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SEASONAL VARIATION OF TRAFFIC-RELATED AIR POLLUTANTS IN BANGKOK METROPOLITAN DAY CARE CENTERS: EFFECTS ON RESPIRATORY SYMPTOMS OF CHILDREN

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CHILDREN**



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List of Abbreviations and Acronyms

ADHD	Attention Deficit Hyperactive Disorder
CCAAPS	Cincinnati Childhood Allergy and Air Pollution Study
CFU	Colony forming unit
CO	Carbon monoxide
COHb	Carboxy hemoglobin
DCCs	Day Care Centers
DSM-5	American Psychiatric Association's Diagnostic and Statistical Manual, Fifth edition
ECAT	elemental carbon attributed to traffic
F _E NO	Fractional Exhaled Nitric Oxide
NO ₂	Nitrogen dioxide
NOS	Nitric oxide synthase
O ₃	Ozone
PM10	particles with an aerodynamic diameter smaller than 10 µm
ppm	Part per million
SO ₂	Sulfur dioxide

CHAPTER I INTRODUCTION

1.1 Rationale

Clean air is a fundamental requirement for human health and well-being [1]. During the past few decades, human activities, industrial activities, and transportation have contributed to increasing contamination of the ambient air environment. Human are exposing to a variety of air pollutants, and air pollution is becoming a major problem throughout the world.

Nowadays, people are spending more time indoors. Indoor air pollution and its effect on health has become an issue of concern worldwide. Indoor air can be contaminated by chemical, physical, and biological agents. There are two important factors that influence the level of indoor air pollutants: the pollutant source, and the amount of ventilation. Sources of indoor air pollution range from combustion of solid fuels, such as biomass and coal, in the developing world, which emit CO and respirable particles, to volatile organic compounds from construction material, furnishing, and consumer products used in modern homes. Outdoor air pollution is also a source of indoor air pollution. Outdoor air can enter and leave a building by infiltration, natural ventilation, and mechanical ventilation. In the presence of indoor pollution source, poor ventilation will aggravate the accumulation of the pollutants [2].

For many reasons, children are more susceptible to health effects from indoor air pollution than adults. Infants and preschool children spend most of their time indoor, and therefore are exposing more to indoor air pollutants. The immature lungs and immune system of children also make them more vulnerable to the environmental insults. A term infant is born with about 25 million alveoli in the lungs, most of new alveoli are formed in the first two years of life, then, alveolar growth continues until age of 8 years old when the number of alveoli reaches 300 million. After that, linear growth of lung volume occurs without forming new alveoli. Exposure to pollutants at this developmental stage can cause adverse effects on lung structure, as well as function. Children's respiratory rate is higher than that of adult, resulting in higher minute ventilation. Children also have larger lung surface area per body weight compared to adult. Therefore, children are exposed to proportionately larger dose of pollutants than adults. Furthermore, many aerosolized pollutants are heavier than air. Children's breathing zone is closer to the ground than that of adults, and they are exposed to higher concentration of a lot of pollutants when staying in the same indoor space [3].

Bangkok is the most highly populated city in Thailand, with over 7 million people and over 2.5 million households. It is also a highly polluted city. There are approximately 300 day care centers (DCCs) in Bangkok. Increasing numbers of preschool children are going to DCCs, and these children spend time at DCCs about 40-45 hours per week. Preschool

children's exposure to air pollution comes primarily from indoor. For children attending DCCs, indoor air in DCCs is a significant source of pollutants exposure.

There have been environmental studies done for assessing level of some indoor air pollutants in DCCs from countries in different parts of the world. A study in Paris found that NO_2 and airborne endotoxin level was significantly higher in DCCs compared to in homes, and 1 out of 28 DCCs had total dust mite allergen in cold season greater than the proposed sensitization threshold level [4]. In Nigeria, a study of particulate matter burden in 48 DCCs found that mean indoor PM_{10} level significantly exceeds WHO (World Health Organization) guideline limit [5]. Dampness and biological indoor air pollution is also a subject of interest. There was a study in Norway in 2005. Inspection of the 175 DCCs was done, and dampness problems, such as sign of molds and water leakage, were observed in 51% of DCCs, but no significant effect on respiratory symptoms of children were found in the study [6]. In Taiwan, airborne fungi and bacteria count in two DCCs was studied, and levels of indoor microbial were above the IAQ (Indoor Air Quality) recommendations [7].

From the Cincinnati Childhood Allergy and Air Pollution Study (CCAAPS), birth cohort study on exposure to Elemental Carbon Attributed to Traffic (ECAT) during infancy and behavioral scores at 7 years of age found that exposure to the highest tertile of ECAT during the child's first year of life was significantly associated with Hyperactivity T-scores in the "at risk" range at 7 years of age. Other previous studies show negative correlation between air pollutants and neurocognitive development in children [8].

In Thailand, only a few studies were carried out on this subject. There was a study in 2005 on 24-hour indoor PM_{10} level done in 25 DCCs in high-polluted area, and 25 DCCs in low-polluted area in Bangkok. No significant difference in PM_{10} concentrations were found between the two areas. Both areas had level of PM_{10} concentration above the WHO recommendation. The study also found positive correlation between PM_{10} concentrations and number of children complaining about cough in the high-polluted area [9, 10]. Another study was done in a child home care center in Bangkok in 2006, where microbial count and PM_{10} levels were measured in 20 households. Almost half of the total air samples had bacterial, fungal counts, and PM_{10} levels are higher than the recommended level [11].

1.2 Objectives

Today, there is still very limited information about the indoor air pollution level in DCCs in Thailand. Although effects of indoor air pollution on health have been widely studied in adults and older children, there are not many studies in the preschool age group, especially in tropical climate regions. Hence, this project aims to combine the results from last year project as follows.

1. To assess the level of indoor air pollutants(including traffic related air pollutants) in DCCs in Bangkok
2. To determine environmental characteristics of DCCS that influence the indoor air pollutants level

3. To determine the association between exposure to indoor air pollution in DCCs in Bangkok and respiratory symptoms, Fractional Exhaled Nitric Oxide (FENO) of Children under 6 years old
4. To determine the association between exposure to indoor air pollution in DCCs in Bangkok and Attention Deficit Hyperactivity Disorder (ADHD) symptoms of children under 6 years old

1.3 Methodology

Details of methodology employed in the present investigation are as follows.

1.3.1 Study Design

Prospective, descriptive study were followed.

1.3.2 Population

Population were included and excluded according to the following criteria.

Inclusion Criteria

- For day care centers (DCCs): DCCs located in Bangkok with signed informed consent to participate in the study
- For Children :Children under 6 years old attending the DCCs for at least a year before the period of study whose parent give signed informed consent to participate in the study

Exclusion Criteria

- Day care centers and children whose parents do not consented for participating
- Children leaving the day care centers during the time of study

1.3.3 Methods of Data Collections

This is a descriptive study. Data will be collected on *Day care environmental characteristics* by direct inspection and question on

- total indoor area,
- type of building material,
- year of construction,
- ventilation system,
- number of children,
- presence of pet indoor,
- indoor cooking,
- proximity to traffic or water resources,
- visible signs of dampness or mold.

Children demographic data and home environment will be collected by questionnaire including

- age, gender, ethnicity of children, as well as history of drug and food allergy,

- immunization for influenza,
- maternal and paternal atopy,
- exposure to smoking at home,
- current pet exposure,
- socio-economic status and
- number of siblings.

Data on housing characteristics will also be collected on

- indoor combustion of fuel for cooking,
- indoor smoking,
- ventilation device used at home,
- molds, dampness or water leakage in home.

Pollutant level will be measured. Samples of indoor air will be collected from DCCs and measure for

- PM₁₀
- CO
- NO₂
- SO₂
- O₃
- Benzene
- bacteria count
- fungi count and
- dust mite level

Respiratory symptoms of children will be assessed by using questionnaire on symptoms including

- Rhinitis symptoms: runny nose, sneezing, blocked nose, itchy nose,
- Cough
- Lower respiratory symptoms: dyspnea
- Absence from school due to respiratory symptoms
- Doctor visit and admission due to respiratory symptoms

For each four-week period the questionnaires will be sent to parents weekly for four times. At DCCs, teachers will also observe and record respiratory symptoms for each child every Monday for four weeks. *Exhaled Nitric Oxide level* representing airway inflammation will be measured in children from each DCCs. Pollutant levels, respiratory symptoms and exhaled nitric oxide levels will be measured three times, each during summer, rainy and winter season due to possible seasonal variation of air pollution. Attention Deficits

Hyperactive Disorder (ADHD) symptoms will be assessed by a standardized questionnaire “Thai ADHD screening scale” filled out by parents and teachers.

1.3.4 Statistical Analyses

Poisson regression and Mann-Whitney U test were performed to find out association among the pollutants level and clinical outcomes. The multiple logistic regression analysis was used to analyze association between benzene, fungi and FENO levels.

CHAPTER 2 LITERATURE REVIEW

2.1 Air Pollutants and Indoor Air Pollution

Air pollution is contamination of indoor or outdoor air environment by chemical, physical or biological agents. The quality of air inside homes, offices, schools, day care centers, public buildings, health care facilities or other private and public buildings, where people spend a large part of their life, is an essential determinant of healthy life and people's well-being.

Indoor air pollutions come from indoor and outdoor sources. Outdoor air can enter and leave building by infiltration, natural ventilation, and mechanical ventilation. For infiltration, air come in and out through any openings, joints, cracks in walls, floors and ceilings, and also around windows and doors. Natural ventilation is when air moves through windows and doors that are opened. These two methods of ventilation occur due to differences in temperature between indoor and outdoor space, and also by wind. There are many mechanical ventilation devices, such as fans, air conditioner, air handling systems. Accumulation of indoor air pollutants occurs when the air exchange rate inside a close space is low.

Exposure to air pollutants can cause both short term and long term health effects. Sources and health effects of the pollutants in this study are shown in Table 1.

Table 1: Sources and health effects of common air pollutants

Pollutant	Source	Health effects
CO	Incomplete combustion of carbonaceous fuels	- Asphyxiating effect - Ischemic heart disease - Respiratory symptoms
NO ₂	Fuel combustion, Tobacco, Traffic smoke	- Increased bronchial reactivity - Airway inflammation
Ozone	Mostly from outdoor	- Airway irritation - Aggravate asthma and COPD
SO ₂	Fossil fuel combustion	- Trigger bronchoconstriction
Benzene	Furnishing material, tobacco, fuel combustion	- Increased lifetime risk for leukemia
PM ₁₀	Combustion, traffic	- Worsening respiratory symptoms - Decreased lung function
Mold	Damp environment, bathroom, kitchen	- Respiratory infection - Allergic reaction

Bacteria	Human activity	- Increased risk for infection - Endotoxin can trigger wheezing
Dust mite	Pillow , mattress, cushion, carpet	- Allergic reaction - Increased risk for asthma

2.1.1 Specific pollutants and their health effects

Carbon monoxide

Carbon monoxide is a product of incomplete combustion of carbonaceous fuels; therefore, a major source is from combustion such as cooking, heating, and traffic. It is a colorless, non-irritant, odorless and tasteless gas. The only exogenous exposure route is by inhalation.

Carbon monoxide strongly binds to hemoglobin, forming carboxyhemoglobin (COHb) and causes hypoxemia and leftward shift in the oxyhemoglobin dissociation curve which means less oxygen release to tissue. Carbon monoxide has asphyxiating effect. Other health effects from carbon monoxide exposure include increase of ischemic heart disease symptoms and decrease of exercise tolerance, hospital admission or emergency department visit due to respiratory symptoms.

Nitrogen dioxide

Nitrogen dioxide is a free radical, a strong oxidant and corrosive, it is heavier than air. The principal source is road traffic; whereas, indoor sources also include tobacco smoke, burning of solid fuel, candles, mosquito coils.

It causes airway injury and inflammation, bronchoconstriction, decreases in immune defense, which can result in respiratory infection. There is suggestive evidence that exposure to indoor nitrogen dioxide is associated with increased respiratory symptoms both in children and adults.

Ozone

Ground level ozone is also a strong oxidant. The source is primarily from outdoor when pollutants such as oxides of nitrogen and volatile organic compounds are exposed to sunlight. It is irritative to airway causing respiratory symptoms, wheezing and shortness of breath, it can also aggravate asthma.

Sulfur Dioxide

Sulfur Dioxide can form harmful compounds such as sulfuric acid, sulfurous acid and sulfate particles. Its major sources come from fossil fuel combustion at power plants and other industrial processes such as extracting metal from ore, and the burning fuel containing sulfur.

SO₂ has many adverse effects on the respiratory system such as bronchoconstriction, increased asthma symptoms, increased emergency room visits, increased hospital admissions due to respiratory diseases, especially in children, elderly, and asthmatics.

Benzene

Benzene is a pollutant that usually has higher level indoors compared to outdoors. The sources are from materials used in building or furnishing such as polymer, rubber, adhesives, particleboard, paints. Fuel combustion and smoking can also be a source of Benzene. High level can result in death. Benzene exposure also impairs respiratory health.

According to the French National Research and Safety Institute, acute benzene exposure will cause no effect at 25 ppm (81 mg/m³), headaches and asthenia from 50 to 100 ppm (162–325 mg/m³), more accentuated symptoms at 500 ppm (1625 mg/m³), tolerance for only 30–60 minutes at 3,000 ppm (9720 mg/m³), and death in 5–15 minutes at 20,000 ppm (64 980 mg/m³) due to asphyxia or central nervous system depression. Exposure to benzene also associated with haematologic diseases such as aplastic anaemia, pancytopenia, thrombocytopenia, granulopenia, lymphopenia and increased lifetime risk for leukaemia.

PM₁₀

PM₁₀ or particles with an aerodynamic diameter smaller than 10 µm represents the particle mass that can enter the respiratory tract. Particulate matter includes coarse and fine particles. Course particles (diameter between 2.5 and 10 µm) can pass through upper airway, while fine particles (diameter less than 2.5 µm, known as PM_{2.5}) are small enough to pass through the lower airway.

Sources of PM₁₀ are dust from traffic, construction, agriculture, and other activities. Particulate matters are associated with worsening respiratory symptoms, decreased lung function recurrent health care utilization, and also shown to have a small but significant adverse effect on cardiovascular and cerebrovascular disease.

Biological Pollutants

Biological pollutants include microorganisms and by-products such as bacteria, fungus, viruses, house dust mite, animal danders. These indoor pollutants level is usually linked to dampness and inadequate air exchange rate in the indoor space. Excess moisture promotes growth of these microbes.

Molds

Molds can cause infection and allergic reaction. They can also produce toxins. Exposure to indoor molds can lead to respiratory symptoms, respiratory infections, and exacerbation of asthma.

House dust mites

House dust mites are usually found in the pillows, mattresses, carpets, curtains, or cushions. Dermatophagoides (Der P1) is most prominent dust mite allergen.

There are evidence that exposure to Dermatophagoides in infancy is associated with increased prevalence of positive skin prick test and increased concentration of IgE specific to dust mite by age 5 in children of atopic parents. [12] Dermatophagoides level higher than 10 mcg/g are associated with 5 fold increased in the risk of asthma by age 11 years old [13]

Bacteria

Many human activities, such as sneezing, coughing, toilet flushing can generate airborne bacteria. Airborne bacteria can cause infections including respiratory infection, skin infection. Gram negative bacilli can produce endotoxin, there is evidence that exposure to high endotoxin levels is associated with wheezing in childhood [14]

Summary of standards for indoor air quality is shown in Table 2.

Table 2: Standards for indoor air quality

Pollutants	Short term (24 hr)	Long term (annual)	Reference
PM ₁₀	50 µg/m ³	20 µg/m ³	WHO
CO	100 mg/m ³ (15 min) 35 mg/m ³ (1 hr) 10 mg/m ³ (8 hr) 7 mg/m ³ (24 hr)	-	WHO
NO ₂	200 µg/m ³	40 µg/m ³	WHO
Ozone	100 µg/m ³ (8 hr)	-	WHO
SO ₂	20 µg/m ³	-	WHO
Microbial count (fungi/bacteria)	500 CFU/m ³		American Industrial Hygiene Association
Dust Mite Antigen	2 µg/g Threshold for sensitization		Peat JK et al.

2.1.2 Indoor air pollution in day care centers

DCCs are important sources of indoor air pollutants exposure for preschool children. There are more than 300 DCCs in Bangkok. Children usually spend five days a week at the DCCs during daytime, which account for approximately 40 – 45 hour per week.

There are a number of studies on the level of indoor air pollutants concentration in DCCs from different part of the world. The studies and results are summarized in Table 3.

Table 3: Previous studies on indoor air pollution in day care centers

Study	Place/Time	Data collected	Pollution data
Microbial counts and particulate matter levels in indoor air samples collected from a child home-care center in Bangkok, Thailand. J Med Assoc Thai 2012; 95 (Suppl. 6): S161-S168	20 households which were part of a child home-care center in Bangkok 2006-2007	- Microbial count - PM10 levels	- 47.2% and 47.6% of total air samples had bacterial and fungal counts higher than recommended levels - 47.0% of total air samples had PM10 levels higher than recommended level
Assessment of indoor environment in Paris child day care centers C. Roda et al. Environmental Research 111 (2011) 1010–1017	28 child day care centers	- Dust mite allergens - Endotoxins - Airborne fungi - Aldehydes - VOCs - NO2 - CO2 - temperature - Relative humidity	- Only one child day care center in cold season had total group1 House Dust Mite allergen levels greater than proposed sensitization threshold level of 2 mg/g - Airborne endotoxin levels in child day care centers were higher than those found in Paris dwellings - the levels for NO2 being significantly higher than those measured in homes
Indoor air quality differences between urban and rural preschools in Korea Environ Sci Pollut Res (2011) 18:333–345	71 classrooms at 17 Korean preschools	- total suspended particulates - respirable particulates - lead - asbestos - volatile organic compounds - formaldehyde - CO2	- Total dust concentration was highest in urban indoor settings followed by urban outdoor, rural indoor, and rural outdoor locations - Lead concentrations were much higher in urban than in rural areas
Inhalable particulate matter burden in selected day-care centres in Ibadan, Nigeria Godson R. E. E. Ana <i>International Journal of Environmental Monitoring and Analysis</i> 2013; 1(6): 296-301	48 day-care centres (DCCs) in Ibadan, South western Nigeria	- PM ₁₀	- Mean indoor PM ₁₀ for wet season 73.4 ± 54.4µg/m ³ and dry season 296.3 ± 61.6µg/m ³ significantly exceeded guideline limit (P<0.05).
Airborne fungi and bacteria in child daycare centers and the effectiveness of weak acid hypochlorous water on controlling microbes	two child daycare centers in Tainan City in Taiwan	- indoor and outdoor airborne fungi and bacteria	- indoor microbial levels were above the IAQ recommendations/ standards of Taiwan, Korea, and Hong Kong (ranging from 500 to 1000 CFU/m ³ for both fungi and bacteria)

	2008		
Nai-Tzu Chen, <i>J. Environ. Monit.</i> , 2012, 14, 2692			
Indoor Concentrations of PM10 and Factors Influencing Its Concentrations in Day Care Centres in Bangkok, Thailand Nareerut Pudpong <i>Asia J Public Health</i> 2011;2(1):3-12	25 DCCs in high-polluted area (H), 25 DCCs in low-polluted area (L) in Bangkok	24-hour indoor PM ₁₀ level	- no significant difference in PM10 concentrations between the H (75.08µg/m ³) and L (69.93µg/m ³) area

2.2 Vulnerability of Children to Air Pollution

Children are more vulnerable to indoor air pollutants than adults, especially infants and preschool children because they spend most of their time indoor, and therefore are exposing more to indoor air pollutants. Children are exposed to proportionately larger dose of pollutants than adults because they have higher respiratory rate and minute ventilation by their physiology, and children also have larger lung surface area per body weight compared to adult. Furthermore, a child's breathing zone is closer to the ground than that of adults, and many aerosolized pollutants are heavier than air. These pollutants tend to have higher concentration near the ground, so children will expose to a higher concentration of many pollutants when staying in the same indoor space with an adult.

Inflammation of the airway caused by exposure to air pollutants can cause airway edema. Airway diameter is smaller in children compared to adults, edema to the airway will have greater effect. Note that resistance to air flow is inversely proportional to the radius of the lumen to the fourth power for laminar flow, and to the radius of the fifth power for turbulent flow. A 1 mm circumferential edema in children results in 75% reduction in cross-sectional area and a 16-fold increase in airway resistance, compared to in adults, the same 1 mm of circumferential airway edema causes 44% reduction in cross-sectional area and only three-fold increase in airway resistance. The difference between effects of airway edema in children and adults is shown in Figure 1.

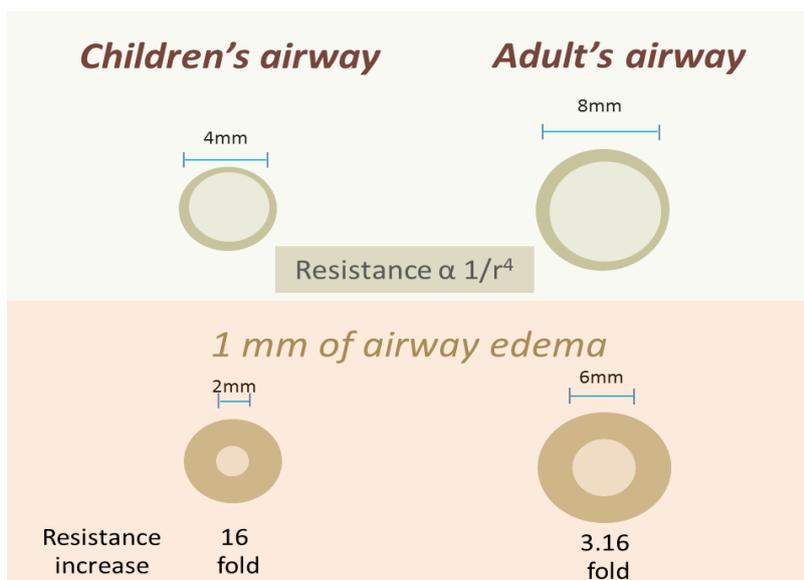


Figure 1: Effect of airway edema in children compared to adults

An adult has approximately 300 million alveoli in the lungs. A term newborn infant is born with 25 million alveoli. Alveolar development occurs rapidly during the first two years of life, after that, it slows down and stops at about 6-8 years old when the alveoli number reaches the same number with that in adult. After 6-8 years of age, linear increase in lung volume takes place until adolescence. Exposure to air pollutants at this developmental stage can cause more severe respiratory symptoms, including death from respiratory causes, and can cause long-term adverse effects on lung structure, as well as function. Furthermore, because of early exposure and longer life span of children compared to adult, children also have higher risk of developing cancers from air pollutants with carcinogenic property, as shown in Figure 2.

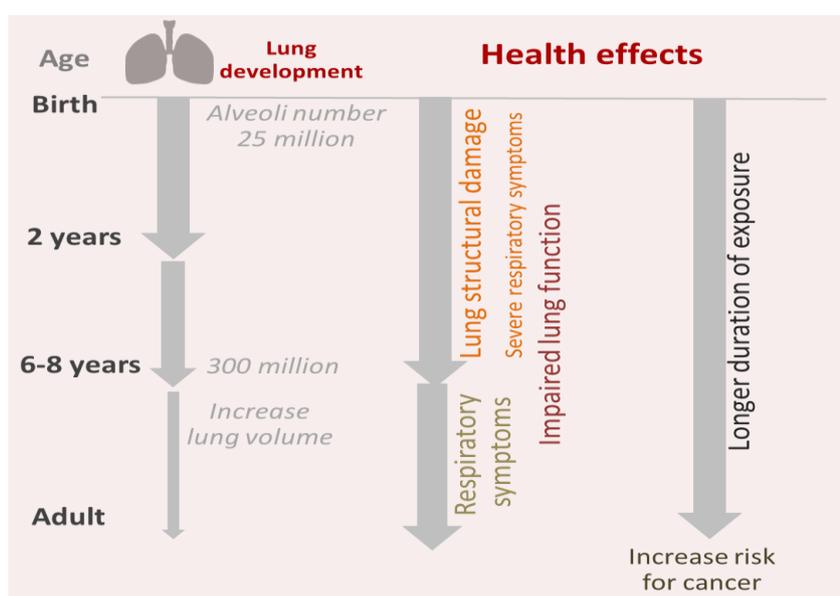


Figure 2: Lung development and risks from air pollution exposure

2.3 FENO (Fractional Exhaled Nitric Oxide)

Nitric oxide is a biological mediator, which is produced in the lung and present in the exhaled breath. Exhaled nitric oxide has been a standardized clinical test. The measurement is quantitative, noninvasive, simple, and safe. Nitric oxide synthase (NOS) is the enzyme required in formation of Nitric oxide.

There are three isoforms of NOS including neuronal cell NOS, inducible NOS, endothelial cell NOS. Inflammator cytokines upregulate inducible NOS, which can generate large quantities of NO.

The American Thoracic Association recommends the use of FENO in the diagnosis of eosinophilic airway inflammation, including asthma. There are also other diseases that can cause elevated level of exhaled NO such as atopy, nonasthmatic eosinophilic bronchitis, chronic obstructive pulmonary disease, exacerbations, noncystic fibrosis bronchiectasis, viral upper respiratory tract infection. Low levels of FENO can occur in cystic fibrosis, smoking, pulmonary hypertension, hypothermia, pulmonary ciliary dysfunction, bronchopulmonary dysplasia. Some of interpretation of FeNO values are shown in Table 4.

Table 4: Clinical guide to interpretation of FeNO values

	Low	Intermediate	High
FENO level for patient <12 years old	<20 ppb	20-35 ppb	>35 ppb
Diagnosis In patient with symptoms during past 6 weeks	Eosinophilic airway inflammation unlikely	Be cautious	Eosinophilic airway inflammation likely
Possible alternative diagnosis	<ul style="list-style-type: none"> ▪ Hyperventiltion ▪ Cardiac disease ▪ COPD ▪ GERD ▪ Noneosinophilic asthma ▪ Rhinosinusitis ▪ Vocal cord dysfunction ▪ Cystic fibrosis ▪ Primary ciliary dyskinesia 	<ul style="list-style-type: none"> ▪ High level of allergen exposure ▪ Infection 	<ul style="list-style-type: none"> ▪ High level of allergen exposure ▪ Infection ▪ Eosinophilic bronchitis

2.4 ADHD (Attention Deficit Hyperactive Disorder)

Attention-deficit/hyperactivity disorder (ADHD) is the most common neurobehavioral disorder of childhood, and can profoundly affect the academic achievement, well-being, and social interactions of children [15].

ADHD is characterized by a persistent pattern of behavior, in multiple settings, that interferes with functioning or development. The behavior is divided into two categories: inattention and hyperactivity-impulsivity. Diagnosis of ADHD is multi-step. The tool used to help in diagnosis is the criteria from DSM-5 [16], as shown in Table 5.

Table 5: DSM-5 criteria for diagnosis of ADHD

DSM-5 Criteria for ADHD
<p>People with ADHD show a persistent pattern of inattention and/or hyperactivity-impulsivity that interferes with functioning or development:</p>
<p>Inattention: Six or more symptoms of inattention for children up to age 16, or five or more for adolescents 17 and older and adults; symptoms of inattention have been present for at least 6 months, and they are inappropriate for developmental level:</p> <ul style="list-style-type: none"> ✓ Often fails to give close attention to details or makes careless mistakes in schoolwork, at work, or with other activities. ✓ Often has trouble holding attention on tasks or play activities. ✓ Often does not seem to listen when spoken to directly. ✓ Often does not follow through on instructions and fails to finish schoolwork, chores, or duties in the workplace (e.g., loses focus, side-tracked). ✓ Often has trouble organizing tasks and activities. ✓ Often avoids, dislikes, or is reluctant to do tasks that require mental effort over a long period of time (such as schoolwork or homework). ✓ Often loses things necessary for tasks and activities (e.g. school materials, pencils, books, tools, wallets, keys, paperwork, eyeglasses, mobile telephones). ✓ Is often easily distracted ✓ Is often forgetful in daily activities.
<p>Hyperactivity and Impulsivity: Six or more symptoms of hyperactivity-impulsivity for children up to age 16, or five or more for adolescents 17 and older and adults; symptoms of hyperactivity-impulsivity have been present for at least 6 months to an extent that is disruptive and inappropriate for the person's developmental level:</p> <ul style="list-style-type: none"> ✓ Often fidgets with or taps hands or feet, or squirms in seat. ✓ Often leaves seat in situations when remaining seated is expected. ✓ Often runs about or climbs in situations where it is not appropriate (adolescents or adults may be limited to feeling restless). ✓ Often unable to play or take part in leisure activities quietly. ✓ Is often "on the go" acting as if "driven by a motor".

- ✓ Often talks excessively.
- ✓ Often blurts out an answer before a question has been completed.
- ✓ Often has trouble waiting his/her turn.
- ✓ Often interrupts or intrudes on others (e.g., butts into conversations or games)

In addition, the following conditions must be met:

- ✓ Several inattentive or hyperactive-impulsive symptoms were present before age 12 years.
- ✓ Several symptoms are present in two or more setting, (e.g., at home, school or work; with friends or relatives; in other activities).
- ✓ There is clear evidence that the symptoms interfere with, or reduce the quality of, social, school, or work functioning.
- ✓ The symptoms do not happen only during the course of schizophrenia or another psychotic disorder. The symptoms are not better explained by another mental disorder (e.g. Mood Disorder, Anxiety Disorder, Dissociative Disorder, or a Personality Disorder).

There have been evidences that, in addition to genetics, other factors that are associated with increased risk for ADHD include brain injury, premature birth, low birth weight, maternal alcohol and tobacco use during pregnancy, environmental exposures such as lead.

Traffic-related air pollution is concerned to have effects on developing brain. There is a cohort study which is a part of the Cincinnati Childhood Allergy and Air Pollution Study (CCAAPS) birth cohort that studied the impact of TRAP exposure on the risk for ADHD was studied using a surrogate elemental carbon attributed to traffic (ECAT). ECAT exposure during the first year of life was estimated based on measurements from 27 air sampling sites. The symptoms of ADHD are assessed at 7 years old by a parents-completed rating scale. The study found that exposure to the highest tertile of ECAT during the child's first year of life was significantly associated with Hyperactivity T-scores in the "at risk" range at 7 years of age, after adjustment [adjusted odds ratio = 1.7; 95% CI: 1.0, 2.7].

CHAPTER 3 RESEARCH PLAN

3.1 Project Schedule

Table 6 shows the project planning schedule. All project members are scheduled to meet regularly to discuss the technical results performed by project research assistant, and directions of the project. Occasionally, the progress report will be presented to the advisors to further seek guidelines and comments of the results and future direction.

Table 6: Project planning schedule

Activity	2015									2016		
	Apr	May	Jun	Jul	Aug	Sep	Oct	Nov	Dec	Jan	Feb	Mar
Literature review												
Questionnaire validation												
Collect data												
Statistical analysis												
Inception report submission	30-Apr											
Progress report presentation				3-Jul								
Interim report presentation						11-Sep						
Interim report submission						30-Sep						
Final report presentation									12-Dec			
Roundtable discussion/workshop										29-Jan		
Final report submission												31-Mar

3.2 Project Expenditure

Table 7 shows the breakdown of the project expenditure, which is mainly composed of the participation of the members and advisors. One research assistants (RAs) will be employed on the part time basis. Main expenses are for remaining measurements of indoor air pollutants from last year project; whereas, other necessary expenses such as transportation to gather data and office/computer supply are included. The project aims to present the preliminary result at a conference upon approval from ATRANS. Lastly, the expenses of secretariat's participation and report publishing are included.

Table 7: Project expenditure

No.	Item	Unit cost	Number of units	Sub total
1	Project leader & co-leader	3,000	12	36,000
2	Advisors participation in project meeting (200 THB/hr x 5 hrs/day x 5 days/month) for 12 months)	5,000	12	60,000
3	Expenses for project meeting	3,000	6	18,000

No.	Item	Unit cost	Number of units	Sub total
4	Expenses for indoor air quality measurement at DCCs	200,000	1	200,000
5	Office & computer supply	3,000	6	18,000
6	Secretariat's participation portion	10,000	1	10,000
7	Publishing proportion of the report book	50,000	1	50,000
Total				392,000

CHAPTER 4 STUDY PROCESS

4.1 Inclusion of day care centers and children

Eleven from approximately 300 day care centers in Bangkok were included into the study. We used computer-generated stratified random sampling to select day care centers from each region of Bangkok. The distribution of day care centers are as shown in Figure 3. Each day care center was contacted and invited to participate in the study. Last year project, a meeting was held in May 2014 to give background information on the study as well as detailed information on data collection and study process. Informed consents were obtained from every day care center. The day care centers included are listed in Table 8. Parents of all children in the 11 day care centers who fit the inclusion criteria listed above were sent an invitation letter to join in the study.

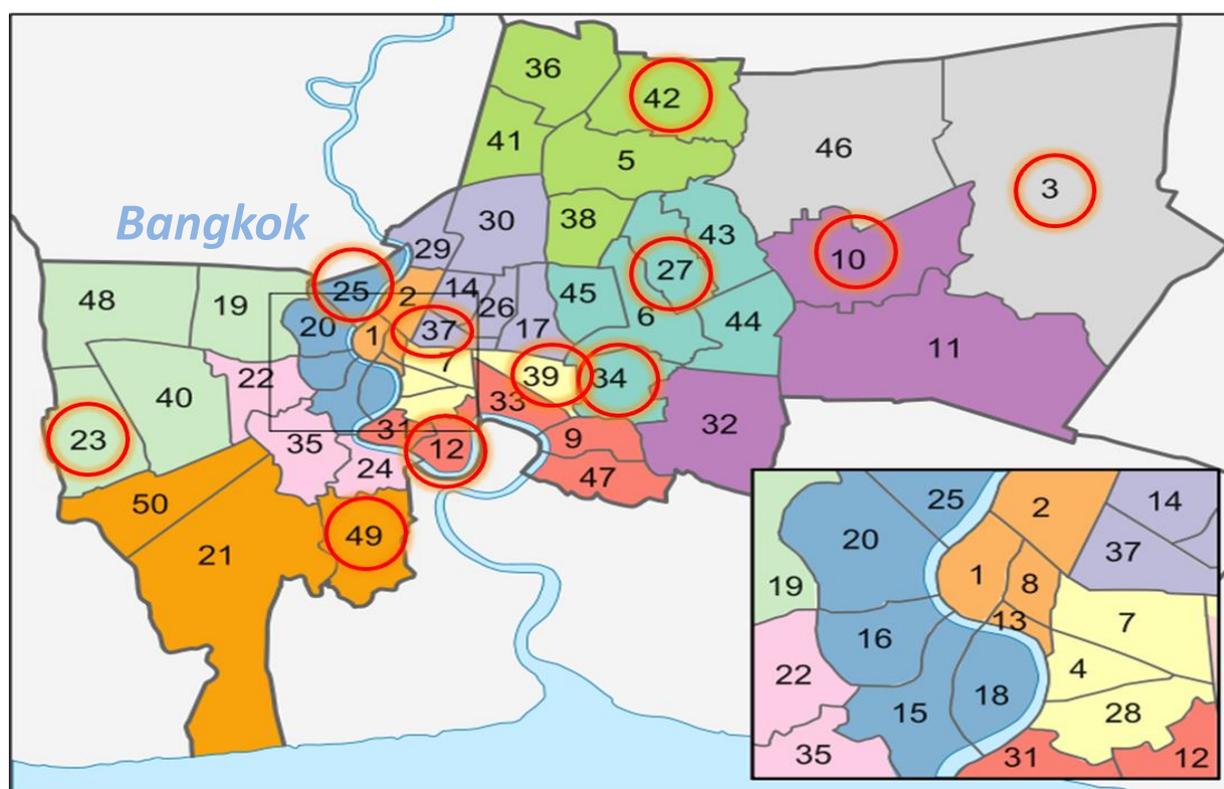


Figure 3: Distribution of Day Care Centers in the Study

Table 8: List of day care centers included in the study

Day Care Center	District	No. of children eligible	No. of children included	Response rate
DCC #1	Nong Jok	85	84	98.8%
DCC #2	Suan Luang	21	21	100%
DCC #3	Mean Buri	22	22	100%
DCC #4	Bung Koom	40	40	100%
DCC #5	Sai Mhai	20	15	75%
DCC #6	Yan Nawa	30	30	100%
DCC #7	Wattana	25	18	72%
DCC #8	Toong Kru	21	21	100%
DCC #9	Bangkok Noi	40	40	100%
DCC #10	Rachathewi	70	70	100%
DCC #11	Nong Khaem	105	75	71.4%

4.2 Day care center visit and survey of environment

The day care centers were re-visited from last year project during their operation time. The research team surveyed every room in each day care center and nearby area, and recorded data on the environmental characteristic record form including total indoor area, type of building material, ventilation system, proximity to traffic or water sources, visible signs of dampness or mold. Some information such as year of construction, number of children, indoor cooking, presence of indoor pets were obtained by interviewing of day care center staff.

4.3 The Questionnaires

Similar to last year, Figure 4 show three sets of questionnaires and one teacher record form used in the process of data collection in this study, including:



แบบสอบถามสิ่งแวดล้อมในบ้าน และอาการทางระบบทางเดินหายใจของเด็กอายุต่ำกว่า 6 ปี
Home environment and respiratory symptoms questionnaire

ข้อมูลทั่วไปของเด็กในครัวเรือน (Basic information)

- ชื่อศูนย์พัฒนาเด็กก่อนวัยเรียน (name of day care center) _____
เด็กในครัวเรือนที่ออกนอกบ้านเข้าศูนย์พัฒนาเด็กก่อนวัยเรียนมาเป็นระยะเวลา _____ ปี เดือน _____
(your child has attended the day care center for _____ year _____ month)
- ชื่อเด็ก (child's name) _____ อายุของเด็ก (age) _____ ปี (years)
วันเดือนปีเกิดของเด็ก (child's date of birth) _____
เพศของเด็ก (gender) ชาย (male) หญิง (female)
น้ำหนักของเด็ก (weight) _____ กิโลกรัม (kg) ส่วนสูงของเด็ก (height) _____ เซนติเมตร (cm)
น้ำหนักแรกเกิดของเด็ก (birth weight) _____ กิโลกรัม (kg) มีพี่น้องรวม (number of siblings) _____ คน
- โรคประจำตัวของเด็ก (child's underlying disease) ไม่มี (no) มี (yes)
ระบุ (please specify) _____
ประวัติการแพ้ยา แพ้อาหารของเด็ก (child's history of drug/food allergy) ไม่มี (no) มี (yes)
ระบุ (please specify) _____
- ใน 1 ปีที่ผ่านมาเด็กได้รับวัคซีนป้องกันโรคไขหวัดใหญ่หรือไม่ (During the past one year, has your child received influenza vaccine?)
 ได้รับ (yes) ไม่ได้รับ (no)
- เด็กเคยได้รับนมมารดาเลี้ยงหรือไม่ (Had the child ever been breast-fed?)
 เคย (yes) นาน (for) _____ เดือน (months) ไม่เคย (no)

ข้อมูลครอบครัวและสิ่งแวดล้อมในบ้าน

- จำนวนคนที่อาศัยในบ้าน (number of house members) _____ คน
จำนวนผู้ใหญ่ (number of adults) _____ คน
เด็กอายุต่ำกว่า 6 ปี (number of children under 6 years old) _____ คน
เด็กอายุมากกว่า 6 ปี (number of children over 6 years old) _____ คน
- ผู้ดูแลหลักของเด็ก (who is the child's main care taker?) คือ บิดา (father) มารดา (mother) อื่นๆ (other) ระบุ please specify _____

บิดาของเด็กสูบบุหรี่หรือไม่ (Does the child's father smoke cigarettes?)

- ใช่ (yes) ไม่ใช่ (no)
ถ้าใช่ บิดาของเด็กสูบบุหรี่ประมาณวันละ _____ มวน
(if yes, how many cigarettes does the child's father smoke each day?) _____ (cigarettes)

- บ้านอยู่ห่างจากศูนย์พัฒนาเด็กก่อนวัยเรียนเป็นระยะทางประมาณ _____ กิโลเมตร
(The distance between your home and the day care center is about _____ kilometer)
เดินทางโดย _____ (mean of transportation)

- สัตว์เลี้ยงในบ้าน (Presence of pet indoors)
ในช่วง 1 ปีที่ผ่านมา, คุณมีสัตว์เลี้ยงในบ้านหรือไม่ (in the past 1 year, have you had these pets in your home?)
สุนัข (dogs) มี (yes) ไม่มี (no) แมว (Cats) มี (yes) ไม่มี (no)
ในช่วง 1 ปีแรกของชีวิตของคุณสัตว์เลี้ยงเหล่านี้ในบ้านหรือไม่ (during the first year of your child's life, have you had these pets in your home?)
สุนัข (dogs) มี (yes) ไม่มี (no) แมว (Cats) มี (yes) ไม่มี (no)

- ท่านใช้เครื่องปรับอากาศในบ้านหรือไม่ (Do you use air-conditioner in your home?)
 ใช่ เครื่องปรับอากาศ (Use air-conditioner)
 ใช้พัดลม (use electrical fan) ไม่มี (none)

- ท่านใช้เชื้อเพลิงชนิดใดในการประกอบอาหารในบ้านของท่าน (in your house, what fuel is usually used for cooking?)
 ไฟฟ้า (electricity) แก๊ส (gas) ก๊าซโพรเพน (open fire)

- ถนนบ้านของท่านมีรถบรรทุกที่ผ่านไปมาบ่อยเพียงใด (how often do trucks pass through the street where you live?)
 ไม่มีเลย (never) นานๆ ครั้ง (seldom)
 มีบ้างในแต่ละวัน (Frequently through the day) มีเกือบทั้งวัน (Almost the whole day)

- บ้านของคุณมีรอยน้ำรั่ว หรือรอยเลอะในบ้านบ้างหรือไม่ (Do you have any water leakage or visible sign of molds in your home?)
 มี (yes) ไม่มี (no)

3. ระดับการศึกษา (Education)

- บิดา (Child's father)
 น้อยกว่าประถมศึกษา (lower than primary school) ประถมศึกษา (primary school)
 มัธยมศึกษา ปว. ปวส. (Secondary school) ปริญญาตรี Bachelor degree
 สูงกว่าปริญญาตรี (higher than Bachelor degree)

- มารดา (Child's mother)
 น้อยกว่าประถมศึกษา (lower than primary school) ประถมศึกษา (primary school)
 มัธยมศึกษา ปว. ปวส. (Secondary school) ปริญญาตรี Bachelor degree
 สูงกว่าปริญญาตรี (higher than Bachelor degree)

- ผู้ดูแลหลัก (หากผู้ดูแลหลักเป็นบิดาหรือมารดา ไม่ต้องตอบ)
Child's main care taker (if the main care taker is father or mother, skip to 4.)
 น้อยกว่าประถมศึกษา (lower than primary school) ประถมศึกษา (primary school)
 มัธยมศึกษา ปว. ปวส. (Secondary school) ปริญญาตรี Bachelor degree
 สูงกว่าปริญญาตรี (higher than Bachelor degree)

- รายได้รวมเฉลี่ยต่อเดือนของครอบครัว (total monthly family income)
 <10,000 บาท (Baht) 10,000-30,000 บาท (Baht) >30,000 บาท (Baht)

- ประวัติโรคภูมิแพ้ของบิดามารดา (parents allergic diseases)
บิดา (Child's father)
 แพ้อาหาร (hay fever) หอบหืด (asthma)
 ผื่นแพ้ผิวหนัง (eczema) ไม่มี (none)

- มารดา (Child's mother)
 แพ้อาหาร (hay fever) หอบหืด (asthma)
 ผื่นแพ้ผิวหนัง (eczema) ไม่มี (none)

- การสูบบุหรี่ในบ้าน (indoor cigarette smoking) มี (yes) ไม่มี (no)
จำนวนคนที่สูบบุหรี่ (Number of smokers in home) _____ คน (persons)

- มารดาของเด็กสูบบุหรี่หรือไม่ (Does the child's mother smoke cigarettes?)
 ใช่ (yes) ไม่ใช่ (no)
ถ้าใช่ มารดาของเด็กสูบบุหรี่ประมาณวันละ _____ มวน
(if yes, how many cigarettes does the child's mother smoke each day?) _____ (cigarettes)

- มารดาของเด็กสูบบุหรี่ในช่วงขวบปีแรกของเด็กหรือไม่ (Does the child's mother smoke cigarettes during the child's first year of life?) ใช่ (yes) ไม่ใช่ (no)

ข้อมูลเกี่ยวกับอาการหอบ (Questionnaire for asthma)

กรุณาทำเครื่องหมาย ✓ ในช่องคำตอบที่ตรงกับอาการของคุณ (Please put a tick ✓ in your answer box)

- ลูกของคุณเคยหายใจเสียงวี๊ด หรือหายใจมีเสียงคล้ายนกหวีดในทรวงอก (หน้าอก) ของลูกคุณมาก่อนหรือไม่
หากไม่เคย เข้าไปข้อ 8
(Has your child ever had wheezing or whistling in the chest at any time in the past?)
(IF YOU HAV ANSWERED "NO" PLEASE SKIP TO QUESTION 8)

- ในช่วงเวลา 12 เดือนที่ผ่านมา ลูกของคุณเคยหายใจเสียงวี๊ด หรือหายใจมีเสียงคล้าย นกหวีดในทรวงอกบ้างหรือไม่
หากไม่เคย เข้าไปข้อ 6
(Has your child had wheezing or whistling in the chest in the past 12 months?)
(IF YOU HAV ANSWERED "NO" PLEASE SKIP TO QUESTION 6)

- ในช่วง 12 เดือนที่ผ่านมา ลูกของคุณมีอาการหายใจเสียงวี๊ด หรือมีอาการหอบหืดเป็นจำนวนกี่ครั้ง
(How many attacks of wheezing has your child had in the past 12 months?)
 1-3 ครั้ง (1-3 times)
 4-12 ครั้ง (4-12 times)
 มากกว่า 12 ครั้ง (over 12)

- ในช่วง 12 เดือนที่ผ่านมา ลูกของคุณเคยตื่นขึ้นมาจากการนอนหลับเพราะหายใจเสียงวี๊ดโดยเฉลี่ยกี่ครั้งต่อ 1 สัปดาห์
(in the past 12 months, how often, on average, has your child's sleep been disturbed due to wheezing?)
 ไม่เคย (never woken with wheezing)
 น้อยกว่า 1 ครั้ง/สัปดาห์ (Less than one night per week)
 เท่ากับหรือมากกว่า 1 ครั้ง/สัปดาห์ (one or more nights per week)

- ในช่วง 12 เดือนที่ผ่านมา ลูกของคุณเคยมีอาการหายใจเสียงวี๊ดรุนแรงจนต้องหยุดพูดและทำให้พูดได้เพียง 1-2 คำต่อเนื่องกันแล้วต้องหยุดหายใจหรือไม่
(in the past 12 months, has wheezing ever been severe enough to limit your child's speech to only one or two words at a time between breaths?)
 ใช่ (yes)
 ไม่เคย (No)

6. ลูกของคุณเคยเป็นโรคหอบหืดหรือไม่ เคย (yes) ไม่เคย (no)
(Has your child ever had asthma?)
7. ในช่วง 12 เดือนที่ผ่านมา ลูกของคุณเคยหายใจเสียงวี๊ดในระหว่างหรือหลังจากการออกกำลังกายหรือไม่ เคย (yes) ไม่เคย (no)
(In the past 12 months, has your child's chest sounded wheezy during or after exercise?)
8. ในช่วง 12 เดือนที่ผ่านมา ลูกของคุณเคยมีอาการไอแห้งๆ ในเวลากลางคืน โดยไม่ได้เป็นหวัดและไม่ได้อาการติดเชื้อของระบบทางเดินหายใจมาก่อนหรือไม่ เคย (yes) ไม่เคย (no)
(In the past 12 months, has your child had a dry cough at night, apart from a cough associated with a cold or chest infection?)

ข้อมูลเกี่ยวกับอาการหวัด (Questionnaire for rhinitis)

กรุณาทำเครื่องหมาย ✓ ลงในช่องคำตอบที่ตรงกับอาการของลูกของคุณ
(Please put a tick ✓ in your answer box)

1. ลูกของคุณเคยมีอาการจาม น้ำมูกไหล แน่นจมูกหรือคัดจมูกในขณะที่ลูกของคุณไม่ได้เป็นหวัดหรือไข้หวัดใหญ่บ้างหรือไม่ เคย (yes) ไม่เคย (no)
(Has your child ever had a problem with sneezing, or a runny, or a blocked nose when he/she DID NOT have a cold or the flu?)
- หากไม่เคย ข้ามไปข้อ 5 (IF YOU HAVE ANSWERED "NO" PLEASE SKIP TO QUESTION 5)
2. ในช่วง 12 เดือนที่ผ่านมา ลูกของคุณเคยมีอาการจาม น้ำมูกไหล แน่นจมูกหรือคัดจมูกในขณะที่ลูกของคุณไม่ได้เป็นหวัดหรือไข้หวัดใหญ่บ้างหรือไม่ เคย (yes) ไม่เคย (no)
(In the past 12 months, has your child had a problem with sneezing, or a runny, or a blocked nose when he/she DID NOT have a cold or the flu?)
- หากไม่เคย ข้ามไปข้อ 5 (IF YOU HAVE ANSWERED "NO" PLEASE SKIP TO QUESTION 5)
3. ในช่วง 12 เดือนที่ผ่านมา อาการของลูกของคุณเคยเกิดขึ้นร่วมกับอาการคันตาและน้ำตาไหลหรือไม่ เคย (yes) ไม่เคย (no)
(In the past 12 months, has this nose problem been accompanied by itchy/watery eyes?)
4. ในช่วง 12 เดือนที่ผ่านมา ปัญหาทางจามรบกวนกิจวัตรประจำวันของลูกของคุณมากน้อย ไม่รบกวน (not at all) เล็กน้อย (a little) ปานกลาง (a moderate amount) มาก (a lot)
(In the past 12 months, how much did this nose problem interfere with your child's daily activities?)
5. ลูกของคุณเป็นโรคแพ้อากาศหรือไม่ เป็น (yes) ไม่เป็น (no)
(Has your child ever had hayfever?)

(a)



แบบสอบถามอาการทางระบบทางเดินหายใจของเด็กโดยผู้ปกครอง
Respiratory symptoms questionnaire by parents

ชื่อ (name) _____ อายุ (age) _____ ที่อยู่ (location) _____
*ในช่วง 1 สัปดาห์ที่ผ่านมา ลูกของคุณเคยมาที่ศูนย์ดูแลเด็กหรือไม่กี่วัน _____ วัน
(*For the past week, your child has been to the day care center for _____ days)

กรุณาทำเครื่องหมาย ✓ ลงในช่องคำตอบที่ตรงกับอาการของลูกของคุณ (Please put a tick ✓ in your answer box)

- ในช่วง 1 สัปดาห์ที่ผ่านมา (During the past 1 week)
- ลูกของคุณเคยมีอาการคัดจมูก (Has your child had blocked nose?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยมีอาการจาม (Has your child had sneezing?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยมีอาการคันจมูก (Has your child had itchy nose?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยมีอาการน้ำมูกไหล (Has your child had runny nose?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยมีอาการไอ (Has your child had cough?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยมีอาการหอบ (Has your child had shortness of breath?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยมีอาการตัวร้อน/ไข้ (Has your child had fever?)
 ใช่ (yes) ไม่ใช่ (No)
 - ลูกของคุณเคยเรียนเพราะอาการใด ๆ หนึ่งข้อ 1-7 หรือไม่ (Has your child been absent from class because of any symptoms from 1-7?)
 ใช่ (yes) ไม่ใช่ (No)
ถ้าใช่ จำนวนวันที่ขาด _____ วัน (If yes, the number of absent days is _____ day(s))
 - ในช่วง 1 สัปดาห์ที่ผ่านมา ลูกของคุณเคยไปพบแพทย์เนื่องจากอาการใด ๆ หนึ่งข้อ 1-7 หรือไม่ (Has your child visited a doctor because of any symptoms from 1-7?)

ใช่ (yes) ไม่ใช่ (No)

ถ้าใช่ การวินิจฉัย (if yes, diagnosis was) _____

ยาที่ได้รับ (medication received) _____

10. ในช่วง 1 สัปดาห์ที่ผ่านมา ลูกของคุณเคยนอนโรงพยาบาลเนื่องจากอาการใด ๆ หนึ่งข้อ 1-7 หรือไม่ (Has your child been admitted to a hospital because of any symptoms from 1-7?)

ใช่ (yes) ไม่ใช่ (No)

ถ้าใช่ การวินิจฉัย (if yes, diagnosis was) _____

จำนวนวันที่นอนโรงพยาบาล (length of hospital stay) _____ วัน (day(s))

(b)

Teacher Respiratory symptoms Report

แบบบันทึกอาการทางระบบทางเดินหายใจรายสัปดาห์โดยครู

ชื่อ(Name) _____ อายุ (Age) _____ ปี (Year)

คำชี้แจง: บันทึกอาการทางระบบทางเดินหายใจของนักเรียน สัปดาห์ละครั้งทุกวันจันทร์ รวม 4 สัปดาห์

กรุณาทำเครื่องหมาย ✓ ลงในช่องหากเด็กมีอาการดังกล่าว ในวันที่ทำการบันทึก

Instructions :Please record respiratory symptoms of the student once a week on every Monday for 4 weeks,

Please check ✓ in the box if the student has these following symptoms

อาการ(Symptoms)	คันหรือเคืองตา หรือตาแดง ตาและ (Itchy or irritated eye/red eye/eye discharge)	คัดจมูก (Blocked nose)	คันจมูก (itchy nose)	จาม (Sneezing)	น้ำมูกไหล (Runny Nose)	ไอ (Cough)	หอบ (Shortness of breath)	ตัวร้อนหรือมีไข้ (Fever)
วันที่ทำการบันทึก (Date)								
วันจันทร์ที่ _____ (Monday)								
วันจันทร์ที่ _____ (Monday)								
วันจันทร์ที่ _____ (Monday)								
วันจันทร์ที่ _____ (Monday)								

(c)

Figure 4: Questionnaires for (a) baseline characteristic and home environment, (b) parent respiratory symptoms and (c) teacher respiratory symptoms

4.3.1 Baseline Characteristic and Home Environment Questionnaire

This questionnaire was distributed at the beginning of the study. The questionnaire is divided into 4 parts. The first part contained questions on the child's general information and health status such as date of birth, current height and weight, birth weight, underlying diseases, drug or food allergy, number of siblings, history and duration of breast feeding, immunization for influenza in the past year, duration of attendant to day care center. The second part focuses on family background and home environment. Questions consist of total number of people in the household, relationship of main caretaker with the child, education level of parents and main caretaker, family income, maternal and paternal atopic diseases, indoor smoking, paternal and maternal smoking, indoor pets exposure, indoor cooking fuels, home ventilation system, visible sign of water leakage or mold, home proximity to traffic, transportation to daycare center, distance between home and day care center. The third and the fourth parts of the questionnaire are about symptoms of asthma and allergic rhinitis, respectively.

4.3.2. Parent Respiratory Symptoms Questionnaire

This questionnaire was distributed weekly for 4 weeks during each season of data collection. The questions are on parent-report respiratory symptoms including blocked nose, sneezing, itchy nose, runny nose, cough, shortness of breath, and also fever to suggest infectious etiology. Questions on the number of days the child was presented to the day care center and the number of days absent for respiratory illness during the week

were also included, as well as doctor visit attributable to respiratory illness, diagnosis, prescribed medication, hospital admission and length of hospital stay.

The two questionnaires above were pilot tested on 30 parents of children attending Ramathibodi hospital's day care center before distribution to parents of all children in the study.

4.3.3. Teacher Respiratory Symptom Record Form

Teachers record respiratory symptoms of children including blocked nose, sneezing, itchy nose, runny nose, cough, shortness of breath, and fever on every Monday for 4 consecutive weeks. The record also includes information on any medication the child took during the day at the day care center.

4.3.4 “Thai ADHD screening scale”

“Thai ADHD screening scale” is a standard questionnaire developed and copyrighted by Department of Psychiatry, Faculty of medicine, Siriraj hospital. The project uses the questionnaire in this study with permission from the authors. The questionnaires will be filled out by parents and teachers at the end of the study. The questionnaire will be scored and compared to the T-score standards, and results are report as the level of risk for ADHD.

4.4 Pollutant measurement

During each day care center visit, research member team collect air sample from the children's room and measure for pollutants, as shown in Figure 5. PM₁₀ was measured by gravimetric techniques. Measurement of benzene, CO and Ozone use gas chromatography technique, whereas NO₂ and SO₂ use ion chromatography technique. For biological pollutants, bacterial and fungus count are carried out by air sampling equipment connected to a cell-culture dish container, microbes then are incubated in the laboratory and measured for colony forming units. Specimen for dust mite antigen level measurement is collected from children's bedding material using vacuum pump and send for laboratory analysis.

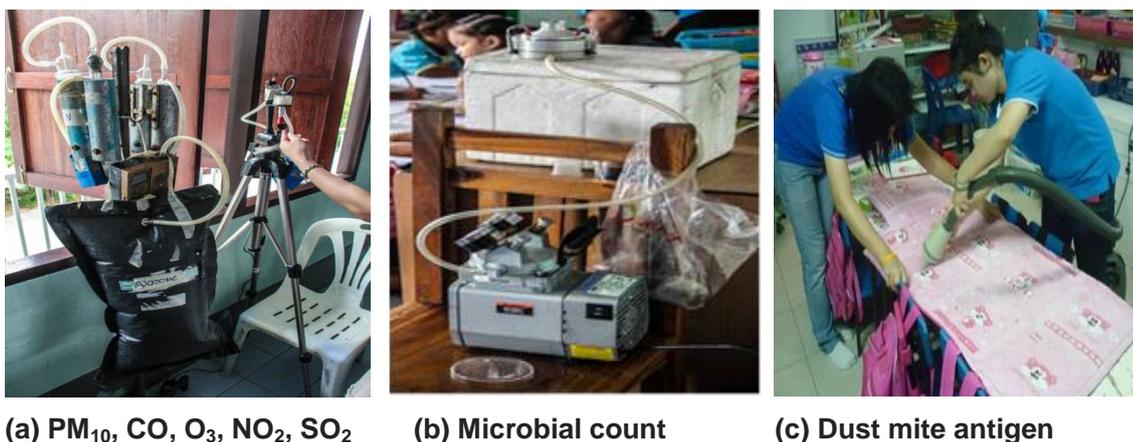


Figure 5: Specimen collection for pollutant measurement

4.5 FENO measurement

Ten children were randomly selected from each day care center and scheduled for exhaled nitric oxide level measurement. Written informed consents by parents were acquired. Details on the test procedure were provided to parents before the scheduled test by letters, and also by demonstration of the process before performing the test. The technique used in children is multiple-breath FENO measurement, requiring each child to breath into a face mask for one minute, as shown in Figure 6. The tests were performed three times on each child to obtain the mean FENO values.



Figure 6: FENO measurement

CHAPTER 5 RESULTS

5.1 Day care center environmental characteristics

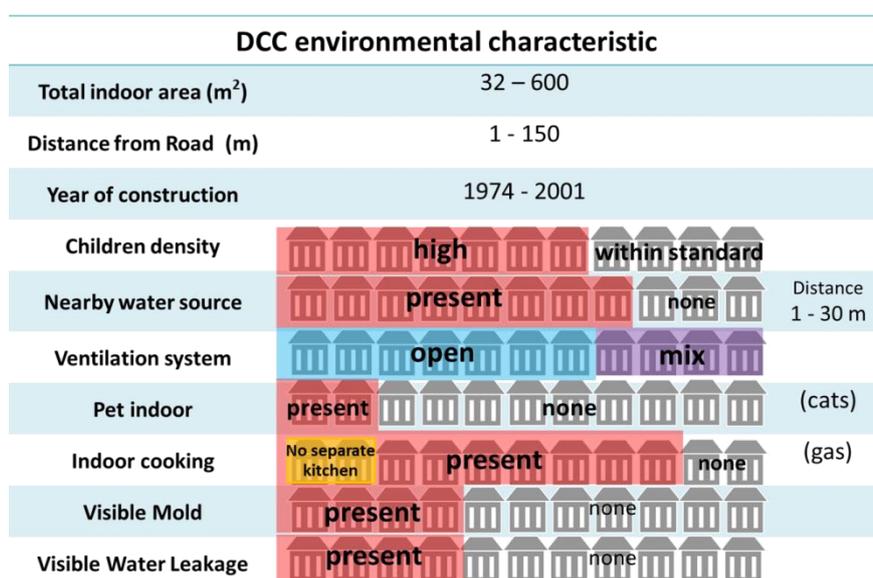
As shown in Figure 7, the size of indoor area of the day care centers in this study range from 32 to 600 meter square (median = 240, IQR = 56 , 400). According to Thailand day care center standard formulated by Ministry of Public Health, each child should have a space of at least 2 meter square. Therefore, children density should be lower than 0.5 person per meter square. In this study, children density range from 0.27 to 2.34 persons per meter square. We find that 6 out of 11 (54.5%) day care centers have higher children density than the standard recommendation.

The building materials of all day care centers in our study include concrete, 36.4% were built from concrete and wood in combination. The time since construction of these day care centers ranges from 13 to 40 years (Median = 19, IQR = 17,30). Most day care centers (81.8%) are within 100 meters distance from road traffic (Median distance = 20, IQR = 2,100). There is one day care center located very close to the railroad traffic. 7 from 11 day care centers (63.6%) have a water resource in the nearby area.

For ventilation system, open ventilation is used exclusively in 7 day care centers, while the rest use both open and mechanical (mixed) ventilation system. Indoor cats are also reported in two day care centers. Total number of 9 day care centers have indoor cooking activity, mostly in a separate kitchen, there are two day cares center that cook with no separate kitchen. Cooking fuel used is gas in all day care centers. More than half (54.5%) of the day care centers in this study have visible sign of dampness or mold indoor.

Between Rainy season and winter, a day care center (DCC5) was temporary closed and data collection was continued in the remaining 10 day care centers in winter and summer.

Figure 7: Environmental characteristics data of day care centers



5.2 Pollutant level

The pollutant levels and seasonal variation are shown in Table 9 and compared in Figure 8, which show that PM₁₀ levels were higher than WHO indoor air quality standard level (50 mcg/m³) in 7 DCCs in rainy season and 6 DCCs in winter and summer.

For chemical pollutants, indoor levels of CO and NO₂ in the day care centers were within the standard values in all seasons. SO₂ levels are slightly higher than recommended level in 3 day care centers in rainy season and summer, and in 1 day care center in winter. Ozone level in rainy season and summer in all DCCs were within normal limit. In winter, ozone level exceeded standard in 8 out of 11 DCCs.

For benzene, due to its carcinogenic effect, World Health Organization states that no safe level of exposure can be recommended. Studies revealed that every 1 mcg/m³ of benzene concentration, excess lifetime risk for leukemia increase by 6×10^{-6} . In Hong Kong, the airborne benzene level of less than 0.0161 mg/m³ is used as an objective for good indoor air quality. In our study, there were two day care centers with benzene level exceeding 0.0161 mg/m³ in rainy season. While in winter and summer, benzene levels were within standard in all day care centers.

Analysis of seasonal variation of pollutant levels revealed significantly higher level of ozone in winter when compared with other seasons ($p < 0.001$). For other pollutants, this study found no significant variation in indoor pollutant levels.

Table 9: Pollutants level data

Indoor Pollutants Levels (Rainy): Jun 2014						
DCC	Particulate matter and Chemicals (mg/m ³)					
	PM ₁₀	Benzene	O ₃	NO ₂	SO ₂	CO
DCC 1	0.081	<0.003	0.018	0.018	0.012	0.502
DCC 2	0.042	<0.003	0.028	0.025	0.014	0.707
DCC 3	0.076	<0.003	0.014	0.020	0.010	0.536
DCC 4	0.082	<0.003	0.012	0.028	0.020	0.524
DCC 5	0.042	<0.003	0.010	0.018	0.010	0.741
DCC 6	0.041	<0.003	0.024	0.032	0.021	0.638
DCC 7	0.168	0.105	0.010	0.018	0.011	0.536
DCC 8	0.084	0.054	0.024	0.015	0.014	0.729
DCC 9	0.089	<0.003	0.044	0.015	0.020	0.559
DCC 10	0.021	<0.003	0.056	0.020	0.011	0.901
DCC 11	0.133	<0.003	0.008	0.028	0.014	0.410
WHO Standard indoor air	0.050 (24hr)	-	0.10 (8hr)	0.040	0.020 (24 hr)	10 (8 hr.)

(a)

Indoor Pollutants Levels (Winter): Nov 2014

DCC	Particulate matter and Chemicals (mg/m ³)					
	PM ₁₀	Benzene	O ₃	NO ₂	SO ₂	CO
DCC 1	0.039	<0.003	0.156	0.020	0.021	0.78
DCC 2	0.058	<0.003	0.196	0.015	0.012	1.10
DCC 3	0.042	<0.003	0.088	0.022	0.012	0.75
DCC 4	0.063	<0.003	0.110	0.020	0.018	0.70
DCC 6	0.205	<0.003	0.124	0.025	0.014	0.97
DCC 7	0.021	<0.003	0.150	0.028	0.014	1.43
DCC 8	0.042	<0.003	0.184	0.024	0.016	0.67
DCC 9	0.105	<0.003	0.164	0.022	0.015	1.17
DCC 10	0.097	<0.003	0.104	0.018	0.012	1.12
DCC 11	0.084	<0.003	0.044	0.018	0.012	0.56
WHO Standard indoor air	0.050 (24hr)	-	0.10 (8hr)	0.040	0.020 (24 hr)	10 (8 hr.)

* DCC 5 was temporary closed and under renovation during winter

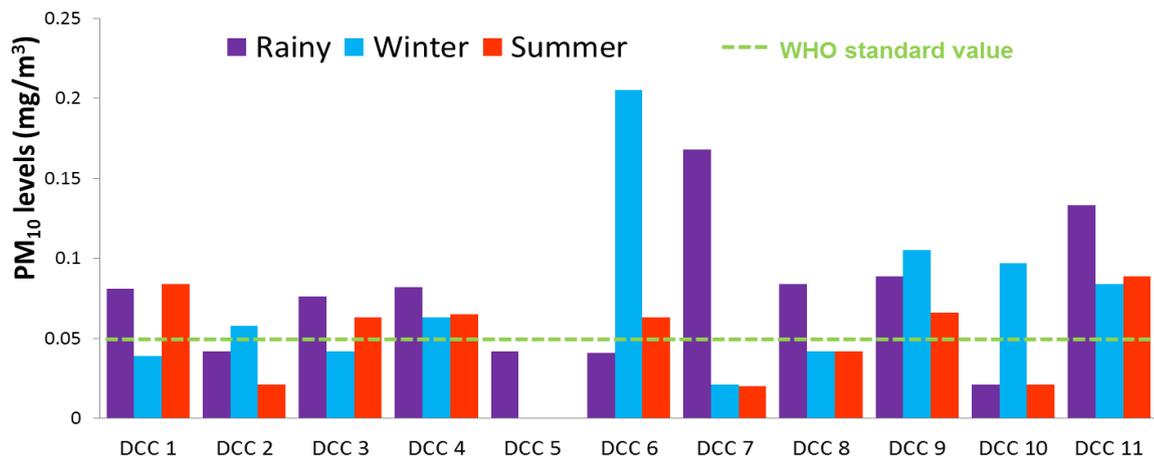
(b)

Indoor Pollutants Levels (Summer): May 2015

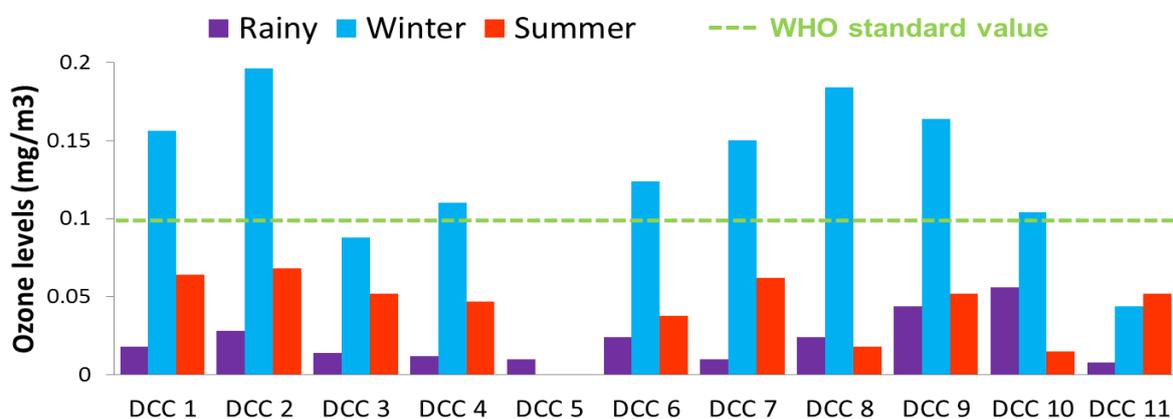
DCC	Particulate matter and Chemicals (mg/m ³)					
	PM ₁₀	Benzene	O ₃	NO ₂	SO ₂	CO
DCC 1	0.084	<0.003	0.064	0.022	0.018	0.79
DCC 2	0.021	<0.003	0.068	0.024	0.014	0.59
DCC 3	0.063	<0.003	0.052	0.024	0.016	0.68
DCC 4	0.065	<0.003	0.047	0.025	0.022	1.18
DCC 6	0.063	<0.003	0.038	0.028	0.02	1.56
DCC 7	0.02	<0.003	0.062	0.026	0.018	0.55
DCC 8	0.042	<0.003	0.018	0.025	0.014	0.45
DCC 9	0.066	<0.003	0.052	0.024	0.015	0.37
DCC 10	0.021	<0.003	0.015	0.032	0.017	0.75
DCC 11	0.089	<0.003	0.052	0.022	0.02	0.25
WHO Standard indoor air	0.050 (24hr)	-	0.10 (8hr)	0.040	0.020 (24 hr)	10 (8 hr.)

* DCC 5 was temporary closed and under renovation during winter and summer

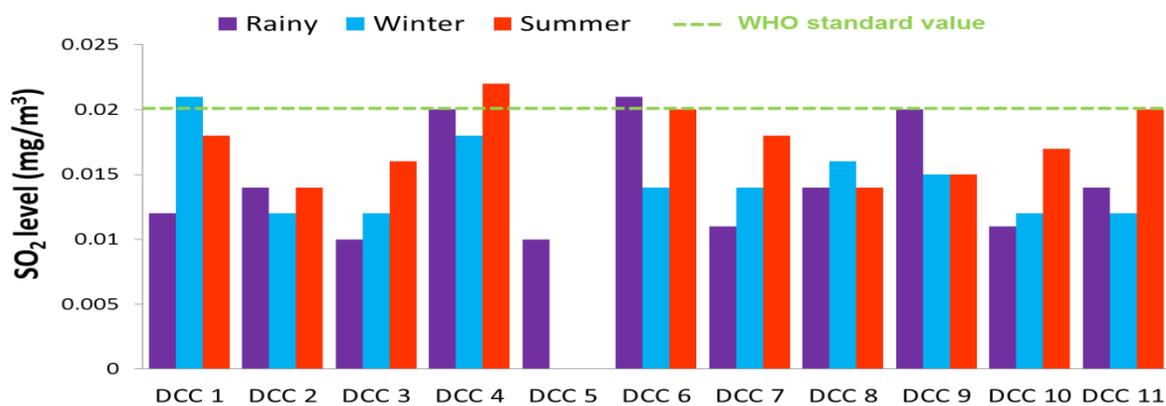
(c)



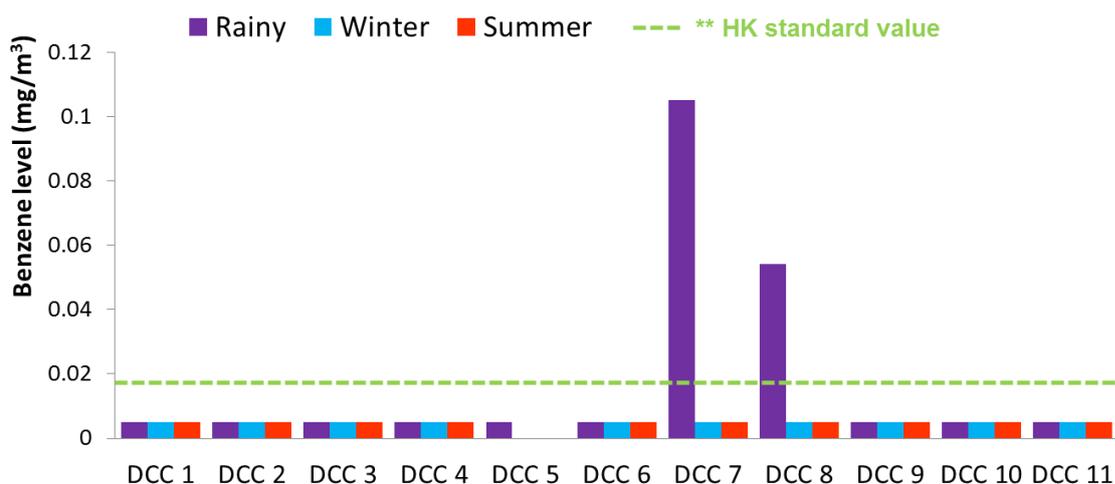
(a)



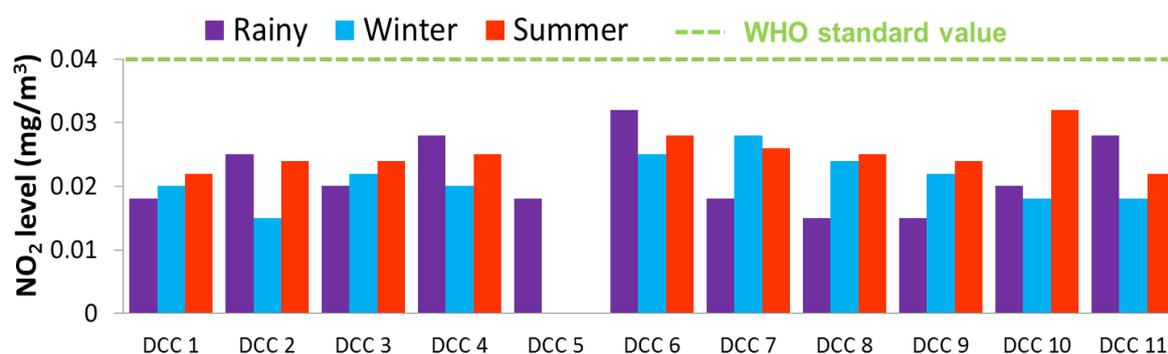
(b)



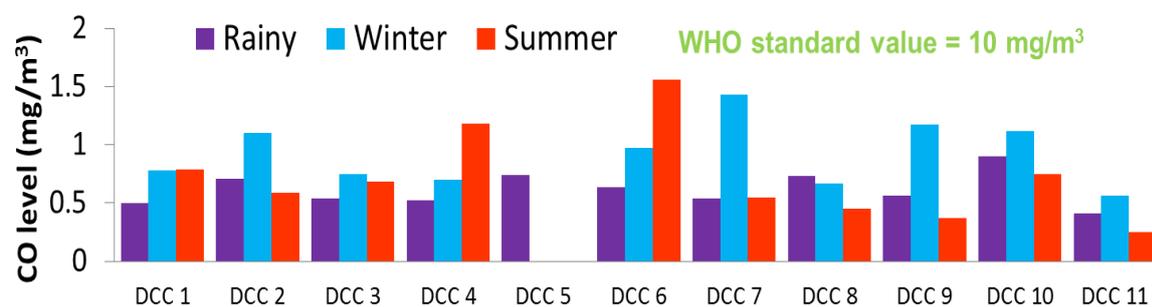
(c)



(d)



(e)



(f)

Figure 8: Levels and seasonal trends of pollutants: (a) PM10, (b) ozone, (c) SO2, (d) Benzene, (e) NO2 and (f) CO in all DCCs during rainy and winter seasons

In order to investigate the effect of outdoor pollutants to indoor level, outdoor and indoor pollutants are compared in 4 DCCs located not far away from Thai Pollution Control Department’s pollution monitoring stations, as shown in Figure 9, where one DCC is as close as 40 meters from the pollution monitoring station. The comparison of indoor and outdoor pollutant levels are shown in Figure 10. Most indoor pollutant levels are within the range between minimum and maximum outdoor levels, except for PM₁₀ levels, which indoor levels exceed maximum outdoor levels in 2 DCCs. The other 7 DCCs are located far away from pollution monitoring station (over 4 kilometers); therefore, the outdoor and indoor levels might not be comparable in these DCCs.

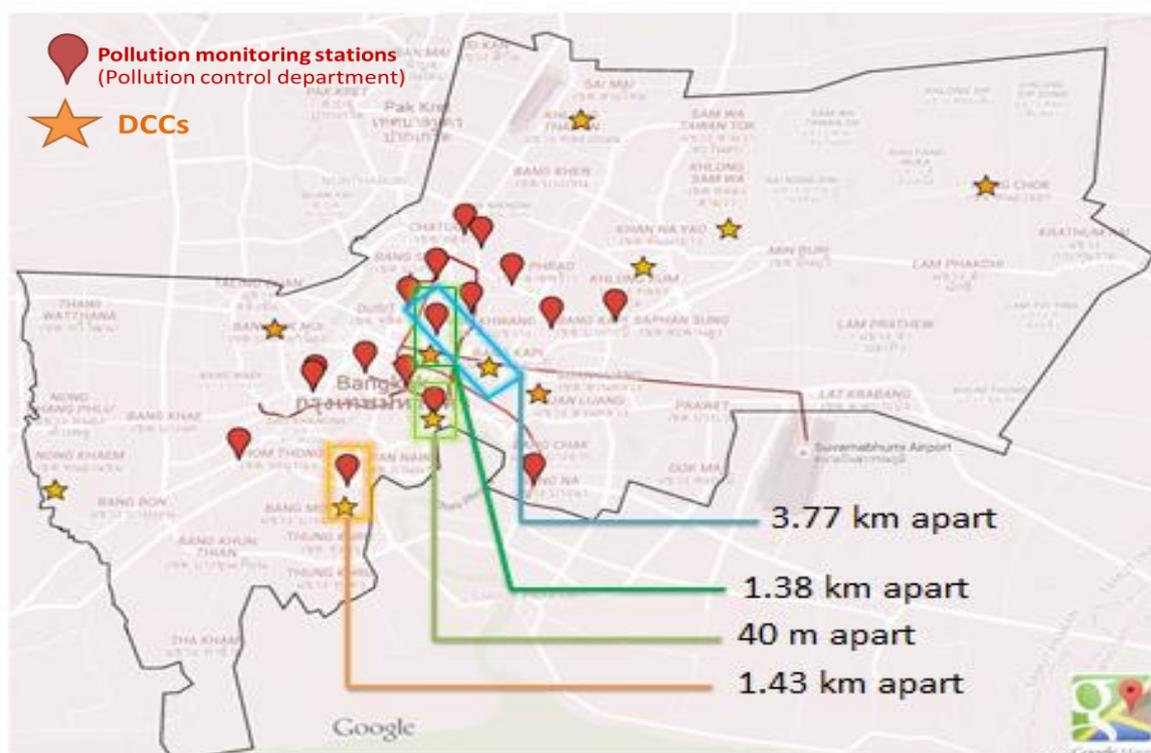
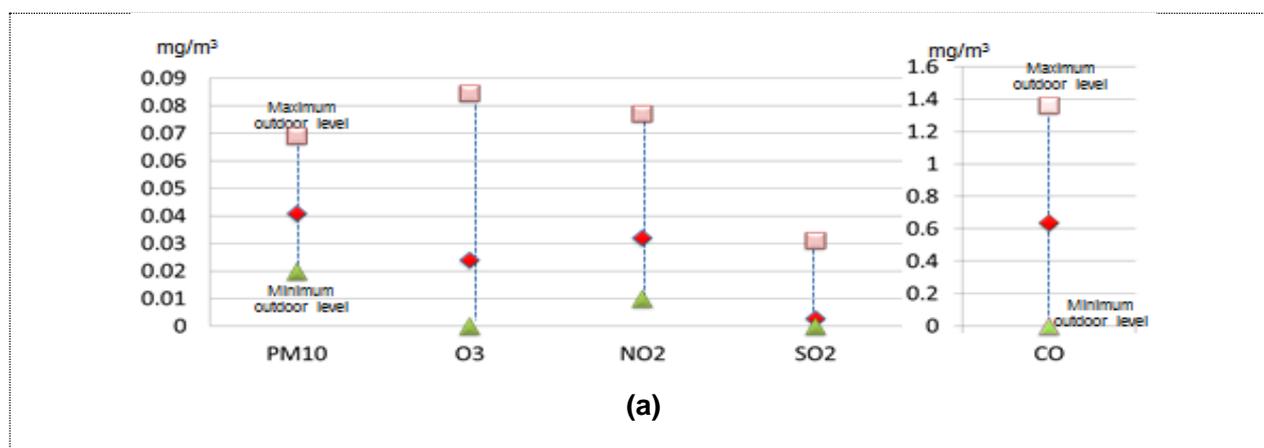
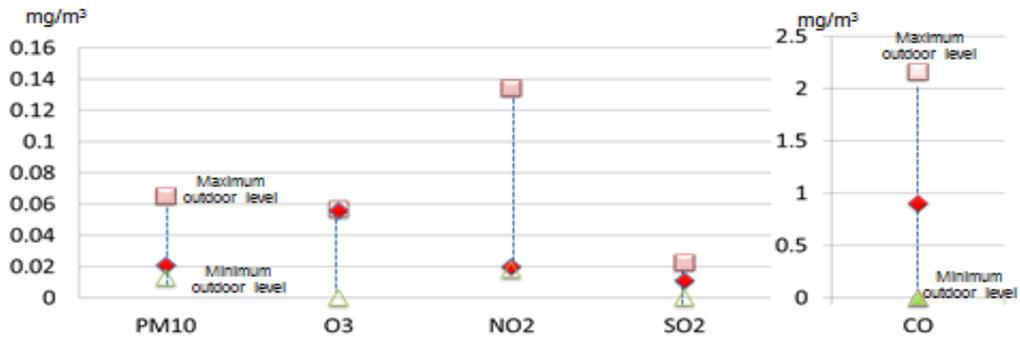


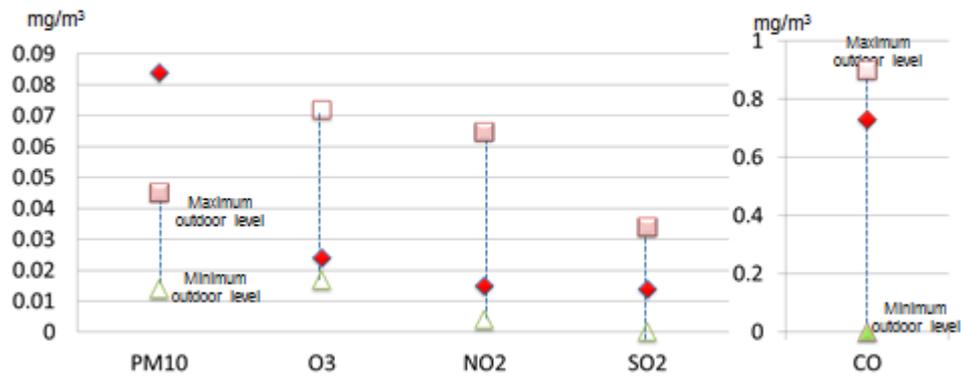
Figure 9: 4 selected DCCs and pollution monitoring stations.



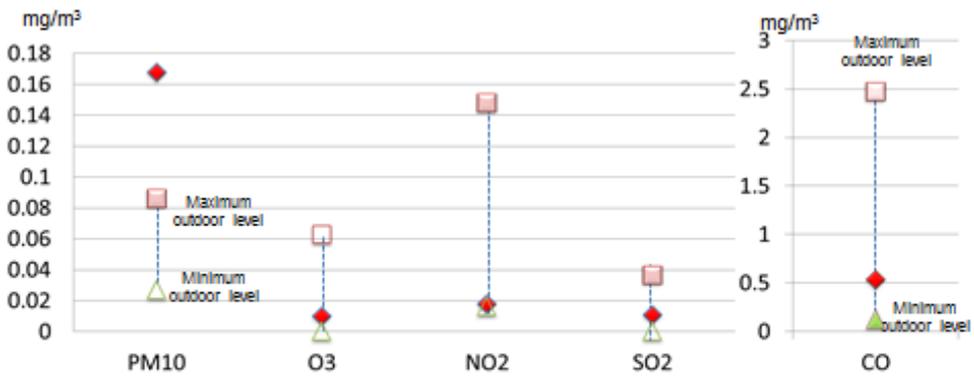
(a)



(b)



(c)



◆ Indoor pollutant level

(d)

Figure 10: Indoor pollutants levels compared to outdoor pollutants levels from nearest pollution monitoring station in 4 selected DCCs

5.3 Statistical analysis

5.3.1 Correlation between DCCs environmental characteristics and pollutant levels

The only association between DCC's environment characteristics and pollutant levels found in this study was children density and bacterial count using multiple regression analysis($\beta = 0.952$, $p < 0.001$).

5.3.2 Children baseline and home environment characteristics

From Table 10, the total number of children participated in this study is 436, where 52% are male and 48% are female. Children have mean age of 50.1 months old, and mean duration of stay in DCC of 17.3 months. Regarding health status, 9.6% of children are known with case of allergic rhinitis while 7.3% are known with case of asthma. Only 25.9% of children receive influenza vaccination during the past year. 24.08% of children have main care taker who are not parents. Parent education level is mainly secondary school, and majority of the family have income of about 10,000 - 30,000 Baht per month.

About home environment, it was found that on average, each household consists of 5 people. Mean distance between home and DCC is 2.81 km, and main transportation to DCC is motorcycle. Reported indoor smoking is as high as 51.4% of the households, where 14.4% have more than one indoor smoker and 43.3% of the fathers smoke. Most houses use open ventilation with electric fan. Air conditioners are used in 21.1% of the houses. 23.3% of children are exposed to indoor cats and 15.6% to indoor dogs. 6.2% of the households still use solid fuel for cooking. Visible sign of dampness or mold are reported in as high as 31.4% of the houses.

Table 10: Children, family and home characteristics

Children characteristics	
Mean age	50.1 months (SD = 9.4)
Mean duration of DCC attendant	17.3 months (SD = 7.5)
Male	229 (52%)
Female	207 (48%)
Underlying diseases	
- Allergic rhinitis	42 (9.6%)
- Asthma	32 (7.3%)
- G6PD deficiency	6 (1.4%)
- Anemia	4 (0.9%)
- Hepatitis B	2 (0.5%)
- Epilepsy	1 (0.2%)
Influenza vaccination during 1 year	113 (25.9%)
Family characteristics	
Father education	
- Below primary school	4 (0.9%)
- Primary school	88 (20.2%)
- Secondary school	289 (66.3%)
- Bachelor degree	52 (11.9%)
- Above Bachelor degree	3 (0.7%)
Mother education	
- Below primary school	11 (2.5%)
- Primary school	129 (29.6%)
- Secondary school	258 (59.2%)
- Bachelor degree	37 (8.5%)
- Above Bachelor degree	1 (0.2%)
Family income	
- <10,000 Baht/month	145 (33.3%)
- 10,000-30,000 Baht/month	251 (57.6%)
- >30,000 Baht/month	40 (9.2%)
Home characteristics	
Indoor smoking	224 (51.4%)
- More than one indoor smokers	63 (14.4%)
- Father smoking	189 (43.3%)
- Mother smoking	5 (1.1%)
Cooking fuel	
- Gas	390 (89.4%)
- Electricity	76 (17.4%)
- Solid fuel	27 (6.2%)

5.3.3 Correlation between pollutant exposure and respiratory symptoms of children

From Poisson regression analysis shown in Table 11,

- ✓ Frequency of parent-report rhinitis correlated with
 - PM₁₀ (IRR 70.3, 95%CI 12.4-399.7, p-value <0.001),
 - CO (IRR 3.2, 95%CI 2.4-4.2, p-value <0.001),
 - benzene (IRR 2.3, 95%CI 1.8-3.2, p-value <0.001) and
 - D. Pteronyssinus level (IRR 2.1, 95%CI 1.7-2.7, p-value <0.001).
- ✓ Frequency of teacher-report rhinitis correlated with
 - CO (IRR = 2.5, 95%CI 1.6-3.8, p-value <0.001) and
 - benzene (IRR = 3.0, 95%CI 2.2-4.3, p-value <0.001) levels.
- ✓ Frequency of parent-report cough correlated with
 - PM10 (IRR 15.2, 95%CI 3.0-78.2, p-value=0.001),
 - CO (IRR 2.8, 95%CI 2.1-3.7, p-value<0.001),
 - benzene (IRR 1.4, 95%CI 1.1-1.9, p-value=0.02).
- ✓ Frequency of teacher-report cough correlated with
 - CO (IRR = 3.5, 95%CI 2.5-4.9, p-value <0.001),
 - benzene (IRR = 2.7, 95%CI 1.9-3.9, p-value <0.001) levels.
- ✓ Parent-report dyspnea frequency correlated with
 - dust mite level. (IRR = 3.9, 95%CI 1.7-9.1, p-value=0.001) and
- ✓ parent-report dyspnea frequency correlated with
 - benzene level (IRR = 37.6, 95%CI 9.0-158.1, p <0.001).

Table 11: Correlation between pollutants exposure and respiratory symptoms

<i>Pollutants</i>	<i>Respiratory symptoms</i>	<i>IRR</i>
PM ₁₀	Parent-report rhinitis frequency	70.26 (12.35 – 399.69)
	Parent-report cough frequency	15.20 (2.95 – 78.23)
CO	Parent-report rhinitis frequency	3.15 (2.36 - 4.21)
	Teacher-report rhinitis frequency	3.04 (1.61 – 3.84)
	Parent-report cough frequency	2.76 (2.06 – 3.69)
	Teacher-report cough frequency	3.51 (2.52 – 4.91)
High Benzene	Parent-report rhinitis frequency	2.33 (1.71-3.18)
	Teacher-report rhinitis frequency	2.49 (1.61 – 3.83)
	Parent-report cough frequency	1.41 (1.06 – 1.89)
	Teacher-report cough frequency	3.16 (2.21 – 4.54)
	Number of absent days due to respiratory symptoms	2.85 (2.29 - 3.55)
DP	Parent-report rhinitis frequency	2.14 (1.67 – 2.74)
	Parent-report dyspnea	3.13 (1.36 – 7.21)

When compared between DCCs, as shown in Table 12, that had pollutant levels higher than standard level for indoor air quality and those within standard levels, using Mann-Whitney U test, it was found that children in DCCs with high PM_{10} levels had significantly higher parent-report and teacher-report rhinitis symptoms, parent-report and teacher-report cough. Wilcoxon Rank Sum Test revealed that children in DCCs with high PM_{10} levels had higher frequency of school absence caused by respiratory symptoms and frequency of hospital visit due to respiratory symptoms. In DCCs with high airborne bacterial levels, children had higher frequency of parent-report rhinitis and cough symptoms when compared to DCCs with airborne bacterial counts within standard using Mann-Whitney U test. For DCCs with high *D. Pteronyssinus* levels, children also had significantly higher parent-report cough frequency.

Table 12: Difference in children’s respiratory symptoms frequency comparison in DCCs with high and normal pollution levels

Pollutants	Symptoms	Respiratory symptom frequency (episode/month) DCCs with high pollutant level, N(%)					Respiratory symptom frequency (episode/month) DCCs with normal pollutant level, N(%)					P-value
		0	1	2	3	4	0	1	2	3	4	
Benzene	Parent-report rhinitis symptoms	6(15.4)	10(25.6)	9(23.1)	10(25.6)	4(10.3)	689(59.6)	271(23.4)	112(9.7)	45(3.9)	4(3.4)	< 0.001
	Teacher-report rhinitis symptoms	19(48.7)	6(15.4)	7(17.9)	3(7.7)	4(10.3)	991(85.7)	91(8.0)	33(2.8)	28(2.4)	13(1.1)	< 0.001
	Parent-report cough	7(17.9)	10(25.6)	12(30.8)	4(10.3)	6(15.4)	611(52.8)	326(28.2)	119(10.3)	60(5.2)	40(3.5)	< 0.001
	Teacher-report cough	16(41.0)	10(20.6)	5(12.8)	3(7.7)	5(12.8)	853(73.8)	222(19.2)	59(5.1)	13(1.1)	9(0.8)	< 0.001
Bacterial count	Absent school caused by respiratory symptoms*	13(33.3)	26(66.7)	-	-	-	774(64.4)	412(35.6)	-	-	-	< 0.001
	Hospital visit*	19(48.7)	20(51.3)	-	-	-	961(83.1)	195(16.9)	-	-	-	< 0.001
Dust Mite (D. Pteronyssinus)	Parent-report rhinitis symptoms	520(62.3)	176(21.1)	80(9.6)	42(5.0)	16(1.9)	175(48.5)	105(29.1)	41(11.4)	13(3.6)	27(7.5)	< 0.001
	Parent-report cough	451(54.1)	232(27.8)	81(9.7)	44(5.3)	26(3.1)	167(46.3)	104(28.8)	50(13.8)	20(5.5)	20(5.5)	< 0.001
	Parent-report cough	168(45.4)	114(30.8)	49(13.2)	25(6.8)	14(3.8)	450(54.5)	222(26.9)	82(9.9)	39(4.7)	32(3.9)	< 0.001

* All of statistical analysis performed by Mann-Whitney U Test, except absent school caused by respiratory symptoms (0=present, 1=absent) and Hospital visit (0= no, 1=yes) was analyses by Wilcoxon Rank Sum Test
The statistically significant is p-value < 0.05.

5.3.4 Correlation between pollutant exposure and fractional exhaled nitric oxide

A total of 110 children in rainy season and 100 children in winter & summer for FENO measurement. The test was successfully performed on 102, 93 and 86 children in rainy season, winter, and summer respectively. The FENO measurement failed in some children due to non-cooperation.

As shown in Table 13, the majority of children have FENO level lower than 20 ppb, which implies non-eosinophilic or no airway inflammation. In this study, there were no children having FENO level above 35 ppb which implies uncontrolled or deteriorating eosinophilic airway inflammation. 9 children in rainy season, 3 children in winter, and 11 children in summer have FENO level between 20 and 35 which is in the intermediate range that need to be cautious for eosinophilic airway inflammation and need monitoring on change in clinical and FENO level. Respiratory infection can also cause elevated level of FENO. Multiple logistic regression analysis showed strong association only between benzene and FENO (OR 5.9 95%CI 1.5-22.9, p-value=0.01).

Table 13: Children numbers according to FENO level

Seasons	Children Enrolled	Successful measurement	F _E NO levels		
			Low (<20 ppb)	Intermediate (20-35 ppb)	High (>35 ppb)
Rainy	110	102	93	9 (8.8%)	0
Winter	100	93	90	3 (3.2%)	0
Summer	100	86	75	11 (12.8%)	0

5.3.5 Correlation between pollutant exposure and ADHD score

Questionnaires were completed in 363 children with the following analysis, as shown in Table 14.

- ✓ T-score of parent-report ADHD symptoms were
 - <50 (normal) in 70.8%,
 - 51-60 (low risk) in 20.1%,
 - 61-70 (moderate risk) in 8%, and
 - >70 (high risk) in 1.1% of children.
- ✓ T-score of teacher-report ADHD symptoms were
 - <50 (normal) in 77.7%,
 - 51-60 (low risk) in 20.7%,
 - 61-70 (moderate risk) in 1.4%, and

- >70 (high risk) in 0.3% of children.

In the present study, there is very modest correlation between ADHD score and PM₁₀ level. PM₁₀ levels correlated with teacher-report ADHD score with R² of 0.025, p = 0.001, and for parent-report ADHD score with R² of 0.037, p < 0.001.

Table 14: ADHD score results

Teacher-report ADHD symptoms	
Normal (T-score <50)	282 (77.7%)
At risk	
- Low risk (T-score 51-60)	75 (20.7%)
- Moderate risk (T-score 61-70)	5 (1.4%)
- High risk (T-score >71)	1 (0.3%)
Parent-report ADHD symptoms	
Normal (T-score <50)	257 (70.8%)
At risk	
- Low risk (T-score 51-60)	73 (20.1%)
- Moderate risk (T-score 61-70)	29 (8%)
- High risk (T-score >71)	4 (1.1%)

5.4 Discussion & Conclusion

Air pollution was a significant problem in Bangkok metropolitan DCCs. In all season, PM₁₀ were a major indoor air pollutant in DCCs in Bangkok. Indoor ozone levels are elevated in winter with similar trend of outdoor ozone levels. Respiratory symptoms of children were correlated with PM₁₀, CO, benzene and dust mite levels. High benzene levels correlated well with children's FENO levels, which implied airway inflammation. PM₁₀ levels had very modest correlation with children's ADHD score. It is noted that children overcrowding is a problem in Bangkok Metropolitan DCCs.

From this study, outdoor pollution including traffic was important source of indoor air pollutant, where exposure to indoor air pollutants significantly affected children respiratory symptoms.

Comparing to the previous studies in DCCs in Bangkok [9, 11], similar results were found that indoor PM₁₀ levels and airborne microbial counts were high in the majority of DCCs. Other pollutants such as SO₂, O₃, CO, NO₂, and dust mite antigens levels had never been studied in DCCs in Bangkok before hence this was the first study to provide data on indoor levels of these pollutants.

In this study, it was found that even though indoor CO was within WHO standard levels, they were significantly correlated with respiratory symptoms frequency (rhinitis and cough) in preschool children. This might suggest that the current standard levels for indoor air pollutants might not be appropriate for children and day care centers.

However, there were some limitations to this study. The total number of DCCs included in this study was only 11 DCCs. Therefore, the power to determine the correlation between DCC environmental characteristics and pollutant levels could be inadequate. One DCC was closed during the study period after the first air pollution level measurement was attempted. The indoor air pollutants levels in this DCC were within standard levels; therefore, this could be assumed no significant affect to the study results. Also, indoor pollutant levels cannot be continuously monitored in this study since a single-day measurement might not represent the actual average exposure to indoor pollutants in a season. Respiratory symptoms frequency assessment by parent and teacher-report questionnaires also had limitation due to possible recall bias.

The finding of correlation between children density and indoor airborne bacterial count was similar to previous studies that found correlation between occupancy rate with bacterial and CO₂ levels [17, 18, 19]. Although In this study did not find any correlation between ventilation system and pollutant levels, there were some other studies [20] that found air-conditioned and mixed ventilation to be correlated with higher level of pollutants with indoor sources when compared to naturally ventilated DCC. In this study, no significant correlation between visible signs of dampness or mold with level of airborne fungi was found; whereas, positive correlation was reported in other study [18].

Apart from the main objectives of this study, it was found that a number of DCCs in Bangkok still have overcrowding problem. In addition, home smoking rate is rather high, and this issue should receive attention and intervention. Also, Separate kitchen should be present in DCC with cooking activity.

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