardiovascular parameters in children.

The few studies performed in the school environment (34,35) confirm that exposure to indoor air pollutants in homes and schools might contribute to the development of long-term health effects in the cardiovascular system. For example, later-life hypertension can correlate with higher blood pressure in early life. A review of the studies published between 2007 and 2017 that examined prenatal or childhood exposure to PM_{10} and $PM_{2.5}$, NO_x , O_3 , black carbon, environmental tobacco smoke and phthalates reported associations between exposure to these pollutants and increased childhood blood pressure (36).

Sughis et al. (35) investigated blood pressure in children at schools located in areas characterized by low and high pollution. The study concluded that children living and attending school in an area of very high TRAP have a substantially higher arterial blood pressure compared to less exposed children. A randomized, double-blind crossover trial among healthy college students (37) also demonstrated that reduction of PM through the use of air purifiers leads to significant decreases of levels of stress hormones (cortisol, cortisone, epinephrine and norepinephrine), blood pressure, insulin resistance, and biomarkers of oxidative stress and inflammation. Children attending schools near an oil refinery with high PAH levels in ambient air had significantly higher systolic and diastolic blood pressure Z-scores than those of children attending schools with low PAH levels, with a 4.36-fold increase in prehypertension (34).

In summary, there is evidence of functional changes to the cardiovascular system in schoolchildren associated with exposure to indoor air pollutants.

Carcinogenic effects

A growing body of research on linkages between indoor air pollutants and carcinogenicity risk suggests that early-life exposures may contribute to cancer development later in life.

Several VOCs found in indoor air, such as benzene, trichloroethylene and formaldehyde, have been classified as known human carcinogens (Group 1) by the International Agency for Research on Cancer (IARC) based on evidence from epidemiologic studies and animal data (38), while styrene and tetrachloroethylene have been classified as possible or probable carcinogen for humans, respectively (39). Benzene is toxic to the bone marrow and is associated with various haematological cancers (40). Formaldehyde may cause nasal cancer and leukaemia (38). Among PAHs, benzo[a]pyrene has been classified as a known human carcinogen, and naphthalene, chrysene, benzo[a]anthracene, benzo[k]fluoranthene and benzo[b]fluoranthene have been classified as possible carcinogens (41). They may induce cancer tumours, primarily in the lungs, skin, bladder, liver and stomach.

Sarigiannis et al. (42) demonstrated that in Europe, indoor VOCs and other compounds, including benzene, formaldehyde, toluene, xylenes, styrene, acetaldehyde, naphthalene,

limonene, alpha-pinene and ammonia, originate mostly from indoor sources (primary indoor pollutants). Cancer risks posed by these chemicals due to indoor exposure were estimated to be up to three orders of magnitude higher than one in 1 million (the threshold of what is considered acceptable).

Young children living close to busy road can be at increased risk of developing leukaemia (43,44). Two recent meta-analyses suggested a link between ambient exposure to TRAP during childhood and leukaemia risk in children (45,46). Several indoor and outdoor hazardous pollutants may play a role in the etiology of childhood leukaemia (47). Children (infants, children 6 and 15 years old) exposed to toxic substances in the air (for example, acetaldehyde, 1,3-butadiene, benzene, ortho-dichlorobenzene, toluene and PAHs) may have increased risk of certain brain tumours (48,49).

A number of studies on exposure to air pollutants in schools and cancer risks, estimated based on child-specific benchmark levels, have been published during last decade.

Chan et al. (50) modelled the risks of cancer from exposures to VOCs in schools in the United States of America. The estimated cancer risks were about 10 per million, which was lower than the risk of cancer predicted for exposures to VOCs in households. Based on investigation of 38 VOCs in 34 early-childhood educational environments, Hoang et al. (51) concluded that child exposures to benzene, chloroform, ethylbenzene and naphthalene exceeded age-adjusted benchmark levels based on California's Proposition 65 guidelines (10-5 lifetime cancer risk) in 71%, 38%, 56%, and 97% of facilities, respectively. Sofuoglu et al. (52) investigated indoor concentrations of VOCs in Turkish primary schools and determined that formaldehyde was the substance of particular concern in terms of high chronic toxic and carcinogenic risks, followed by naphthalene, benzene and toluene.

In summary, some studies revealed the associations between exposure to certain pollutants in indoor air and risks of childhood leukaemia and some central nervous system tumours.

Effects on the endocrine system

Children are exposed to endocrine-disrupting chemicals (EDCs) through contaminated drinking-water and food, through dermal contact with these chemicals, and through breathing polluted air indoors. EDCs are present in indoor environments mostly in particulate forms and in dust. Phthalates, bisphenol A, PBDEs and PCBs, environmental phenols, dioxins, PAHs and perfluorocarbons often occur in indoor air of homes and classrooms (53–55). Data collected by Oziol et al. (56) confirmed the presence of EDCs in a gaseous state and highlighted their indoor origin and concentration, especially in the cold season.

Evaluating the contribution of indoor air quality in residences to daily intake of phthalates in Japanese children, Yoshida et al. (57) concluded that inhalation did not seem to contribute very much as an absorption pathway of the phthalates. A study by Bekö et al. (58) of inhalation, dust ingestion and dermal absorption of diisobutyl phthalate (DiBP), benzyl butyl phthalate and di(2-ethylhexyl) phthalate (DEHP) in 431 Danish children 3–6 years old resulted in a different conclusion: exposures to phthalates present in the air and dust indoors meaningfully contribute to a child's total intake of certain phthalates.

Using a cumulative risk-assessment approach for evaluating the contribution of indoor air pollution by phthalates, Fromme et al. (59) investigated phthalates in indoor air and dust in

German day-care centres. Estimates using a cumulative risk-assessment approach for the sum of DEHP, di-n-butyl phthalate (DnBP) and DiBP, revealed that 20% of the children had concentrations exceeding the hazard index of one. However, the contribution of different phthalates in indoor air and dust varied significantly (24–70%). Mechanisms of action, adverse effects and dose–response relationships between exposure to these chemicals from air (indoor and outdoor) are poorly understood. As no systematic screening of common chemicals for endocrine-disrupting effects is currently underway, questions remain as to the health impacts of these exposures (60).

Unfortunately, until now, evidence of health effects in children from pollution of indoor air in public settings is very limited and relates mainly to obesogenic effects of EDCs (61,62). In Portugal, 815 children in 20 schools in Porto were involved in an investigation of exposure to EDC and the prevalence of asthma and obesity (61). The results confirmed that even low levels of indoor exposure may influence the risk of asthma, respiratory symptoms and obesity.

In summary, EDCs in both air and dust can contribute to overall exposure. However, the evidence of health effects in children from exposure to EDCs in schools and day-care centres is limited.

Summary

This overview of the literature on health outcomes associated with indoor air pollution was prepared to support discussion and selection of priority adverse-effect endpoints for inclusion in the screening tool. In this context, the evidence was considered consistent for the association between air pollution and respiratory system effects (most often allergic rhinitis, asthma development or exacerbation, chronic airway inflammation, and acute respiratory infections) and nervous system effects (impairments in different neuropsychological development outcomes or effects on the nervous system observed by neuroimaging) in children.

Evidence of potential cardiovascular effects in children (for example, elevated blood pressure and heart rate, higher levels of stress hormones and biomarkers of oxidative stress) is more limited and less consistent.

Data on potential effects on the immune system of children are limited (except for asthma and allergic rhinitis described above), varied and difficult to interpret, due in part to the commonly nonspecific nature of the endpoints considered (for example, infections such as pneumonia or otitis media; absenteeism from school due to sore throat, cough and cold; development of allergies).

The following priority adverse-effect endpoints were identified for inclusion in the screening tool based on this literature review and through expert consultations (2,3):

- respiratory
- ♦ cardiovascular
- ♦ nervous
- irritation
- carcinogenicity.

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SECTION 2: POLLUTANTS MOST COMMONLY DETECTED IN INDOOR AIR

A list of chemical pollutants in indoor air is broad: VOCs, SVOCs, PM, phthalates, musks, PAHs, brominated flame retardants, organophosphate flame retardants, chlorinated paraffins and inorganic compounds. These pollutants originate from outdoor and indoor sources, and their concentrations indoors are determined by many factors and vary significantly. Given the high health relevance of indoor air pollution, national monitoring programmes/studies have been conducted in many countries. A large number focus on chemicals expected to be released from the known indoor sources, such as VOCs and SVOCs. Given the unavoidable penetration of outdoor pollutants into indoor environments, studies also commonly include an assessment of the main pollutants of outdoor air (PM, NO, PAHs, etc.) (1,2).

Scope

This literature review aimed at creating a list of chemicals most commonly detected in indoor air in public settings for children (schools, kindergartens and day-care centres), based on the review of studies conducted in the WHO European Region and published from 2012. The list was shared with the expert group to consider chemicals of concern for inclusion in the WHO screening tool for assessment of health risks from combined exposure to hazardous chemicals in indoor air, referred to earlier in this publication (3,4).

Methods

Priority was given to the published reviews: Malliari (5), Morawska (6) and Annesi-Maesano (SINPHONIE) (7). Regarding the SINPHONIE project, results from the final report from 2014 were retrieved, with a total of 114 schools and 342 classrooms investigated across Europe. Peer-reviewed journal articles and conference papers were sought using Google Scholar, Scopus and Science. In total, more than 150 papers were reviewed.

The keywords for the sampling locations were "school" OR "kindergarten" OR "nursery" OR "day care". The keywords regarding the target substances were "pollutant" OR "particle" OR "substance". Overall, the keywords for the search were "indoor air quality" combined with (AND) the sampling location keywords and target substance keywords. For the purpose of this review, measurements in settled dust were excluded.

Several rejection criteria were defined for the overview:

- ◆ a substance-type criterion: bio-contaminants, radiation and tobacco smoke were excluded;
- ♦ a geographical criterion: only studies performed in European countries were retrieved; and
- ♦ a temporal criterion: only studies published between 2012 and 2017 were retrieved, including reviews dwelling on a much broader time period.

Among the more than 150 publications reviewed, 26 were selected (1,2,5–28) based on the following exclusion criteria.

- Among the selected articles, different words referred to public settings for children, such as elementary school, primary school, nursery school, kindergarten or day-care centre. Studies involving universities or engineering schools were excluded, as were those involving high and middle schools.
- Studies originating from specific contexts, for example, particle characterization near schools where the uncontrolled combustion of a tire landfill took place, or a school near a steel factory, were excluded as they may misrepresent the common situation depicted in this publication.
- ♦ Five studies that took place in facilities not used by children (for example, teaching rooms or laboratories at a university) were also excluded.

Regarding data compilation (see Annex 1), several parameters were retrieved from the articles when available:

- measured substances
- corresponding Chemical Abstracts Service (CAS) registration number
- ◆ type of building investigated (school, kindergarten or day-care centre)
- number of buildings investigated
- number of rooms (in each building or in total)
- measurement unit
- measurement data: minimum, maximum, arithmetic mean and median
- country where the measurements took place
- year when the measurements took place
- reference information: year of publication, authors, title and digital object identifier (doi).

In several cases, studies did not report all the requested data (measurement results, number of sampling location, sampling year, etc.). In order not to be too restrictive, the articles were considered as long as at least one of the following measurement values was given: minimum, maximum, arithmetic mean or median.

If data were not provided or were reported too partially (that is, distribution P_{25} , P_{75} and nothing else), or if only plots were given without thresholds or figure descriptions, they could not be considered for the compilation.

Lastly, some results were properly given but combined with other types of buildings, such as offices and/or homes. If it was not possible to obtain separate values specifically for schools, kindergartens or day-care centres, data were not considered.

Results

As a result, the search identified 90 substances prioritized by frequency of detection (see Table 1).

Table 1. List of the most commonly detected indoor air pollutants in public settings for children

No.	Substance (alphabetical order)	CAS number	No.	Substance (alphabetical order)	CAS number
1	1-butanol	71-36-3	34	CO	630-08-0
2	1,2,3-trimethylbenzene	526-73-8	35	d-limonene	5989-27-5
3	1,4-dichlorobenzene	106-46-7	36	Di-isodecyl phthalate	26761-40-0
4	4-(1,2-dibromoethyl)-1,2-dibromocyclohexane	3322-93-8	37	DEHP	117-81-7
5	4,4'-dichloro- diphenyltrichloroethane	50-29-3	38	Dibenz[a,h]anthracene	53-70-3
6	Acenaphthene	83-32-9	39	Dibenzo[a,l]pyrene	191-30-0
7	Acenaphthylene	208-96-8	40	DiBP	84-69-5
8	Acetaldehyde	75-07-0	41	Diethyl phthalate (DEP)	84-66-2
9	Alpha hexachlorocyclohexane (HCH) (or α-HCH)	319-84-6	42	Diisononyl phthalate	28553-12-0
10	Anthracene	120-12-7	43	Dimethyl phthalate	131-11-3
11	Arsenic	7440-38-2	44	DnBP	84-74-2
12	Brominated diphenyl ether (BDE) 28	41318-75-6	45	Ethylacetate	141-78-6
13	BDE 47	5436-43-1	46	Ethylbenzene	100-41-4
14	BDE 99	60348-60-9	47	Fluoranthene	206-44-0
15	BDE 100	189084-64-8	48	Fluorene	86-73-7
16	BDE 153	68631-49-2	49	Formaldehyde	50-00-0
17	BDE 183	207122-16-5	50	Galaxolide	1222-05-5
18	BDE 209	1163-19-5	51	Gamma HCH (or γ-HCH)	134237-52-8
19	Benz[a]anthracene	56-55-3	52	Heptane	142-82-5
20	Benzene	71-43-2	53	Hexabromobenzene	201-773-9
21	Benzo[a]pyrene	50-32-8	54	Hexaldehyde	66-25-1
22	Benzo[b+ j]fluoranthene	205-99-2/205- 82-3	55	Indeno[1,2,3-cd]pyrene	193-39-5
23	Benzo[b+j+k]fluoranthene	205-99-2/205- 82-3/207-08-9	56	Limonene	138-86-3
24	Benzo[e]pyrene	192-97-2	57	m,p-xylenes	108-38-3/106- 42-3
25	Benzo[ghi]perylene	191-24-2	58	Methylacetate	79-20-9
26	Benzo[k]fluoranthene	207-08-9	59	n-butylbenzene	104-51-8
27	Benzyl butyl phthalate	85-68-7	60	n-decane	124-18-5
28	Butyl acetate	123-86-4	51	Naphthalene	91-20-3
29	Butyraldehyde	123-72-8	62	Nickel	7440-02-0
30	Cadmium	7440-43-9	63	NO ₂	10102-44-0
31	Camphene	79-92-5	64	o-xylene	95-47-6
32	Carbon	124-38-9	65	O ₃	10028-15-6
33	Chrysene	218-01-9	66	Particles (number)	-

Table 1 cont.

No.	Substance (alphabetical order)	CAS number	No.	Substance (alphabetical order)	CAS number
67	PCB 28	7012-37-5	79	Styrene	100-42-5
68	PCB 31	16606-02-3	80	Tetrachloroethylene	127-18-4
69	PCB 52	35693-99-3	81	Toluene	108-88-3
70	PCB 101	37680-73-2	82	Tonalide	21145-77-7
71	PCB 138	35065-28-2	83	Total suspended particles	_
72	Permethrin	52645-53-1	84	Tri-(2-butoxyethyl)- phosphate	78-51-3
73	Phenanthrene	85-01-8	85	Tributylphosphate	126-73-8
74	PM ₁	_	86	Trichloroethylene	79-01-6
75	PM _{2.5}	_	87	Tris(1-chloro-2-propyl) phosphate	13674-84-5
76	PM ₁₀	_	88	Tris(2-chloroethyl) phosphate	115-96-8
77	Pyrene	129-00-0	89	Ultrafine particles	_
78	SO ₂	7446-09-5	90	α-pinene	80-56-8

To reduce the list of substances to consider, a prioritization process was applied following different criteria that fit the objectives and scope of the study:

- preselection based on WHO guidance and existing documents (that is, guidelines for indoor or ambient air quality);
- number of studies and measurements in European Union schools over the past years to spot the most common chemical pollutants; and
- expert judgment regarding toxicological or exposition potential.

A final list of 30 substances measured in 26 countries was considered for the selection of chemicals for inclusion in the screening tool for assessment of health risks from combined exposure to multiple chemicals in indoor air, based on frequency of their recording (see Table 2).

Table 2. List of 30 chemicals selected for further discussion

No.	Substance	Number of recordings	No.	Substance	Number of recordings
1	Formaldehyde	24	17	Limonene	4
2	NO ₂	20	18	n-decane	4
3	PM ₁₀	19	19	Acetaldehyde	3
4	PM _{2.5}	17	20	BDE 47	3
5	Benzene	11	21	Benzo(a)pyrene	3
6	O ₃	10	22	Naphthalene	3
7	CO	7	XX	o-xylene*	3
8	m,p-xylenes	7	XX	d-limonene*	2
9	Styrene	7	23	Phenanthrene	2
10	Toluene	7	24	Trichloroethylene	2
11	1,4-dichlorobenzene	6	25	DEP	2
12	Ethylbenzene	6	26	DiBP	2
13	Butyl acetate	5	27	DnBP	2
14	Tetrachloroethylene	5	28	Galaxolide	1
15	α-pinene	5	29	SO ₂	1
16	1,2,3-trimethylbenzene	4	30	Tonalide	1

^{*} The two substances in grey font (o-xylene and d-limonene) are accounted respectively with m,p-xylenes and limonene.

The process for the selection of priority chemicals is described in the reports on the first and the second expert consultations (3,4). The final list of chemicals included in the screening tool is also available in the publication on methods for sampling and analysis of chemicals (29).

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SECTION 3: CO-OCCURRENCE OF CHEMICAL POLLUTANTS IN INDOOR AIR

According to the WHO–IPCS framework (1), the potential impact of the co-occurrence of, and concomitant exposure to, multiple chemicals should always be taken into account in problem formulation for combined exposures risk assessment. A critical question in that context is whether or not the co-occurrence of chemicals is likely within a relevant timeframe.

Scope

Relevant literature was reviewed to answer the following question: *Do chemicals co-occur in indoor air in schools, kindergartens and day-care centres, based on the review of information on chemical pollutants in indoor air found in public settings for children?*

In the review, the frequency of detection was used as the best way to identify whether pollutants are present simultaneously in air samples. If the frequency of detection for a pollutant is lower than 100%, it means that it is not systematically present in the indoor air, according to the sampling and analytical methods used.

Methods

Peer-reviewed journal articles and conference papers were sought in the PubMed, Science Direct and Google Scholar search engines, with publication dates from 2014 until 30 October 2018 and without geographical limitation. Studies that presented data for at least two chemicals measured simultaneously in indoor air in schools (preschools or elementary schools) and day-care centres or kindergartens were analysed.

The Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) approach (2) was used to identify selected studies.

The three following search strings were used in the search engines.

- ("Indoor air" OR "indoor air quality") AND ("air pollutants") AND ("schools" OR "classroom" OR "preschools" OR "day-care center" OR "nursery")
- 2. ("indoor air" OR "indoor air quality") AND ("multipollutants" OR "multi-pollutants" OR "multi-pollution") AND ("schools" OR "classroom" OR "preschools" OR "day-care center" OR "nursery")
- 3. ("indoor air" OR "indoor air quality") AND ("air pollutants") AND ("co-occurrence") AND ("schools" OR "classroom" OR "preschools" OR "day-care center" OR "nursery").

Several parameters were retrieved from the articles:

- ◆ reference information: year of publication, authors, title, journal or book doi, etc.
- year when the measurements took place
- ◆ country where the measurements took place
- ◆ type of building investigated (that is, school, kindergarten or day-care centre)

- number of buildings investigated
- number of rooms (in each building or in total)
- number of air samples
- measured compounds
- frequency detection of the measured compounds (%)
- corresponding CAS registration number.

The frequencies of detection of each measured chemical were recorded to preview their simultaneous presence in air samples (see Table 3).

Table 3. Number of measured compounds in the reviewed studies according to chemical family

Chemical family (alphabetical order)	Number of measured compounds	Chemical family (alphabetical order)	Number of measured compounds
VOCs		SVOCs	
Alcohols	4	Brominated flame retardants	28
Aldehydes	15	Organophosphates	8
Alkanes	14	PAHs	19
Aromatic hydrocarbons	12	PCBs	9
Ester alcohols	2	Perfluorinated compounds	11
Esters	2	Phthalates	7
Glycol ether	1	Pyrethroids	5
Halogenated hydrocarbons	6	Musks	2
Ketones	4	Non-phthalate plasticizers	8
Siloxanes	3	Organochlorine pesticides	7
Terpenes	5		

Some studies did not report all the requested data (detection frequencies, number of sampling location, sampling year, etc.). If detection frequencies were not reported but the distribution (P_{25} , median P_{75}) and limit of detection were mentioned, an estimate of frequency (a range) of detection from the distribution was indicated in the file. If detection frequencies were not provided, studies could not be considered for the compilation.

Lastly, some results were properly given but combined with other types of building such as offices and/or homes. If it was not possible to obtain separate values specifically for schools, data were not considered.

In total, 1656 studies were identified in the search.

 ("Indoor air" OR "indoor air quality") AND ("air pollutants") AND ("schools" OR "classroom" OR "preschools" OR "day-care center" OR "nursery")

a. PubMed: 66 references

b. Science Direct: 536 referencesc. Google Scholar: 1000 references

("Indoor air" OR "indoor air quality") AND ("multipollutants" OR "multi-pollutants" OR "multi-pollution") AND ("schools" OR "classroom" OR "preschools" OR "day-care center" OR "nursery")

a. PubMed: 2 references

b. Science Direct: 21 referencesc. Google Scholar: 13 references

3. ("Indoor air" OR "indoor air quality") AND ("air pollutants") AND ("co-occurrence") AND ("schools" OR "classroom" OR "preschools" OR "day-care center" OR "nursery")

a. PubMed: 0 references

b. Science Direct: 5 referencesc. Google Scholar: 13 references

The first step of the reviewing process was to screen the titles and abstracts of these studies. As a result, 1585 papers were excluded according to the exclusion criteria mentioned above. Seventy-three studies were identified for full-text screening (3–76).

The second step was to remove irrelevant studies. Thirty-seven studies were removed because frequencies of detection or distributions were not provided, and five were removed because they took place in facilities not used by children (teaching rooms or laboratories at a university).

The number of selected studies published since 2014 was two times higher in 2016 and 2017 than in the other years (see Fig. 1). Twenty studies were carried out in schools (elementary or nursery) and 10 in day-care centres or children's facilities in 16 countries (Canada, China, France, Germany, Greece, Iran, Japan, Korea, Malaysia, Norway, Poland, Portugal, Serbia, Spain, Sweden and the United States of America).

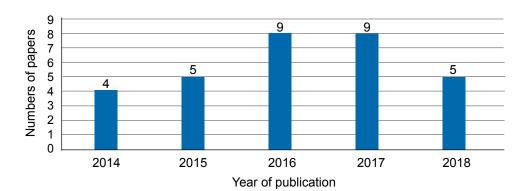


Fig 1. Number of publications per year

The most measured chemical family was VOCs, which were included in 15 studies. This was followed by aldehydes (11 studies), PAHs (6 studies) and flame retardants (6 studies). In total, 177 pollutants were considered (see Table 3).

Summary

At least two chemicals were detected in all reviewed studies. No specific pairs or group of chemicals were identified. However, the likelihood of co-occurrence of multiple chemicals in indoor air in schools, kindergartens and day-care centres is high. This is a strong argument in favour of applying the WHO framework for assessment of risks from combined exposure to multiple chemicals (1) in indoor air in public settings for children.

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ANNEX 1: DATABASE OF CHEMICAL POLLUTANTS IN INDOOR AIR IN SCHOOLS, KINDERGARTENS AND DAY-CARE CENTRES

Table A1. Chemical pollutants in schools, kindergartens and day-care centres $(2010-2017)^2$

Reference			Annesi-Maesano I, Baiz N, Banerjee S, Rudnai P, Rive S & on behalf of the SINPHONIE Group. Indoor	air quality and sources in schools and related health effects. J	Toxicol Environ Health B Crit Rev.	_ 2013;16:8:491–550. doi:10.1080/10 _ 937404 2013 853609				Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Indoor air quality in preschools (3- to 5-year-old children) in the northeast of Portugal during spring-summer season: pollutants and comfort parameters. J Toxicol Environ Health A. 2017;80(13–15):740–55. doi:10.1080/15287394.
Maximum Location Sampling year	/des		V V	NA	NA	NA	NA	NA	¥	2013
Location	s) – aldehy		A A	Ą	NA	NA	NA	Ą	Ϋ́	Portugal
Maximum	Oxygenated volatile organic compounds (oxyVOCs) - aldehydes		55	I	85.8	68.04	56.62	76.2	47.3	0.00
Median	compound		28	21.5	4.8	I	I	I	ı	I
Arithmetic Median mean	ile organic		I	ı	7.8	35.13	30.34	40.04	ı	0.035
Minimum	nated volat		12	ı	0.3	2.21	4.05	3.81	12.9	0.01
Unit	Oxyge		µg/m³	µg/m³	µg/m³	µg/m³	µg/m³	hg/m³	hg/m³	mg/m³
Number of rooms		-00-0)	NA	NA	NA	NA	NA	NA	AN	1 playroom 1 classroom
Number of buildings		Formaldehyde (CAS 50-00-0)	Not applicable (NA)	ΑĀ	NA	NA	NA	ΑN	ΑĀ	2
. Type of building		Formalder	School	School	School	School	School	School	School	Nursery school
o N		_								

² Chemicals agreed at the first expert consultation (3–4 December 2018) are included in the database.

Table A1 cont.

Number Number Officers Of	Sampling Reference year		2009/2010 Ramalho O, Wyart G, Mandin C, Blondeau P, Cabanes P, Leclerc N et	 a. Association of carbon dioxide with 2009/2010 indoor air pollutants and exceedance of health guideline values. 	2009/2010 Build Environ. 2015;93:115–24. doi:10.1016/j.buildenv.2015.03.018.	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.12222.	2011/2012 European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.	2011/2013 Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk
of pulldings Number of rooms of rooms Unit of rooms Minimum mean mean mean mean mean mean mean mea								
Number Number Unit Minimum Arithmetic	Maximum		181	86	70	66.2	99	126.9
of number of rooms buildings Number of rooms of rooms of rooms buildings Unit Minimum of rooms of rooms buildings Idehyde (CAS 50-00-0) 46 1 or 2 in µg/m³ 2.5 each each rooms 2.5 each rooms ach each rooms 1.6 each rooms Idehyde (CAS 50-00-0) 53 1 or 2 in µg/m³ 6.8 each rooms 1.6 each rooms 1.6 each rooms Idehyde (CAS 50-00-0) 17 51 µg/m³ 6.8 1.6 each rooms 1.6 each rooms Idehyde (CAS 50-00-0) 17 51 µg/m³ 8.24 1	Median		12	19	17	19.2	5	17.5
Number Number Unit of rooms Unit of rooms	Arithmetic mean		41	22	19	25.1	5	19.8
Idehyde (CAS 50-00-0) Idehyde (CAS 50-00-0) / 46 1 or 2 in each garten 61 1 or 2 in each tary 53 1 or 2 in each // 17 51 tary 114 300	Minimum		2.5	2	1.6	<u>ω</u>	_	8.24
of buildings ldehyde (CAS 50-00 / 46 / 46 / 17 / 17 tary 53 / 114	Unit		hg/m³	hg/m³	hg/m³	µg/m³	µg/m³	µg/m³
ling of buildings buildings all buildings and buildings ary 46 old argarten 61 antary old 114	Number of rooms	(0-00	1 or 2 in each	1 or 2 in each	1 or 2 in each	51	300	72
ing ling sary argarten entary ol	Number of buildings	Je (CAS 50-	46	61	53	11	41.	20
Nurse school Sch	Type of building	Formaldehyd	Nursery school	Kindergarten	Elementary school	Nursery/ elementary school	School	School

Table A1 cont.

O	Type of building	Number of buildings	Number of rooms	Unit	Minimum	Arithmetic Median mean	Median	Maximum	Location	Sampling year	Reference
7	Acetaldehyo	Acetaldehyde (CAS 75-07-0)	0-20								
	Nursery/ elementary school	17	15	hg/m³	2.7	6.3	6.1	10.7	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65.
	School	20	72	µg/m³	1.92	9.31	7.65	64.6	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j.buildenv.2015.11.031.
				Volat	ile organic	Volatile organic compounds (VOCs) - aromatic hydrocarbons	s (VOCs) -	- aromatic h	ydrocarbo	sus	
က	Benzene (CAS 71-43-2)	AS 71-43-2)									
'	Nursery school	46	1–2 per building	µg/m³	0.7	2.5	2.3	6.5	France	2009/2010	Ramalho O, Wyart G, Mandin C, Blondeau P, Cabanes P, Leclerc N et
	Kindergarten	61	1–2 per building	hg/m³	9.0	2.4	2.2	7.9	France	2009/2010	 a. Association of carbon dioxide with indoor air pollutants and exceedance of health quideline values.
'	Elementary school	53	1–2 per building	µg/m³	0.2	2.5	2.2	21.7	France	2009/2010	Build Environ. 2015;93:115–24. doi:10.1016/j.buildenv.2015.03.018.
	Nursery	-	2	hg/m³	1	2.745	ı	I	Czechia	ı	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-899X/245/8/082027.

Table A1 cont.

o V	Type of building	Number of buildings	Number of rooms	Chiit	Minimum	Arithmetic Median Maximum Location mean	Median	Maximum	Location	Sampling year	Reference
3	Benzene (C	Benzene (CAS 71-43-2)									
	Nursery/ elementary school	17	15	mg/m³	< Limit of quantitation (LOQ)	2.1	4.	8.5	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.1222.
	School	41	300	°μ/bπ	< Limit of detection (LOD)	4	2	38	Europe	2011/2012	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.
,	Elementary school	1	ဇ	µg/m³	0.55	I	I	0.65	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
,	Elementary school	1	ဇ	µg/m³	0.47	I	I	1.09	Italy	I	outdoor monitoring of volatile organic compounds in school buildings: indicators based on health risk
	Elementary school	1	ဇ	µg/m³	0.11	I	I	0.25	ltaly	I	assessment to single out critical issues. Int J Environ Res Public
	Elementary school	1	ဇ	hg/m³	0.04	I	I	0.11	ltaly	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.
	School	50	72	°µ/br	ਨ.	2.2	25.5	2.7	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.

Table A1 cont.

Ö	Type of building	Number of buildings	Number of rooms	Chit	Minimum	Arithmetic mean	Median	Maximum	Location	Sampling year	Reference
4	Ethylbenzer	Ethylbenzene (CAS 100-41-4)	-41-4)								
	Nursery/ elementary school	17	12	hg/m³	1.2	2.2	7.7	ဖ	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.1222.
	Nursery	-	2	µg/m³	1	6.57	ı	1	Czechia	₹ Z	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-899X/245/8/082027.
	Elementary school	_	က	hg/m³	0.5	I	I	0.62	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
	Elementary school	~	က	hg/m³	0.53	I	I	1.53	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
	Elementary school	←	က	hg/m³	0.16	I	I	0.28	Italy	I	assessment to single out critical issues. Int J Environ Res Public
	Elementary school	~	က	hg/m³	0.19	I	I	0.22	Italy	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.
2	1,2,3-trimeth	1,2,3-trimethylbenzene (CAS 526-73-8)	3AS 526-73	(8-							
	Elementary school	~	က	րց/m³	0.39	l	I	61	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor
	Elementary school	~	က	րց/m³	0.37	I	I	0.86	Italy	I	and outdoor monitoring of volatile organic compounds in school
	Elementary school	-	င	րց/m³	0.11	I	I	0.18	Italy	I	health risk assessment to single out critical issues. Int J Environ Res
	Elementary school	~	က	hg/m³	0.14	ı	I	19	Italy	I	Public Health. 2013;10:6273–91. doi:10.3390/ijerph10126273.

Table A1 cont.

Reference		Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.12222.	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-899X/245/8/082027.	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and	outdoor monitoring of volatile organic compounds in school buildings:	assessment to single out critical issues. Int J Environ Res Public	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.
Sampling year		2010	₹ Z	1	I	I	I	2011/2013
Location		France	Czechia	Italy	Italy	Italy	Italy	Portugal
Maximum Location		14.9	1	2.01	5.84	0.82	0.59	365.2
Median		2.8	I	I	I	I	I	വ
Arithmetic Median mean		4.	1.825	I	I	l	l	17.7
Minimum		9.	ı	1.58	1.66	0.48	0.56	1.2
Unit	12-3)	µg/m³	µg/m³	րց/m³	hg/m³	րց/m³	րց/m³	µg/m³
Number Unit of rooms	-38-3/106-4	12	2	က	က	က	က	22
Number of buildings	s (CAS 108	17	-	_	-	~	~	50
Type of building	m-,p-xylenes (CAS 108-38-3/106-42-3)	Nursery/ elementary school	Nursery	Elementary school	Elementary school	Elementary school	Elementary school	School
No.	9		ı	I	I		I	1

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Table A1 cont.

	lype of building	Number of buildings	Number Unit of rooms	Onit	E I	Arithmetic Median mean	Median	Maximum	Location	Sampling year	Keterence
	o-xylene (CAS 95-47-6)	S 95-47-6)									
	Nursery/ elementary school	17	52	µg/m³	C LOD	9.	0.7	6.8	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.12222.
	Nursery	-	2	µg/m³	1	9.42	I	1	Czechia	₹ 2	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-
	School	50	22	µg/m₃	1.	ල. වෙ	2.3	52.4	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.
	Styrene (CAS 100-42-5)	3 100-42-5)									
	Nursery/ elementary school	17	51	µg/m³	0.9	2.	£.	4	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.1222.
!	Nursery school	~	2	µg/m₃	ı	3.325	ı	I	Czechia	Υ V	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757- 899X/245/8/082027.

Table A1 cont.

o N	Type of building	Number of buildings	Number Unit of rooms	Unit	Minimum	Arithmetic Median Maximum Location Sampling Reference mean	Median	Maximum	Location	Sampling year	Reference
œ	Styrene (C.	Styrene (CAS 100-42-5)									
	Elementary school	_	က	hg/m³	0.27	I	I	0.35	Italy	ı	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and outdoor
	Elementary school	-	က	րց/m³	0.32	I	I	0.59	Italy	I	monitoring of volatile organic compounds in school buildings: indicators based
	Elementary school	-	က	hg/m³	0.12	I	I	0.24	Italy	I	of fleatiff fisk assessment to single out critical issues. Int J Environ Res Public Health, 2013:10:6273–91, doi:10.3390/
	Elementary school	_	က	րց/m³	0.12	I	I	0.14	Italy	I	ijerph10126273.
	School	20	72	ng/m₃	-	4.	1.2	2.7	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.
တ	Toluene (C	Toluene (CAS 108-88-3)									
	Nursery/ elementary school	17	75	hg/m³	1.7	5.2	3.2	24.4	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.1222.
	Nursery school	~	2	µg/m³	ı	3.2	I	1	Czechia	۷ ۷	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-899X/245/8/082027.

Table A1 cont.

Type of building	Number Numbe of of buildings rooms	Number Unit of rooms	Unit	Minimum	Arithmetic Median mean	Median	Maximum Location Sampling Reference year	Location	Sampling year	Reference
Toluene (C	Toluene (CAS 108-88-3)									
Elementary school	-	က	րց/m³	2.55	I	I	4.84	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
Elementary school	-	က	µg/m³	4.12	I	I	6.81	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
Elementary school	←	က	hg/m³	0.73	I	I	0.97	Italy	I	indicators based on health fisk assessment to single out critical issues, Int J Environ Res Public
Elementary school	~	က	µg/m³	0.83	I	I	0.86	Italy	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.
School	20	72	m/gn	8.	15.1	4.9	202.5	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j.buildenv.2015.11.031.
1,4-dichlor	1,4-dichlorobenzene (CAS 106-46-7)	4S 106-46	(2-9							
Nursery/ elementary school	17	51	µg/m³	< LOD >	1.8	< LOQ	6.6	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.12222.

Table A1 cont.

Type of Numbuilding of	Nun	Number of	Number Unit of	Unit	Minimum	Arithmetic Median mean		Maximum Location Sampling year	Location	Sampling year	Reference
buildings rooms	puilding	10	rooms								
1,4-dichlorobenzene (CAS 106-46-7)	penzene	9	AS 106-4	3-7)							
Elementary 1 school	_		က	µg/m³	0.01	I	I	0.01	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
Elementary 1 school	-		က	hg/m³	0.04	I	I	0.04	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
Elementary 1 school	←		က	µg/m³	0.01	I	I	0.02	Italy	I	indicators based of Health risk assessment to single out critical issues. Int J Environ Res Public
Elementary 1 school	_		က	µg/m³	0.01	I	I	0.01	Italy	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.
					Ŏ	VOCs - polycyclic aromatic hydrocarbons	lic aromati	c hydrocarb	suo		
Naphthalene (CAS 91-20-3)	e (CAS 97	14	(6-3)								
School 114	1		300	µg/m³	~ LOD	2	COD >	31	Europe	2011/2012	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.
Nursery 2 school	7		1	ng/m³	13.1	I	14.8	62.7	Portugal	2015	Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Assessment of exposure to polycyclic aromatic hydrocarbons in preschool children: levels and impact of preschool indoor air on excretion of main urinary monohydroxyl metabolites. J Hazard Mater. 2017;322:357–69. doi:10.1016/j. jhazmat.2016.10.004.

Table A1 cont.

Number Duilding of Incoms Number Of Incoms Unit Actinimum Arithmetic Median Maximum Location Sampling Paternics Sampling Paternics 11 Namely Duildings Duildings Duildings CAS 91-20-31 1 1.16												
Naphthalene (CAS 91-20-3) Nursery 1 2 µg/m³ - 1.16 - - Czechia NA School 1 3 µg/m³ 2.51 - 4.01 Italy - Elementary 1 3 µg/m³ 2.24 - 4.79 Italy - Elementary 1 3 µg/m³ 4.21 - 6.1 Italy - Elementary 1 3 µg/m³ 1.3 - - 6.1 Italy - Elementary 1 3 µg/m³ 0.92 - - 2.62 Italy - School 1 3 µg/m³ 0.56 - - 1.21 Italy - Elementary 1 3 µg/m³ 0.56 - - 5.14 Italy - Elementary 1 3 µg/m³ 0.37 - - 5.29	N	Type of building	Number of buildings	Number of rooms	Unit	Minimum	Arithmetic mean	Median	Maximum	Location	Sampling year	Reference
Nursery 1 2 µg/m³ - 1.16 - - Czechia NA school Elementary 1 3 µg/m³ 2.54 - 4.79 Italy - Elementary 1 3 µg/m³ 4.21 - 4.79 Italy - Elementary 1 3 µg/m³ 4.21 - 6.1 Italy - Elementary 1 3 µg/m³ 0.92 - - 6.1 Italy - Elementary 1 3 µg/m³ 0.92 - - 6.1 Italy - Elementary 1 3 µg/m³ 0.56 - - 5.14 Italy - Elementary 1 3 µg/m³ 0.56 - - 5.14 Italy - Elementary 1 3 µg/m³ 0.37 - - 5.24 Ital	7	Naphthalen	ne (CAS 91-2	0-3)								
Limonene (CAS 138-86-3) Elementary school 1 3 µg/m³ 2.51 - - 4.01 Italy - School Elementary 1 3 µg/m³ 2.24 - - 4.79 Italy - School Elementary 1 3 µg/m³ 1.3 - - - 4.79 Italy - Elementary 1 3 µg/m³ 0.92 - - 6.1 Italy - Elementary 1 3 µg/m³ 0.56 - - 1.21 Italy - School 1 3 µg/m³ 0.56 - - 5.14 Italy - School 1 3 µg/m³ 0.36 - - 5.14 Italy - School 1 3 µg/m³ 0.37 - - 2.29 Italy - School 1		Nursery school	-	7	µg/m³	ı	1.16	1	Ī	Czechia	Ā	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-899X/245/8/082027.
Limonene (CAS 138-86-3) Elementary school 1 3 µg/m³ 2.51 - 4.01 Italy - Elementary school 1 3 µg/m³ 2.24 - - 4.79 Italy - Elementary school 1 3 µg/m³ 1.3 - - 6.1 Italy - Elementary school 1 3 µg/m³ 0.92 - - 2.62 Italy - Elementary school 1 3 µg/m³ 0.56 - - 1.21 Italy - Elementary school 1 3 µg/m³ 0.56 - - 4.05 Italy - School -							VOC	s – terpen	se			
Elementary 1 3 µg/m³ 2.51 - - 4.01 Italy - Elementary 1 3 µg/m³ 2.24 - - 4.79 Italy - Elementary 1 3 µg/m³ 4.21 - - 6.1 Italy - Elementary 1 3 µg/m³ 0.92 - - 6.1 Italy - Elementary 1 3 µg/m³ 0.56 - - 1.21 Italy - Elementary 1 3 µg/m³ 0.56 - - 5.14 Italy - School 1 3 µg/m³ 0.56 - - 5.14 Italy - Elementary 1 3 µg/m³ 0.37 - - 5.29 Italy - School - - - - - - - - <	12	Limonene (CAS 138-86-	-3)								
Elementary school 1 3 µg/m³ 2.24 - - 4.79 Italy - Elementary school 1 3 µg/m³ 4.21 - - 6.1 Italy - Elementary school 1 3 µg/m³ 0.92 - - 2.62 Italy - Elementary school 1 3 µg/m³ 0.92 - - 1.21 Italy - Elementary school 1 3 µg/m³ 0.56 - - 5.14 Italy - Elementary school 1 3 µg/m³ 0.36 - - 4.05 Italy - Elementary school 1 3 µg/m³ 0.37 - - 2.29 Italy -		Elementary school	_	က	hg/m³	2.51	I	ı	4.01	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
Elementary school 1 3 μg/m³ 4.21 - 6.1 Italy - Elementary school 1 3 μg/m³ 1.3 - - - 6.1 Italy - A-pinene (CAS 80-56-8) 3 μg/m³ 0.92 - - 2.62 Italy - Elementary school 1 3 μg/m³ 0.56 - - 5.14 Italy - Elementary school 1 3 μg/m³ 1.13 - - 4.05 Italy - Elementary school 1 3 μg/m³ 0.37 - - 2.29 Italy -		Elementary school	_	က	hg/m³	2.24	I	I	4.79	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
Elementary school 1 3 µg/m³ 1.3 - - 2.62 Italy - A-pinene (CAS 80-56-8) A-pinene (CAS 80-56-8) <th></th> <td>Elementary school</td> <td>-</td> <td>က</td> <td>hg/m³</td> <td>4.21</td> <td>I</td> <td>I</td> <td>6.1</td> <td>Italy</td> <td>I</td> <td>assessment to single out critical issues. Int J Environ Res Public</td>		Elementary school	-	က	hg/m³	4.21	I	I	6.1	Italy	I	assessment to single out critical issues. Int J Environ Res Public
Ca-pinene (CAS 80-56-8) A composition of the pine of the		Elementary school	1	က	hg/m³		I	I	2.62	Italy	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.
Itary 1 3 µg/m³ 0.92 - - 1.21 Italy - Itary 1 3 µg/m³ 0.56 - - 5.14 Italy - Itary 1 3 µg/m³ 1.13 - - 4.05 Italy - Itary 1 3 µg/m³ 0.37 - - 2.29 Italy -	13	α-pinene (C	AS 80-56-8)									
1 3 $\mu g/m^3$ 0.56 5.14 Italy - 1 3 $\mu g/m^3$ 0.37 - 2.29 Italy -		Elementary school	_	က	hg/m³	0.92	I	I	1.21	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
1 3 μg/m³ 1.13 – – 4.05 Italy – 1 3 μg/m³ 0.37 – – 2.29 Italy –		Elementary school	_	က	hg/m³	0.56	I	I	5.14	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
1 3 μg/m³ 0.37 – – 2.29 Italy –		Elementary school	1	က	hg/m³	1.13	I	I	4.05	Italy	I	assessment to single out critical issues. Int J Environ Res Public
		Elementary school	-	က	hg/m³	0.37	1	ı	2.29	Italy	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.

Table A1 cont.

Type of building		Number of buildings	Number Unit of rooms	Unit	Minimum	Arithmetic mean	Median	Maximum	Location	Arithmetic Median Maximum Location Sampling Reference mean	Reference
α-pinene (α-pinene (CAS 80-56-8)									
School		50	72	µg/m³	-	4.6	8.	35	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ . PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.
						VOCs - chlorinated hydrocarbons	rinated hyd	Irocarbons			
Tetrachlo	=	Tetrachloroethylene (CAS 127-18-4)	AS 127-18	<u>4</u>							
School		411	300	µg/m³	< LOD	-	< FOD >	18	Europe	2011/2012	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.
Elementary school	_	_	က	hg/m³	0.19	I	I	0.19	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
Elementary school	`	1	က	hg/m³	0.26	I	I	0.31	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
Elementary school	_	1	က	µg/m³	0.09	I	l	0.13	Italy	I	assessment to single out critical issues. Int J Environ Res Public
Elementary school	>	<u></u>	ო	hg/m³	0.14	I	I	0.15	Italy	ı	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.

Table A1 cont.

Trichloroethylene (CAS 79-01-6) Nursery/school 17 51 µg/m³ <lod< th=""> 2.3 <lod< th=""> 28.2 School 114 300 µg/m³ <lod< th=""> 3 <lod< th=""> 126 School 114 300 µg/m³ <lod< th=""> 3 <lod< th=""> 126 But/I acetate (CAS 123-86-4) 1 3 µg/m³ 0.48 - - 1.08 Elementary 1 3 µg/m³ 0.48 - - 0.79 School 1 3 µg/m³ 0.19 - - 0.29 School 1 3 µg/m³ 0.19 - - 0.29 School 1 3 µg/m³ 0.19 - - 1.01 Nursery 1 3 µg/m³ - 9.315 - -</lod<></lod<></lod<></lod<></lod<></lod<>		Type of building	Number of buildings	Number Unit of rooms		Minimum	Arithmetic Median mean	Median	Maximum Location	Location	Sampling Reference year	Reference
ttary 17 51 µg/m³ <lod (cas="" 0.19<="" 1="" 11="" 123-86-4)="" 2="" 2.3="" 3="" 300="" <lod="" esters="" itary="" little="" m³="" rectate="" th="" µg=""><th></th><th>Trichloroethy</th><th>lene (CAS</th><th>(9-10-6)</th><th></th><th></th><th></th><th></th><th></th><th></th><th></th><th></th></lod>		Trichloroethy	lene (CAS	(9-10-6)								
114 300 µg/m³ < LOD 3 < LOD 11	I	Nursery/ elementary school	71	15	µg/m³	C LOD	2.3	< LOD >	28.2	France	2010	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.12222.
Esters Esters scetate (CAS 123-86-4) Action (CAS 123-86-4)		School	4	300		C LOD	က	<pre></pre>	126	Europe	2011/2012	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.
tectate (CAS 123-86-4) ntary 1 3 µg/m³ 0.85 - - ntary 1 3 µg/m³ 0.19 - - ntary 1 3 µg/m³ 0.19 - - Y 1 2 µg/m³ - 9.315 -								Esters				
Itary 1 3 µg/m³ 0.85 - - Itary 1 3 µg/m³ 0.19 - - Itary 1 3 µg/m³ 0.19 - - Y 1 2 µg/m³ - 9.315 -		Butyl acetate (CAS 123-86	4								
Itary 1 3 µg/m³ 0.48 - - Itary 1 3 µg/m³ 0.19 - - Itary 1 3 µg/m³ 0.19 - - Itary 1 2 µg/m³ - 9.315 -		Elementary school	-	က	րց/m³	0.85	I	I	1.08	Italy	I	de Gennaro G, Farella G, Marzocca A, Mazzone A, Tutino M. Indoor and
Itary 1 3 µg/m³ 0.19 - - Itary 1 3 µg/m³ 0.19 - - - Image: Itary and the properties of the p	1	Elementary school	_	က	րց/m³	0.48	I	I	0.7	Italy	I	outdoor monitoring of volatile organic compounds in school buildings:
tfary 1 3 µg/m³ 0.19 – – – – – – – – – – – – – – – – – – –		Elementary school	-	က	hg/m³	0.19	I	I	0.29	Italy	I	indicators based on health lisk assessment to single out critical issues. Int J Environ Res Public
/ 1 2 µg/m³ – 9.315 –		Elementary school	-	က	hg/m³	0.19	I	I	1.01	Italy	I	Health. 2013;10:6273–91. doi:10.3390/ ijerph10126273.
		Nursery school	-	7	µg/m³	ı	9.315	ı	ı	Czechia	٧ ٧	Senitkova IJ. Occurrence of indoor VOCs in nursery school – case study. IOP Conference Series: Materials Science and Engineering. 2017;245:082027. doi:10.1088/1757-899X/245/8/082027.

Table A1 cont.

				l	I	
Reference				Mortamais M, Pujol J, van Drooge BL, Macià D, Martínez-Vilavella G, Reynes et al. Effect of exposure to polycyclic aromatic hydrocarbons on basal ganglia and attention-deficit hyperactivity disorder symptoms in primary school children. Environ Int. 2017;105:12–19. doi:10.1016/j.envint.2017.04.011.	Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Assessment of exposure to polycyclic aromatic hydrocarbons in preschool children: levels and impact of preschool indoor air on excretion of main urinary monohydroxyl metabolites. J Hazard Mater. 2017;322:357–69. doi:10.1016/j. jhazmat.2016.10.004.	Pacitto A, Stabile L, Viana M, Scungio M, Reche C, Querol X et al. Particlerelated exposure, dose and lung cancer risk of primary school children in two European countries. Sci Total Environ. 2018;616–17;720–9. doi:10.1016/j. scitotenv.2017.10.256.
Maximum Location Sampling year				2012/2013	2015	2012/2013
Location		VOCs)		Spain	Portugal	Spain
Maximum		Semi-volatile organic compounds (SVOCs)		425	0.109	1
Median		organic cor		96	0.0754	0.1
Arithmetic Median mean		ni-volatile c		105	1	1
Minimum		Ser		23	0.0501	ı
Unit				pg/m³	ng/m³	ng/m³
Number Unit of	rooms		50-32-8)	35	I	ı
Number of	buildings rooms		Benzo[a]pyrene (CAS 50-32-8)	35	8	36
Type of building			Benzo[a]p	School	Nursery	School
Š			17			

48

Table A1 cont.

Se_01-8) Se_01-8)	No.	Type of	Number	Number Unit	Unit	Minimum	Arithmetic Median		Maximum	Location	Sampling	Reference
Phenanthrene (CAS 85-01-6) Nuvsery/ 30 62 ng/m³ 1.6		building	of buildings				mean				year	
Nursery 30 62 ng/m³ 1.6	18	Phenanthr	rene (CAS 8	5-01-8)								
Nursery 2		Nursery/ elementary school		62	ng/m³	9.	ı	8.7	> 50	France	2017	Raffy G, Mercier F, Blanchard O, Derbez M, Dassonville C, Bonvallot N et al. Semivolatile organic compounds in the air and dust of 30 French schools: a pilot study. Indoor Air. 2017;27:114–27. doi:10.1111/ina.12288.
Diethyl phthalate (CAS 84-66-2) Phthalate (CAS 84-66-2) Action (CAS 84-66-2)		Nursery	2	ı	ng/m³	6.1	1	22	56.1	Portugal	2017	Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Assessment of exposure to polycyclic aromatic hydrocarbons in preschool children: levels and impact of preschool indoor air on excretion of main urinary monohydroxyl metabolites. J Hazard Mater. 2017;322:357–69. doi:10.1016/j. jhazmat.2016.10.004.
Diethyl phthalate (CAS 84-66-2) Nursery/selementary 30 53 ng/m³ 40 - 221 684 France 2010 elementary school school Day-care 63 63 μg/m³ 0.095 0.516 0.468 2.613 Germany 2011/2012								Phthalat	es			
30 53 ng/m³ 40 – 221 684 France 2010 63 63 μg/m³ 0.095 0.516 0.468 2.613 Germany 2011/2012	19	Diethyl ph	thalate (CAS	\$ 84-66-2)								
ıre 63 μg/m³ 0.095 0.516 0.468 2.613 Germany 2011/2012		Nursery/ elementary school		53	ng/m³	40	ı	221	684	France	2010	Raffy G, Mercier F, Blanchard O, Derbez M, Dassonville C, Bonvallot N et al. Semivolatile organic compounds in the air and dust of 30 French schools: a pilot study. Indoor Air. 2017;27:114–27. doi:10.1111/ina.12288.
		Day-care centre	63	63	µg/m³	0.095	0.516	0.468	2.613	Gегтапу	2011/2012	Fromme H, Lahrz T, Kraft M, Fembacher L, Dietrich S, Sievering S et al. Phthalates in German day-care centers: occurrence in air and dust and the excretion of their metabolites by children (LUPE 3). Environ Int. 2013;61:64–72. doi:10.1016/j.envint.2013.09.006.

Table A1 cont.

eference		Raffy G, Mercier F, Blanchard O, Derbez M, Dassonville C, Bonvallot N et al. Semi-volatile organic compounds in the air and dust of 30 French schools: a pilot study. Indoor Air. 2017;27:114–27. doi:10.1111/ina.12288.	Fromme H, Lahrz T, Kraft M, Fembacher L, Dietrich S, Sievering S et al. Phthalates in German day-care centers: occurrence in air and dust and the excretion of their metabolites by children (LUPE 3). Environ Int. 2013;61:64–72. doi:10.1016/j. envirt.2013.09.006.		Raffy G, Mercier F, Blanchard O, Derbez M, Dassonville C, Bonvallot N et al. Semi-volatile organic compounds in the air and dust of 30 French schools: a pilot study. Indoor Air. 2017;27:114–27. doi:10.1111/ina.12288.	Fromme H, Lahrz T, Kraff M, Fembacher L, Dietrich S, Sievering S et al. Phthalates in German day-care centers: occurrence in air and dust and the excretion of their metabolites by children (LUPE 3). Environ Int. 2013;61:64–72. doi:10.1016/j.
Maximum Location Sampling Reference year		2010 R. all all all all all all all all all al	2011/2012 Fe et		2010 R. all	2011/2012 Fr
Location		France	Germany		France	Germany
Maximum		800	2.613		800	1.276
Median		800	0.468		228	0.227
Arithmetic Median mean		1	0.516		I	0.283
Minimum		207	0.095		37	0.049
	2)	ng/m³	µg/m³	2)	ng/m³	µg/m³
Number Unit of rooms	CAS 84-69-	62	63	:AS 84-74-	62	63
Number Numbe of of buildings rooms	Diisobutyl phthalate (CAS 84-69-5)	30	63	Di-n-butyl phthalate (CAS 84-74-2)	30	63
Type of building	Diisobutyl	Nursery/ elementary school	Day-care centre	Di-n-butyl	Nursery/ elementary school	Day-care centre
o O N	20			21		

Table A1 cont.

;							:		:	:	
o Z	lype or building	Number of buildings	Number Unit of rooms			Arithmetic Median mean	Median	Maximum	Location	Location Sampling Reference year	Kererence
						Brominat	Brominated flame retardants	stardants			
22	Brominated	Brominated diphenyl ether 47 (CAS 5436-43-1)	her 47 (CA	S 5436-	43-1)						
	School	2	1	pg/m³	ı	1	131	459	Norway	2017	Malliari E, Kalantzi O-I. Children's exposure to brominated flame retardants in indoor environments – a review. Environ Int. 2017;108:146–69. doi:10.1016/j.envint.2017.08.011.
	School	8	м	hg/m³	I	I	0.000131	I	Norway	2016	Cequier E, Ionas AC, Covaci A, Marcé RM, Becher G, Thomsen C. Occurrence of a broad range of legacy and emerging flame retardants in indoor environments in Norway. Environ Sci Technol. 2014;48(12):6827–35. doi:10.1021/es500516u.
							Musks				
23	Galaxolide (Galaxolide (CAS 1222-05-5)	15-5)								
	Nursery/ elementary school	30	62	ng/m³	4	I	> 50	> 50	France	2010	Raffy G, Mercier F, Blanchard O, Derbez M, Dassonville C, Bonvallot N et al. Semi-volatile organic compounds in the air and dust of 30 French schools: a pilot study. Indoor Air. 2017;27:114–27. doi:10.1111/ina.12288.
24	Tonalide (CA	Tonalide (CAS 21145-77-7)	(2								
	Nursery/ elementary school	30	06	ng/m³		ſ	20	> 50	France	2010	Raffy G, Mercier F, Blanchard O, Derbez M, Dassonville C, Bonvallot N et al. Semi-volatile organic compounds in the air and dust of 30 French schools: a pilot study. Indoor Air. 2017;27:114–27. doi:10.1111/ina.12288.

Table A1 cont.

Reference			Annesi-Maesano I, Baiz N,	Banerjee S, Kudnai P, Kive S & on behalf of the SINPHONIE	Group. Indoor air quality	and sources in schools and related health effects. I Toxicol	Environ Health B Crit Rev.	2013;16:8:491–550. doi:10.10 80/10937404.2013.853609.	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.		Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Indoor air quality in preschools (3- to 5-year-old children) in the northeast of Portugal during spring-summer season: pollutants and comfort parameters. J Toxicol Environ Health A. 2017;80(13- 15):740-55. doi:10.1080/1528
Sampling R year			NA A	NA AN	NA	NA RE	NA E	NA 8	2011/2012 E H H H H H H H H H H H H H H H H H H		2013 O O O E E E E E E E E E E E E E E E E E
Maximum Location			NA	NA	NA	NA	NA	NA	Europe		Portugal
Maximum			230	170	250	47	37	33.2	88		ı
Median	Inorganic compounds		I	I	ı	I	I	I			ı
Arithmetic Median mean	Inorganic		44.26	32.96	59.52	29	I	I	4-		0.67
Minimum			99.8	9.9	13.9	13	12.5	15.2	<		I
Unit			hg/m³	hg/m³	hg/m³	µg/m₃	µg/m³	hg/m³	hg/m³		mg/m³
Number Unit of rooms		0102-44-0)	NA	A A	ΑN	ΝΑ	NA	NA	300	(0-80-089	∢ Z
Number of buildings		Nitrogen dioxide (CAS 10102-44-0)	NA	NA	NA	NA	NA	NA	41.	Carbon monoxide (CAS 630-08-0)	₹ Z
Type of building		Nitrogen dic	School	School	School	School	School	School	School	Carbon mon	School
No.		25		-	-	. '	. '			56	

Table A1 cont.

Type of Nu building of	Number	Number	Unit	Minimum	Arithmetic Median mean	Median	Maximum Location	Location	Sampling	Sampling Reference vear
D	sguipling								no.	
Carbon m	Carbon monoxide (CAS 630-08-0)	AS 630-08-(((
School	411	300	Parts per million (ppm)	< FOD	-	C LOD	122	Europe	2011/2012	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014, doi:10.2788/99220.
School	20	72	mg/m³	0.01	0.48	0.38	1.7	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j.buildenv.2015.11.031.
Ozone (C	Ozone (CAS 10028-15-6)	2-6)								
Nursery	2	1 playroom 1 classroom	mg/m³	0.02	0.08	ſ	ſ	Portugal	2013	Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Indoor air quality in preschools (3-to 5-year-old children) in the northeast of Portugal during spring-summer season: pollutants and comfort parameters. J Toxicol Environ Health A. 2017;80(13–15):740–55. doi:10.1080/15287394.2017.1286932.

Table A1 cont.

Reference		European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.			Annesi-Maesano I, Baiz N, Banerjee	S, Rudnai P, Rive S & on behalf of the	SINPHONE Group. Indoor air quality and sources in schools and related	health effects. J Toxicol Environ Health B	Crit Rev. 2013;16:8:491–550. doi:10.108	0/1093/404.2013.853609.			Vilcekova S, Meciarova L, Kridlova Burdova E, Katunska J, Kosicanova D, Doroudiani S. Indoor environmental quality of classrooms and occupants' comfort in a special education school in Slovak Republic. Build Environ. 2017;120:29–40. doi:10.1016/j. buildenv.2017.05.001.
		Europ Gene Direct Centr Health (SINF Public			Anne	S, Ru	ANIS	healt	Crit R	0/109			,
Sampling year		2011/2012			2010	2010	2010	2010	2010	2010	2010	NA	2013/2014
Location		Europe			Italy	Italy	Norway	Sweden	Denmark	France	Italy	NA	Slovakia
Maximum		142	ter (PM)		247	260	131	53	233	151	247	300	2061.2
Median		က	Particulate matter (PM)		141	154	43	32	160	106	141	70	1
Arithmetic Median Maximum mean		ω	Partic		148	158	54	33	169	112	148	79.3	287.84
Minimum		< LOD		1,0)	69	92	17	14	112	98	69	6	32.5
Unit		µg/m³		0 µm (PN	hg/m³	µg/m₃	hg/m³	hg/m³	hg/m³	hg/m³	hg/m³	µg/m₃	µg/m³
Number Unit of rooms	2-6)	300		ess than 1	NA	NA	NA	NA	NA	ΝΑ	NA	NA	2
Number Numbe of of buildings rooms	Ozone (CAS 10028-15-6)	411		PM with diameter of less than 10 µm (PM ₁₀)	ΑN	ΑN	ΑN	ΑN	ΑN	ΑΝ	ΑN	ΑN	-
Type of building	Ozone (C/	School		PM with d	School	School	School	School	School	School	School	School	School
O	27			28									

Table A1 cont.

Type of Number	Numb	er	Number Unit	Unit	Minimum	Arithmetic Median	Median	Maximum	Location	Maximum Location Sampling	Reference
sduiplind	puilding	S	rooms								
PM with diameter of less than 10 µm (PM ₁₀)	liameter	of k	ess than 10	md) um (10)						
School 10	10		4	_s m/gh	32	82.24	70	197	Serbia	۲	Matic B, Rakic U, Jovanovic V, Dejanovic S, Djonovic N. Key factors determining indoor air PM10 concentrations in naturally ventilated primary schools in Belgrade, Serbia. Zdr Varst. 2017;56(4):227–35. doi:10.1515/sjph-2017-0031.
School 39	36		I	hg/m³	ı	I	60.7	I	Spain	2012/2013	Pacitto A, Stabile L, Viana M, Scungio M, Reche C, Querol X et al. Particlerelated exposure, dose and lung cancer risk of primary school children in two European countries. Sci Total Environ. 2018;616–17;720–9. doi:10.1016/j. scitotenv.2017.10.256.
Day-care 30 centre	36		I	hg/m³	I	75.91	I	I	Ϋ́	ΑN	Morawska L, Ayoko GA, Bae GN, Buonanno G, Chao CYH, Clifford
School 7	7	78	I	hg/m³	I	116.92	I	I	ΑN	NA	 S et al. Airborne particles in Indoor environment of homes, schools, offices and aged care facilities: the main routes
School 281	28	<u>5-</u>	I	hg/m³	ı	182.03	I	I	A A	Y V	of exposure. Environ Int. 2017;108:75–83. doi:10.1016/j.envint.2017.07.025.
Nursery school		2	24	µg/m³	I	122.11	I	ı	Poland	2013/2014	Mainka A, Zajusz-Zubek E. Indoor air quality in urban and rural preschools
Nursery school		7	24	hg/m³	I	103.89	I	I	Poland	2013/2014	in Upper Silesia, Poland: particulate matter and carbon dioxide. Int J Environ Res Public Health. 2015;12:7697–711. doi:10.3390/ijerph120707697.

Table A1 cont.

Maximum Location Sampling Reference year		2011/2013 Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ . PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.		Annesi-Maesano I, Baiz N, Banerjee	S, Rudnal P, Rive S & on behalf of the SINPHONIE Group. Indoor air quality and sources in schools and related health effects. J Toxicol Environ Health B Crit Rev. 2013;16:8:491–550. doi:10.108 0/10937404.2013.853609.	Oliveira M, Slezakova K, Delerue-Matos C, do Carmo Pereira M, Morais S. Indoor air quality in preschools (3- to 5-year-old children) in the northeast of Portugal during spring-summer season:
Sam year		2011,		ΑN	∢ Z	2013
Location		Portugal		NA	₹ Z	Portugal
Maximum		320		13.7	28	28.1
		127		I	I	I
Arithmetic Median mean		139		I	I	17.5
Minimum	· ·	99	,,,	1.6	ω	7.58
	Jum (PM	µg/m³	5 µm (PM	hg/m³	µg/m³	µg/m³
Number Unit of rooms	less than 10	72	less than 2.	ΑN	Ą Z	1 playroom 1 classroom
Number of buildings	PM with diameter of less than 10 µm (PM10)	20	PM with diameter of less than 2.5 μm (PM $_{2.5}$)	Ϋ́	Ϋ́	7
Type of Nur building of buil	PM with di	School	PM with di	School	School	Nursery school
o O	28		53			

Table A1 cont.

Reference		Vilcekova S, Meciarova L, Kridlova Burdova E, Katunska J, Kosicanova D, Doroudiani S. Indoor environmental quality of classrooms and occupants' comfort in a special education school in Slovak Republic. Build Environ. 2017;120:29–40. doi:10.1016/j. buildenv.2017.05.001.	Canha N, Mandin C, Ramalho O, Wyart G, Riberon J, Dassonville C et al. Assessment of ventilation and indoor air pollutants in nursery and elementary schools in France. Indoor Air. 2016;26:350–65. doi:10.1111/ina.12222.	Morawska L, Ayoko GA, Bae GN, Buonanno G, Chao CYH, Clifford	S et al. Airborne particles in indoor	environment or normes, schools, offices and aged care facilities: the main routes of exposure. Environ	Int. 2017;108:75–83. doi:10.1016/j. envint.2017.07.025.	European Commission Directorate General for Health and Consumers, Directorate General Joint Research Centre. Schools Indoor Pollution and Health Observatory Network in Europe (SINPHONIE). Final report. Luxembourg: Publications Office of the European Union; 2014. doi:10.2788/99220.
Sampling Reference year		2013/2014	2010	NA A	NA	NA	ΑN	2011/2012
Location		Slovakia	France	ΑN	NA	NA	ΝΑ	Europe
Arithmetic Median Maximum mean		50.1	I	I	I	I	I	250
Median		I	I	I	I	I	I	37
Arithmetic mean		17.12	52	44.27	40.47	52.31	50.14	4
Minimum		<u>υ</u>	ı	I	I	I	I	4
Unit	ım (PM _{2.5})	µg/m³	µg/m³	րց/m³	hg/m³	րց/m³	µg/m³	µg/m³
Number of Unit rooms	PM with diameter of less than 2.5 μ m (PM $_{2.5}$)	ro.	75	I	I	I	I	300
Number of buildings	ameter of le	-	17	304	18	63	159	1. 4
Type of building	PM with dia	School	Nursery/ elementary school	Day-care centre	School	Day-care centre	School	School
No.	53		•	•	,		'	

Table A1 cont.

o N	Type of building	Type of Number Numbe building of buildings rooms	Number Unit of rooms	Unit	Minimum	m Arithmetic Median Maximum Location Sampling Reference mean	Median	Maximum	Location	Sampling year	Reference
59	PM with di	PM with diameter of less than 2.5 $\mu m \; (\text{PM}_{2.5})$	ess than 2.	5 µm (P	M _{2.5})						
	Nursery school	2	24	hg/m³	I	94.08	l	I	Poland	2013/2014	Mainka A, Zajusz-Zubek E. Indoor air quality in urban and rural preschools
	Nursery school	2	24	hg/m³	l	66.72	I	I	Poland	2013/2014	in Upper Silesia, Poland: particulate matter and carbon dioxide. Int J Environ Res Public Health. 2015;12:7697–711. doi:10.3390/ijerph120707697.
	School	20	72	hg/m³	38	46	85	244	Portugal	2011/2013	Madureira J, Paciencia I, Rufo J, Severo M, Ramos E, Barros H et al. Source apportionment of CO ₂ , PM ₁₀ and VOC levels and health risk assessment in naturally ventilated primary schools in Porto, Portugal. Build Environ. 2016;96:198–205. doi:10.1016/j. buildenv.2015.11.031.

58

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