1	Efficacy of newly invented temperature and humidity control machine on house dust							
2	mite allergen and clinical symptom of dust mite sensitized allergic rhinitis children: a							
3	pilot study							
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22 Abstract

23 Background:

24 House dust mite avoidance is advised in dust mite sensitized patients to decrease their risk

of developing symptoms. Maintaining a relative humidity (RH) of less 50% is recommended

26 for reducing the growth of dust mite.

27 **Objective:**

- 28 This study was to investigate the efficacy of temperature and humidity control machine on
- the level of dust mite allergens and total nasal symptom score (TNSS) in dust mite
- 30 sensitized allergic rhinitis children.

31 **Method:**

- 32 Children (8-15 years) with dust mite sensitized persistentAR were enrolled. The temperature
- and humidity control machine was installed in the bedroom for 6 months. TNSSwas
- 34 accessed before and every month after machine installingand the level of dust mite allergen
- 35 (Der p1 and Der f1) from the mattress were measured before and every 2 months after
- 36 machine installingusing enzyme-linked immunosorbent assay (ELISA).

37 Results:

- 38 A total of 7 children were enrolled. The reduction of Der f1 was demonstrated as early as 2
- 39 months after installing the machine but the significant difference was seen at 4 months and it
- 40 was sustained low throughout 6 months (p<0.05). There was a modest but
- 41 significant reduction in TNSS at 2 and 4 months (p=0.003). Seventy percent of the patients
- 42 were able to stop using their intranasal corticosteroids. However, there was no correlation
- 43 between TNSS and the level of dust mite antigen.

44 **Conclusions:**

- The level of house dust mite in the mattress decreases after using temperature and humidity control machine. This machine mayhave a role in controlling clinical symptom of dust mite sensitized AR children.
- 48 Key words: Allergic rhinitis, Dust mite, Der p1, Der f1, children, relative humidity
- 49

51 Introduction

Allergic rhinitis is a common allergic disease worldwide. It has been proven that more than
600 million patients suffering worldwide. In Thailand, the prevalence of allergic rhinitis
increased from 37.9% in the year 1995 to 50.6% in the year 2001.¹

55 Clinical symptoms of allergic rhinitis consist of nasal itching, sneezing, watery nasal discharge, rhinorrhea, and blocked nose. Allergic rhinitis can be classified as intermittent or 56 57 persistent and the severity from mild to severe. Medication treatments include antihistamine, 58 oral leukotriene receptor antagonists, and topical nasal steroid depending on disease severity.² However, allergen avoidance is advocated as a central role for the managements 59 of allergic rhinitis.³House dust mite is the most common aeroallergen sensitization in patients 60 with respiratory allergy in tropical climate country such as Thailand and Singapore.^{4, 5} A 61 62 recent practice parameter has advised dust mite sensitized patients with asthma or rhinitis to 63 minimize exposure to dust mite allergens to decrease their risk of developing of symptoms. 64 Water balance is critical for house dust mite survival. The optimum relative humidity (RH) 65 more than 65% isneeded to maintain their water balance and growth. If humidity decreases 66 below 50%, mite proliferation decreases and survival is decreased. The temperature also 67 has an impact on RH, as a result maintaining stable temperature is required to maintain a 68 stable RH⁶. The present study was to investigate the effect of newly invented temperature 69 and humidity control machine on the level of dust mite allergen and clinical symptom of dust 70 mite sensitized allergic rhinitis children. This machine was invented by Dr.

VeeraponMonyakul,KingMongkut's University of Technology Thonburi. This machine has
 been shown to provide a precise temperature and desired humidity in the room.⁷

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76 Patients and Methods

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78 Patients

79 Children aged 8-15 year with persistent allergic rhinitis according to ARIA guideline who 80 have skin prick test positive to house dust mites (Dermatophagoidespteronyssinus 81 and/orDermatophagoides farina) and had symptom of allergic rhinitis related to the exposure 82 to house dust mite were eligible for the study. Positive skin test was defined as a wheal 83 diameter at least 5 mm larger than the control. Patients with multiple sensitization and 84 previously or current on specific immunotherapy were excluded. Since the temperature and 85 humidity control machine operate with air-conditioning and the machine was installed in the 86 bedroom, the enrolled patients' must have air-conditioning in the bedroom.

87 Study protocol

88 This study is a 6 month experimental study with 1 month run in period. During running, 89 intranasal corticosteroids were not allowed, but the patients were allowed to use 90 decongestant and antihistamine and nasal irrigation. At V0, the patients were evaluated for 91 total nasal symptom scores(TNSS) and dust mite collection from the mattress in the 92 bedroom for dust mite antigen measurement. Then, clinical visit and TNSS was assessed 93 every month after machine installation in the bedroom. The dust mite antigens(Der p1 and 94 Der f1) were measured every 2 months (Figure 1). TNSS was defined as the sum of the 95 average score in the past 4 weeks for the 4 individual patients-reported nasal symptoms of 96 congestion, itching, rhinorrhea, and sneezing (0-3 from mild to severe). After enrollment, patients were treated their rhinitis according to ARIA guideline³ and received standard 97 98 recommendation for dust mite control. The study was reviewed and approved by the human 99 rights and ethic committee of Faculty of medicine, Ramathibodi Hospital, Mahidol 100 University.All enrolled patients and guardian were explained and consented for the study.





112 Figure 1: Study protocol

113 The temperature and humidity control machine

114 The temperature and humidity control machine invented by Dr. VeeraponMonyakul, King Mongkut's University of Technology Thonburi, Bangkok, Thailand.⁷The temperature and 115 116 humidity control machine consists of a unit of humidifier and a unit of dehumidifier within the 117 same case. The device has the dimension of 0.3 x 1.0 x 1.2 meters (Width x Length x 118 Height)and is similar to a 10,000 BTU air condition (type TURBO A.P.S., Saijo Denki 119 International Co., MuangNonthaburi, Thailand). This device does operate with air-120 conditioningunder the automatic control by microprocessor in order to keep the relative 121 humidity and temperature constant at all times, 55% RH and 25 degrees Celsius. The device 122 is installed and running incorporated with air-conditioning under the automatic control by 123 microprocessor (Figure 2). Therefore, the temperature and humidity in testing room was

- 124 controlled to be as 25°C and 55% RH. This device has never been used as a commercial
- 125 product.



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Figure 2: The newly invented temperature and humidity control machine (A). Diagram
 demonstrating the operation system of the machine with the air condition(B).

129 Mite Allergen Measurement

130 Levels of Der p 1, the major allergen of Dermatophagoidespteronyssinus mites and Der f1,

131 the major allergen of Dermatophagoides were measured using the commercial sandwiched

132 ELISA reagents (Indoor Biotechnology, U.K.) at Siriraj Dust Mite Center for Services and

133 Research Department of Parasitology, Faculty of Medicine Siriraj Hospital Mahidol

134 University, Bangkok, Thailand. The lower detection limit is 0.2 microgram of Der f1 and Der

135 p1 per gram dust.

136 Statistical analysis

Descriptive analysis was used to report the mean value and standard deviation (sd) of the data. Comparative analysis between pre and post machine installation was analyzed using paired Student's t test or ANOVA repeated tests. Correlation between TNSS and the level of dust mite antigen was analyzed using Pearson correlation. The differences with a p value

- 141 less than 0.05 were considered statistically significant. Data were analyzed using SigmaPlot142 12.
- 143

144 Results

- 145 A total of 7 children with house dust mite sensitized allergic rhinitis were enrolled. Four
- 146 children were male, mean age was 9.8 years, mean age of onset was 4 years and mean
- 147 duration of symptom was 4 years. Three children also had asthma. Baseline characteristic of
- 148 the patients was shown in table 1.
- 149 Table 1: Baseline characteristic of participating children

	Age	Sex	Duration	Age of	Other	Severity	Wheal	Wheal	Baseline
	(yr)		of AR	onset	allergic	of AR	size for	size for Df	TNSS
			(yr)	(yr)	diseases		DpbySPT	By SPT	
1	12	М	1	7	Drug	Mild	6*5	8*6 with	3.45
					allergy	persistent		pseudopod	
2	12	F	7	4	AD,food	Mild	9*7	12*6 with	1
					allergy	persistent		pseudopod	
3	8	М	2	6	No	Moderate	10*5 with	10*5 with	6
						persistent	pseudopod	pseudopod	
4	8	М	6	2	Asthma	Mild	7*7	10*6	1
						persistent			
5	11	F	4	7	Asthma	Moderate	10*6	16*15	7.6
						persistent			
6	10	М	1	8	No	Mild	10*25	10*8	3
						persistent			
7	8	F	7	1	Asthma	Mild	17*15	23*13	1.83
						persistent			

150 AR : Allergic Rhinitis, SPT : Skin Prick Test, TNSS: Total Nasal Symptom Scores

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153 Total nasal symptom score after installing the machine

Total nasal symptom score was significantly decreased after 2 months of installation the machine. However due to the variation in the total nasal symptom score and the small sample size, the statistically significant difference was seen only at 2 months and 4 months after installation (Figure 1). Five out of seven patients were able to stop using their intranasal corticosteroids.

159 The level of Der p1 and Der f1 after installation

There was are a reduction of the level of Der f1 as early as 2 months after installing the machine but the significant difference was demonstrated at 4 months and it was sustained low throughout 6 months (p<0.05) (Figure 2). However, the level of Der p1 in the mattress was lower than 0.2 ug/g dust in 6/7 houses resulting in the non-significant changes in the level of Der p1 (Figure 3).

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166 The level of dust mite antigen and TNSS

167 There was no correlation between the TTNS and the level of Der f1 or Der p1 (Figure 4 168).Comparison between patient s who had baseline TNSS more than 3 and less than 3, there 169 was no significant difference between the baseline level of Der f1 level (1.4 vs 2.5 ug/ gram 170 dust, p = 0.4)

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Figure 1 Clinical symptom score in each visit after machine installation. V0=before machine
installation, V1,2,3,4,5, and 6 represent 1,2,3,4,5 and 6 months after installation respectively.
Same symbol represents data from same subject.



Figure 2 The level of house dust mite antigen Der f1 (A) and Der p1 (B) in each visit after machine installation. V0=before machine installation, V2,4,6 represent 2,4 and 6 months after installation respectively. Samesymbol represents data from same subject.



Figure 3: The level of house dust mite antigen Der f1 (A) and Der p1 (B) and total nasalsymptom scores.

187 **Discussion**

188 House dust mite is the major allergen causing symptoms in respiratory allergy especially in tropical countries especially Thailand.⁵More than 70% of children with respiratory allergy 189 were sensitized to house dust mites.⁸Current guidelines for allergic rhinitis and asthma 190 191 management have emphasized on allergen avoidance and exposure control as a 192 fundamental treatment in all step of treatment. Environmental exposure of dust mite can be 193 controlled by removal of facilitative factor for mite growth and survival, clean or remove mite reservoirs and get rid of the mites.⁶The most important facilitative factor for mite survival is 194 relative humidity (RH) in the ambient environment.⁹Maintaining RH in the house to less than 195 196 55% has been recommended to control dust mites and their allergen in homes in temperate climate¹⁰ If there is a changing in temperature that results in increasing the RH only few 197 hours can promote the growth of dust mites.⁶ 198

199 The present study has demonstrated our newly invented temperature and humidity control 200 machine was able to decrease the level of dust mite allergen in the bedroom of allergic 201 rhinitis patients from Thailand, the high humidity country. Previous studies have shown the ineffectiveness of portable dehumidifiers in reduction the level of dust mite antigens.^{11,} 202 ¹²Since RH is temperature dependent. The advantage of this newlyl invented machine is the 203 204 ability to maintain stable RH in conjunction with stable temperature by running incorporated with air-conditioning under the automatic control using microprocessor.⁷ Previous study has 205 206 been demonstrated that maintaining daily indoor relative humidity below 50% RH but 207 allowing RH above 50% for more than 2 hours daily resulted in dust mite's population growth.¹³ As a result, a portable dehumidifier in home use may not be able to control the 208 209 dust mite population due to the inability to maintain stable temperature and RH throughout 210 24 hours.

After installing this novel machine, there was a modest but significantly improvement in total nasal symptom score as early as 2 months. However, there was no correlation with the level 213 dust mite antigen. Previous studies have recommended decreasing the level of Der p 1 and 214 Der f 1 to lower than 2 microgram per gram dust in order to lower the risk of dust mite sensitization¹⁴. Exposure to 10 microgram of Der p 1 and Der f 1 per gram dust has been 215 216 suggested as an exposure threshold for the development of asthma symptoms in already sensitized children¹⁵. In the present study, the levels of Der p 1 and Der f 1 from the 217 218 bedrooms of the participating patients before machine installation were varied from 0.2-4 219 microgram per gram dust. Three out of seven subjects exposed to dust mite antigen lower 220 than 2 microgram per gram dust. Consequently, the lower level of initial dust mite antigen may result in poorer correlation with the symptom improvements. In addition, forty percent 221 222 of the participating patients had mild but persistent symptoms. The efficacy of the machine in 223 reduction TTNS may be stronger if there were more severe patients enrolled. Since, the 224 objective measurements such as rhinomanometry or nasal peak flow were not evaluated in 225 the present study. The improvement in these objective measurements may be demonstrated 226 early than the significant changes of clinical symptoms. There was a significant reduction of 227 the level of dust mite antigen after machine installation. As a result, the significant 228 improvement of clinical symptom may be demonstrated if the machine has been used for the 229 longer duration.

230

231 The limitation of the present study is this novel machine is not commercially available and under developed. There are limited numbers of the machine which resulted in little number 232 233 of enrolled patients. In addition due to the study period is 6 months, this may result in the 234 difference in the season among patients. Nevertheless, this machine was demonstrated to be able to control the stable temperature and humidity,⁷ the difference in the season should 235 236 have minimal effect on the RH. There is no control patients in the present study, the placebo 237 effect cannot be excluded for the improvement of the clinical symptoms. In addition, other 238 methods that can control or decrease the level of dust mite were not monitor in the present

- 239 study.Further study on the efficacy of the machine in clinical practice needs more
- 240 participating patients, longer duration of study and non-intervention group.

241 In conclusion, this pilot study has demonstrated that the newly invented temperature and

- 242 humidity control machine was able to decrease the level of dust mite antigen in the bedroom
- 243 of dust mite sensitized allergic rhinitis children. There was also a modest improvement in
- 244 clinical symptoms of allergic rhinitis. Larger and well control studies are required to confirm
- the efficacy of this machine.
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- Bunnag, C., P. Jareoncharsri, P. Tantilipikorn, P. Vichyanond, and R. Pawankar, *Epidemiology and current status of allergic rhinitis and asthma in Thailand -- ARIA Asia-Pacific Workshop report.* Asian Pac J Allergy Immunol, 2009. 27: 79-86.
- Brozek, J.L., J. Bousquet, C.E. Baena-Cagnani, S. Bonini, G.W. Canonica, T.B. Casale, et al.,
 Allergic Rhinitis and its Impact on Asthma (ARIA) guidelines: 2010 revision. J Allergy Clin
 Immunol, 2010. 126: 466-76.
- Bousquet, J., J. Reid, C. van Weel, C. Baena Cagnani, G.W. Canonica, P. Demoly, et al.,
 Allergic rhinitis management pocket reference 2008. Allergy, 2008. 63: 990-6.
- Andiappan, A.K., K.J. Puan, B. Lee, A. Nardin, M. Poidinger, J. Connolly, et al., *Allergic airway diseases in a tropical urban environment are driven by dominant mono-specific sensitization against house dust mites.* Allergy, 2014. 69: 501-9.
- 2585.Sritipsukho, P., Aeroallergen sensitivity among Thai children with allergic respiratory259diseases: a hospital-based study. Asian Pac J Allergy Immunol, 2004. 22: 91-5.
- Portnoy, J., J.D. Miller, P.B. Williams, G.L. Chew, J.D. Miller, F. Zaitoun, et al., *Environmental assessment and exposure control of dust mites: a practice parameter.* Ann Allergy Asthma
 Immunol, 2013. 111: 465-507.
- Sookahaiya T, M.V., Thepa S., Assessment of the thermal environment effect a human comfort and health for the development of novel air conditioning system in tropical regions. Energy Build, 2010. 42: 1692-1702.
- Anantasit N., K.W., Direkwattanachai C., Manuyakorn W., Vilaiyuk S., Benjaponpitak S.,
 Aeroallergen Sensitization Among Children with Respiratory
- Allergy in Ramathibodi Hospital, Bangkok. J Allergy Clin Immunol, 2010. 125: AB17.
- Arlian, L.G., *Water balance and humidity requirements of house dust mites.* Exp Appl Acarol, 1992. 16: 15-35.
- Arlian, L.G., J.S. Neal, M.S. Morgan, D.L. Vyszenski-Moher, C.M. Rapp, and A.K. Alexander,
 Reducing relative humidity is a practical way to control dust mites and their allergens in homes in temperate climates. J Allergy Clin Immunol, 2001. 107: 99-104.
- Custovic, A., S.C. Taggart, J.H. Kennaugh, and A. Woodcock, *Portable dehumidifiers in the control of house dust mites and mite allergens.* Clin Exp Allergy, 1995. 25: 312-6.
- Hyndman, S.J., L.M. Vickers, T. Htut, J.W. Maunder, A. Peock, and T.W. Higenbottam, A
 randomized trial of dehumidification in the control of house dust mite. Clin Exp Allergy, 2000.
 30: 1172-80.

- Arlian, L.G., J.S. Neal, and D.L. Vyszenski-Moher, *Reducing relative humidity to control the house dust mite Dermatophagoides farinae.* J Allergy Clin Immunol, 1999. 104: 852-6.
- 14. Kuehr, J., T. Frischer, R. Meinert, R. Barth, J. Forster, S. Schraub, et al., *Mite allergen exposure is a risk for the incidence of specific sensitization.* J Allergy Clin Immunol, 1994. 94:
 44-52.
- 15. Custovic, A. and M. Chapman, *Risk levels for mite allergens. Are they meaningful?* Allergy,
 1998. 53: 71-6.

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