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International Journal of Biometeorology The impact of heat waves on children's health: A systematic review --Manuscript Draft--

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Abstract:	Young children are thought to be particularly sensitive to heat waves, but relatively less research attention has been paid into this field to date. A systematic review was conducted to elucidate the relationship between heat waves and children's health. Literature published up to August 2012 were identified using the following MeSH terms and keywords: "heatwave", "heat wave", "child health", "morbidity", "hospital admission", "emergency department visit", "family practice", "primary health care", "death" and "mortality". 628 publications were identified and 12 met selection criteria. Existing literature does not consistently suggest that mortality among children increase significantly during heat waves, even though infants were associated with more heat-related deaths. Exposure to heat waves in the perinatal period may pose a threat to children's health. Pediatric diseases or conditions associated with heat waves include renal disease, respiratory disease, electrolyte imbalance and fever. Future research should focus on how to develop a consistent definition of a heat wave from a children's health perspective, identifying the best measure of children's birth outcomes, and understanding the possible impacts of heat waves on children's birth outcomes, and understanding the differences in vulnerability to heat waves among children of different ages and from different income countries. Projection of children's disease burden caused by heat waves under climate change scenarios, and development of effective health measures, are also strongly recommended.					

9/12/2012

Dear Professor Sheridan,

Re: "The impact of heat waves on children's health: A systematic review"

(Ms. No. IJBM-D-12-00185)

Thank you very much for your letter and attached reviewers' comments on our manuscript. We have revised the manuscript according to your advice and the reviewers' comments. The changes have been marked red.

In response to Reviewer 1:

Reviewers' comments:

- give a clearer definition of the classes children, infants... at the beginning of the introduction, and not at the end

Response: A clearer definition of children has been shifted to the beginning of the second paragraph as suggested (Page 3, Lines 60-62).

- use these classes to organize the paragraph on the sensitivity to heat waves in the introduction, the results, and the discussion. This is already done but not systematically, especially in the discussion.

Response: We have used the same classes to organize the paragraph on the sensitivity to heat waves across the whole manuscript.

- in the Discussion, mention if different mortality impacts occur for adults. Indeed, some heat waves don't have an impact even on adults, so it would be nice to know if the papers you quote have found no impact in children but large impacts in adults, or no impacts at all. Response: The comparison between the impacts of heat waves on mortality among adults and children, and the possible reasons for the distinction, have been added to the Discussion of the revised manuscript (Page 10, Lines 234-241). We found that very young children, especially those aged under one year old, rather than old children, are more vulnerable to the impact of heat waves when compared with adults.

- when possible, can you also give the mean daily number of deaths reported in each study, as it is likely to be very low, and therefore raises statistical issues.

Response: Nearly all of the studies looking at the impact of heat waves on children's mortality did not give the mean daily number of deaths, except for the study in Great Britain. The Great Britain study found that the number of deaths among children was lower than adults during heat waves, but they used excess death number rather than relative risk to evaluate the possible risk posed by heat waves, so there may be no statistical issue in this study.

- in the methods, the key words heat wave seem restricted, you may also try extreme temperature, or heat; to see if yields different search results.

Response: The major objective of this review was to look at the impact of heat waves on children's health. We used extreme temperature and heat as the search terms, and found some papers documenting the impact of hot and cold temperatures on children's health, which is beyond the scope of this study.

- your search mostly include studies from developed countries, but the drivers of children mortality are very different in developing countries, with a high burden paid to infectious diseases and dehydration. This should be discussed in more detail in your Discussion. In developed countries you may want to mention reports from ancient heat waves, if available, as the impacts of heat on children might have been larger at the beginning of the 20th century, when infectious diseases were a major cause of children mortality. For instance, the 1911 heat wave in France was associated to 40 000 deaths, including 29 000 young children. It resulted in a large campaign to reduce children mortality (<u>http://www.cairn.info/resume.php?ID_ARTICLE=ADH_120_0105</u>). The same was observed in UK and US, so there may be references in English.

This would be interesting, because it would show that the lower impacts observed on children now are also explained by improvements in care.

Response: We've tried very hard to find the published papers regarding the impact of ancient heat waves on children's mortality, but have not found any paper which meets our paper inclusion criteria. We've discussed more about the situations of mortality drivers in developed and developing countries. Further, we used the 1911 heat wave in France as an example to elucidate the high number of deaths among children associated with ancient heat waves and discussed the effect of improved care in reducing the heat-related deaths (Page 11, Lines 250-258).

Minor comments:

In the abstract, you may not need to develop all the bases you have used, this could save space to give more results.

Response: In the revised Abstract, we've shortened the descriptions about the bases and added more results (Page 2, Lines 31-32).

Line 47, global temperatures have risen at a faster rate than what? This sentence is too general; you could be more precise about the increases in global temperature.

Response: We've removed the previous sentence: "global temperatures have risen at a faster rate" and added a new sentence to the Introduction (Page 3, Line 47).

Line 313, the problem of heat waves is not only that heat stroke is a fatal diseases, but also that many causes of mortality may increase.

Response: We agree that the major problem of heat waves is because many causes of mortality may increase during heat wave episodes, and we've added this in the revised manuscript (Page 14, Lines 327-330). In this paragraph, we intended to highlight that the mortality rate of some diseases is very high even when treated, so we remain the example of heat stroke in the revision.

Line 220, what do you mean by "environmental infrastructures"?

Response: "Environmental infrastructures" refers to the infrastructures which can help people to cool down or rehydrate during heat waves, such as water fountain for public use etc..

Line 251, aren't most heat waves occurring during summer holidays, with low impact on missed school days?

Response: Firstly, in some countries, heat waves do not always occur during summer holidays. Secondly, paediatric diseases usually last quite long. Even though heat waves sometimes happen during holidays, heat-related diseases may still result in missed school days.

In response to Reviewer 2:

Reviewer #2: General comments:

- I like your point on page 11 that these outcomes represent only the most extreme outcomes, while other important outcomes (e.g., missed school days) are also important for further investigation.

Specific comments:

- The authors use the term 'heat wave,' but included 'heatwave' and 'heat-wave' only in literature search terms. Maybe a typo?

Response: We included both "heat wave" and "heatwave" in literature search terms because "heatwave" were written as "heat wave" in some published papers. "Heat-wave" has been revised to "heat wave" in the revision (Page 2, Line 27; Page 4, Line93).

- The authors refer to 'English articles' -- perhaps more clear as 'English-language articles.'

Response: Done (Page 4, Line 91).

- Typo in Table 1 reporting of Knowlton et al (2009) RR point estimate reads 1.09, should be 1.19.

Response: It has been revised in Table 1. Thanks.

Yours sincerely,

Zhiwei Xu (On behalf of all co-authors)

1 The impact of heat waves on children's health: A systematic review

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22 Abstract

Young children are thought to be particularly sensitive to heat waves, but relatively less 23 research attention has been paid into this field to date. A systematic review was conducted to 24 25 elucidate the relationship between heat waves and children's health. Literature published up to August 2012 were identified using the following MeSH terms and keywords: "heatwave", 26 "heat wave", "child health", "morbidity", "hospital admission", "emergency department 27 visit", "family practice", "primary health care", "death" and "mortality". 628 publications 28 were identified and 12 met selection criteria. Existing literature does not consistently suggest 29 30 that mortality among children increase significantly during heat waves, even though infants were associated with more heat-related deaths. Exposure to heat waves in the perinatal period 31 may pose a threat to children's health. Pediatric diseases or conditions associated with heat 32 33 waves include renal disease, respiratory disease, electrolyte imbalance and fever. Future research should focus on how to develop a consistent definition of a heat wave from a 34 35 children's health perspective, identifying the best measure of children's exposure to heat waves, exploring sensitive outcome measures to quantify the impact of heat waves on 36 children, evaluating the possible impacts of heat waves on children's birth outcomes, and 37 38 understanding the differences in vulnerability to heat waves among children of different ages and from different income countries. Projection of children's disease burden caused by heat 39 waves under climate change scenarios, and development of effective heat wave mitigation 40 41 and adaptation strategies that incorporate other child protective health measures, are also strongly recommended. 42

43 **Keywords:** Heat wave; Child health; Mortality; Morbidity

44

46 Introduction

There is a widespread consensus that climate is rapidly changing. Earth's average surface 47 temperature will increase by 1.8 to 4.0°C by the end of this century relative to the 1961-1990 48 level (IPCC 2007). Heat waves, sporadic periods of elevated temperatures outside the normal 49 50 range of climate variability for a specific region, occur throughout the world and are projected to become more frequent and intense in the future (Meehl and Tebaldi 2004). The 51 increasing urbanization globally compounds the potential risk due to the urban heat island 52 53 effect which can increase urban core temperatures disproportionately (Balogun et al. 2010; O'Neill and Ebi 2009). Heat waves are a significant threat to population health. For example, 54 the 2003 heat wave caused nearly 15,000 excess deaths during the period of 1-20 August in 55 France alone (Poumadère et al. 2005). To protect the population from the adverse impact of 56 heat waves, identifying who is most vulnerable for heat-related illness and death and how to 57 reduce their exposure is imperative. 58

59

Children are usually defined as human under 18 years of age (American Academy of 60 Pediatrics Committee on Environmental Health 2003), and infants refer to those who are 61 62 under one year of age. In a number of ways, children differ from adults thereby potentially increasing their sensitivity to heat waves: (i) Physiological modality: Children have less 63 64 developed thermoregulatory systems and a greater body surface area-to-mass ratio compared to adults, allowing greater heat and cold transfer between the environment and the body 65 66 (Blum et al. 1998). (ii) Metabolic modality: children have a higher metabolic rate that may render them more sensitive to heat waves (Bunyavanich et al. 2003). (iii) Cardiovascular 67 modality: children at a given activity level have a lower cardiac output than adults (Turley 68 and Wilmore 1997). Besides, children, especially infants, have lower cardiac index than 69 70 adults, which may result in differing physiologic adaptive capacity to heat waves; (iv)

71 Behavior modality: at some developmental stages, children spend more time outdoors and 72 participate in more vigorous activities than adults, which can result in more exposure to outdoor heat (United States Environmental Protection Agency 2011). (v) Self-care ability 73 74 modality: children, especially infants, and children less than two years, cannot take care of 75 themselves, and they are dependent on others to protect them from unsafe environments (Danks et al. 1962). (vi) Life expectancy modality: more expected future years of life 76 77 provides both an greater potential exposure period and also a longer period to experience delayed adverse health impacts from extreme heat exposure (Landrigan et al. 1999; Perera 78 79 2008).

80

There has been increasing interest in assessing the impact of heat waves on children's health (Basagaña et al. 2011; Knowlton et al. 2008). Nevertheless, to our knowledge, no literature reviews are available on the specific relationship between heat waves and children's health to date. A systematic review was conducted to explore whether children are more likely to be associated with heat-related deaths and elucidate some key pediatric diseases associated with heat waves.

87

88 Methods

Literature regarding heat wave and children's health published up to August 1st 2012 was
retrieved using the databases Pubmed, ProQuest, ScienceDirect, Scopus and Web of Science.
Peer-reviewed English-language journal articles were included in the initial search. The
primary search used the following U.S. National Library of Medicine's Medical Subject
Headings (MeSH terms) and keywords: "heatwave", "heat wave", "child health",
"morbidity", "hospital admission", "emergency department visit", "family practice",
"primary health care", "death" and "mortality". References and citations of the relevant

96 articles were manually inspected to make sure that all relevant articles are included.

97 Eligibility included any studies which used original data and appropriate effect estimates

98 (e.g., regression coefficient, relative risk, odds ratio, percentage change in morbidity, and

99 morbidity or excess morbidity following heat waves); where heat wave was a main exposure

100 of interest, and where children's morbidity or mortality were analyzed.

101

102 **Results**

103 628 papers were identified in the initial search. Finally, 12 studies were included (Figure 1).

104 The characteristics of the 12 studies meeting all inclusion criteria are summarized in Table 1.

105

106 The impact of heat waves on children's mortality

Nine studies examined the impact of heat waves on children's mortality. In Catalonia, Spain,
Basagaña et al. assessed the impact of heat waves on total and cause-specific mortality in
infants during the warm season from 1983 through 2006 (Basagaña et al. 2011). They found
that the effect of heat waves in infants was observed on the same day (Lag 0) of exposure and
was detected for conditions originating in the perinatal period (Relative risk (RR):1.53; 95%
CI: 1.16–2.02). The major condition originating in the perinatal period associated with heat
waves was digestive system diseases (RR: 3.85; 95% CI: 1.02–14.5).

114

In Shanghai, Huang and colleagues quantified the effect of 2003 heat wave on mortality
(Huang et al. 2010). This is one of the few English studies looking at the impact of heat
waves on mortality in China. They controlled for air pollution and did not find significant
increase in total mortality among children aged 0–4 years old (RR: 0.67; 95% CI: 0.24–1.87)
in heat waves.

Fouillet et al. investigated the effect of 2003 heat waves on mortality in France (Fouillet et al. 2006). The observed mortality during 2003 heat waves was compared to those expected on the basis of the mortality rates observed from 2000 to 2002. They did not find the significant increase in mortality among infants (observed mortality/expected mortality: 1.1; 95% CI: 0.9–1.3) or children aged 1–14 years (observed mortality/expected mortality: 1.0; 95% CI: 0.8–1.3).

127

Kysely et al. examined the effect of heat waves on daily mortality during 1991–2005 in South
Korea (Kysely and Kim 2009). They computed the excess mortality based on calculating
deviations of the observed number of deaths and the expected number of deaths for each day
of the examined period, and found that during the heat wave in 1994, the relative increase in
mortality was larger in children aged 0–14 years (+27.5%; 183 excess deaths, 95% CI: 133–
234) than that in any other age group.

134

Hutter and colleagues investigated the effect of heat waves on daily mortality during 1998– 2004 in Vienna, Austria (Hutter et al. 2007), and found that the point estimate for the relative risk of deaths during heat wave days was the highest in infants, even though the confidence interval was broad because of the low number of deaths in infants and the effect was therefore not significant (RR: 1.25; 95% CI: 0.82–1.90). Sex-specific analysis revealed that male infants had increased risk compared to female babies. However, the reason for such difference remains unclear.

142

143 Nitschke et al. investigated morbidity and mortality associated with heat waves from 1993 to
144 2006 in Adelaide using ambulance transport, hospital admission, and mortality data (Nitschke
145 et al. 2007). They classified the total population into five age groups: 0–4, 5–14, 15–64, 65–

146 74, and \geq 75 years. They found that mortality increased (even though not significant) among

147 children aged 0-4 years (RR:1.19; 95% CI:0.82-1.71) and 5-14 years (RR:1.15; 95%

148 CI:0.58–2.29) but decreased in other three age groups during heat waves. They also explored

the impact of 2008 and 2009 Adelaide heat waves on hospital admissions, ambulance call

150 out, emergency department presentations and mortality, and found that there was a significant

151 mortality rise among children aged 0–4 years (RR: 3.23 ; 95% CI:1.30–7.99) during 2008

heat waves (Nitschke et al. 2011).

153

Rooney et al. investigated the effect of 1995 heat waves on mortality in England, Wales and Greater London (Rooney et al. 1998). They analyzed the mortality variation in daily mortality and found that deaths among children aged 0–15 years increased during heat waves in England and Wales (percent change in deaths: 4.6%), and Greater London (percent change in deaths: 13.0%). In this study, air pollution was not controlled when assessing the effect of heat waves.

160

Son et al. examined mortality from heat waves in seven major South Korean cities from 2000
to 2007 (Son et al. 2012). They also investigated effect modification by individual
characteristics and heat wave characteristics (intensity, duration, and timing in season). They
found no significant percent change in mortality among children aged 0–14 years during heat
waves in Seoul.

166

167 The impact of heat waves on children's morbidity

168 Five studies have examined the relationship between heat waves and children's morbidity

- 169 (Knowlton et al. 2008; Kovats et al. 2004; Leonardi et al. 2006; Nitschke et al. 2007;
- 170 Nitschke et al. 2011). Kovats and colleagues examined the effect of heat waves on emergency

hospital admissions during April 1994 – March 2000 in London, using a time-series design
(Kovats et al. 2004). After adjusting for long term trend, season, day of week, public
holidays, the Christmas period, influenza, relative humidity, air pollution, and overdispersion,
they found no relation between total emergency hospital admissions and heat waves but they
did observe heat-related increases in emergency hospital admissions for respiratory and renal
disease in children under five years of age.

177

Leonardi et al. investigated the relationship between heat waves and calls to National Health 178 179 Service Direct – a nurse-led helpline which provides health-related information and advice and directs callers to the appropriate health service and self care, during December 2001-180 May 2004 in England (Leonardi et al. 2006). They used a time-series design and mainly 181 182 focused on calls for fever, vomiting, difficulty breathing and heat stroke and sunstroke. Potential confounders such as ozone, PM₁₀ and seasonally varying factors were controlled in 183 the data analysis. They found that total calls were moderately increased as environmental 184 temperature increased, and a rise in fever calls (RR: 2.5% per 10 °C increase in mean 185 temperature; 95 % CI: 1.8%–3.3%) was seen only for children 0–4 years in Greater London 186 and South East regions. 187

188

Knowlton et al. investigated the effect of heat waves on hospital admissions and emergency
department visits during July – August 2006 in 58 counties of California, USA, using a
descriptive design (Knowlton et al. 2008). They found that emergency department visits for
all ages were increased but the effect was greatest in the 0–4 year age group (RR: 1.05; 95%
CI: 1.04–1.07) with emergency department visits for heat-related (RR: 6.17; 95%CI: 2.58–
17.88) and electrolyte imbalance diagnoses (RR: 1.19; 95% CI: 1.10–1.30) specifically being
elevated among 0–4 year age group during the heat wave period.

196 Nitschke et al. quantified the impact of heat waves from 1993 to 2006 on morbidity and mortality in Adelaide using ambulance, hospital admission, and mortality data (Nitschke et 197 al. 2007). They found that the hospital admissions for respiratory diseases decreased during 198 199 heat waves in children aged 0-4 years (RR: 0.86; 95% CI: 0.76-0.97). They also assessed the associations between heat waves and hospital admissions, ambulance call out, emergency 200 department presentations and mortality from 2008 to 2009 in Adelaide, Australia (Nitschke et 201 202 al. 2011), and found that during the heat waves, there was a significant rise of renal hospital admissions in 5–14 year age group (RR: 2.64; 95% CI: 1.47–4.73). Besides, significant rises 203 204 of renal emergency department presentations in 0-4 (RR: 1.74; 95% CI: 1.06-2.45) and 5-14 year (RR: 1.51; 95% CI: 1.02–2.23) old groups was also detected during heat waves occurred 205 206 in 2008 to 2009.

207

208 **Discussion**

Existing literature does not consistently suggest that heat waves increase the risk of deaths 209 among children. Some studies found that heat waves had a significant impact on children's 210 211 mortality in Australia (Nitschke et al. 2011), Great Britain (Rooney et al. 1998), Spain 212 (Basagaña et al. 2011) and South Korea (Kysely and Kim 2009), but other studies did not find a significant effect of heat waves on children's mortality (Son et al. 2012; Nitschke et al. 213 214 2007; Hutter et al. 2007; Huang et al. 2010; Fouillet et al. 2006). In the setting of extreme heat, young children experience greater risk of renal disease, respiratory disease, electrolyte 215 imbalance, fever. 216

217

The inconsistencies in the impact of heat waves on children's mortality across regions couldbe explained by the following reasons. (i) Different adaptabilities to heat waves: Due to some

220 factors, such as caregiver behavior, air conditioning use (Ostro et al. 2010), nutritional status, vaccination status and access to environmental infrastructures, the adaptability to heat waves 221 varies worldwide. (ii) Different characteristics of heat waves: Even a small change in the heat 222 223 wave definition had an appreciable effect on the estimated health impact (Tong et al. 2010). The existing literature looking at the impact of heat waves on children's health used various 224 definitions of heat waves, which might render the inconsistent results. Further, even for the 225 226 same heat wave definition (Nitschke et al. 2007; Nitschke et al. 2011), the intensity, duration, timing of every heat wave is different from each other, which also may cause different health 227 228 outcomes in children (Anderson and Bell 2009). (iii) Different age groups: Current studies assessed the effects of heat waves on the health of children of different ages. Some 229 researchers analyzed children aged 0-4 years (Huang et al. 2010) or 0-14 years (Nitschke et 230 231 al. 2007), and others focused on infants (Basagaña et al. 2011). Apparently, children of 232 different ages have exclusive characteristics, including their abilities to adapt to heat waves.

233

The findings of our review results illustrate that studies in Australia (Nitschke et al. 2007; 234 Nitschke et al. 2011), Austria (Hutter et al. 2007), South Korea (Kysely and Kim 2009), and 235 Spain (Basagaña et al. 2011) supported the assumption that heat waves had a greater effect on 236 237 mortality among children than adults, while the studies in Great Britain (Rooney et al. 1998) and Korea (Son et al. 2012), which considered children aged 0–14 years as a whole group, 238 challenged this assumption. This finding may indicate that very young children, especially 239 those aged under one year old, rather than old children, were more vulnerable to heat wave 240 impact when compared with adults. This age-specific vulnerability could be partly due to 241 their less developed thermoregulation ability and their low self-care ability. A study found 242 that heat-related mortality among infants was only detected for conditions originating in the 243 perinatal period in Catalonia, Spain, especially for digestive system diseases (Basagaña et al. 244

245 2011). This result indicates that the exposure to heat waves in the perinatal period may pose a
246 threat to children's health. The impact of maternal exposure to high temperature on adverse
247 birth outcomes has attracted an increasing research attention (Strand et al. 2011), but no study
248 has elucidated the relationship between heat waves and birth outcomes to date.

249

The published English studies regarding the impact of heat waves on children's mortality are 250 251 mostly from developed countries, but the drivers of mortality in developing countries are very different, with a high burden paid to infectious disease and dehydration. In the early 20th 252 253 century, when infectious disease also largely contributed to mortality in developed countries, heat waves were associated with greater number of deaths in the whole population 254 (Infoplease 2007), and also in children. For example, the 1911 heat wave in France was 255 256 associated with 40000 deaths, in which 29000 deaths were children (Rollet 2010). To some extent, the relatively lower impact of recent heat waves on deaths observed in children 257 reflects the improvement in care. 258

259

The key pediatric diseases or conditions significantly affected by heat waves include renal 260 disease, respiratory disease, electrolyte imbalance and fever (Knowlton et al. 2008; Kovats et 261 al. 2004; Leonardi et al. 2006; Nitschke et al. 2007; Nitschke et al. 2011). A recent analysis 262 of the contribution of extreme temperatures on years of life lost in Australia draws attention 263 264 to the concept that consideration of age and life expectancy are important in mortality studies (Huang et al. 2012), but to date there is no research concerning the impacts of heat waves on 265 children's years of life lost. The outcomes examined in these studies likely represent the most 266 267 extreme effects of a continuum of health impacts from heat. Other outcomes such as missed school days and impaired cognitive performance are other potentially important parts of the 268 total burden of disease from extreme heat. 269

270 Pediatric renal disease is an important adverse consequence of heat waves among children. Several studies have reported increases in hospital admissions for renal dysfunction during 271 periods of high ambient temperatures (Dematte et al. 1998; Kovats and Kristie 2006). 272 273 Exposure to extreme hot weather can induce heat-related conditions including hyperthermia and heat stress in children (Semenza et al. 1999), and the thermoregulatory physiological and 274 circulatory adjustments necessary to cope with extreme heat can place stress on the kidneys 275 276 and compromise the function of the renal system. Physiologically, children have poor ability to cope with heat, which can make them more vulnerable to the impact of heat waves. Heat-277 278 related renal dysfunction has also been attributed to some other factors, including direct thermal injury, prerenal insult, rhabdomyolysis, and disseminated intravascular coagulation 279 (Kew et al. 1967; Raju et al. 1973). Persons with diabetes have an increased susceptibility to 280 281 extreme heat (Semenza et al. 1999) and heat-related renal dysfunction, possibly due to preexisting renal conditions resulting in compromised kidneys (Mogensen et al. 1983). 282

283

284 Respiratory disease is another adverse consequence of heat waves in children (Kovats et al. 2004). It seems that the very young children are more influenced by heat in terms of 285 respiratory function. While the underlying mechanisms through which high temperatures may 286 increase the risk of hospitalization for respiratory diseases are unclear, we assume that young 287 children's susceptibility to respiratory disease during heat waves can be partly due to their 288 289 still developing respiratory system and generally smaller airways. For young children, exposure to heat waves may result in the exacerbation of their chronic respiratory disease, 290 which will result in mortality increases during the following hot days (Stafoggia et al. 2008). 291 292 One mechanism is through the effect of heat on formation of ozone, a known respiratory irritant. One recent study estimated that ozone-related asthma emergency department visits 293 for children could increase as much as 7% in a major metropolitan area due to temperature 294

driven changes in ozone concentrations (Sheffield et al. 2011). Further understanding of the
underlying mechanisms through which high temperatures influence respiratory disease is an
area where further research and development are clearly needed, especially because the
burden of such diseases is expected to grow as climate change continues (Mannino and Buist
2007).

300

During a heat wave period, in an effort to prevent hyperthermia and dehydration, the body's physiological mechanisms attempt to regulate electrolyte and water imbalance. In the setting of unreplaced fluid losses through perspiration and respiration, children, in particular, may face electrolyte imbalance (Knowlton et al. 2008). Electrolyte imbalance can precipitate heat exhaustion or heat cramps which in turn can further intensify electrolyte imbalance in the setting of continued exposure to intense heat.

307

Fever calls increased in heat wave days, especially for children 0-4 years (Leonardi et al. 308 309 2006). Ambient temperatures in excess of 41°C were often associated with hyperthermia (Feld and Jeffrey 2005). When the hypothalamus receives information that the body 310 temperature is lower than the setting of the internal thermostat, thermoregulatory responses 311 that conserve or produce heat are put into action. Heat is generated by shivering and is 312 conserved by vasoconstriction. If the body temperature is higher than the internal thermostat 313 314 setting, heat is lost by vasodilatation and increased sweating (Feld and Jeffrey 2005). Other responses include extracellular fluid volume regulation via arginine vasopressin, and 315 behavioral responses such as seeking a warmer or cooler environmental temperature (Feld 316 317 and Jeffrey 2005). If a body is involved in a sustained heat environment (eg. heat wave) and cannot seek a cooler environment, the physiologic responses may not suffice and increased 318 body temperature (i.e. hyperthemia) may occur. 319

Susceptibility to disasters decreases through activities such as prevention and mitigation measures that prevent or limit a population's exposure to the hazard, which is particularly important for children. Preparedness, response, and recovery capacity building increase resilience. Heat wave resilience is composed of (1) the absorbing capacity; (2) the buffering capacity; and (3) response to heat wave and recovery from the damage sustained (Boer and Dubouloz 2000).

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In the process of building resilience to cope with heat wave impacts, preventive measures are 327 328 essential not only because many causes of death may increase but also due to the fact that the mortality rate of some diseases, such as heatstroke, is high even when treated (Bouchama and 329 Knochel 2002). The mortality of patients with heat stroke admitted to intensive care units 330 331 during the 2003 heat wave in France was 62.6% (Misset et al. 2006). In addition, for those 332 patients who recovered from heat stroke, they suffered from severe sequelae (eg. persistent neurological sequelae) (Rav-Acha et al. 2007; Omero et al. 2000). The best method for 333 334 handling heat waves is through primary prevention, which means preventing exposure to extreme heat in the first place, rather than treating symptoms. Primary prevention includes 335 health education of children, and families about the risk factors of heat-related illnesses, their 336 signs and symptoms, and how to recognize and treat affected children. When they are aware 337 of the potential health effects of heat waves and the special physical and emotional 338 339 vulnerabilities of children, parents and caregivers can do a lot to protect children from the 340 potential harm (Luber and McGeehin 2008).

341

There is a growing appreciation amongst policy makers and societal actors that the policy context in which climate change adaptive decisions are made must be considered (Burton et al. 2002). The importance of policy on preventing children from the impact of heat waves

should be specifically focused, even though the heat wave prevention policy made for
children is scarce currently (Department of Human Services 2009). Governments at all levels
should make great efforts to reduce carbon emissions, including heat waves, and to mitigate
their impact on children's health.

349

Similar to studies of risk factors for heat vulnerability among the elderly (Semenza et al.
1996), heat vulnerability in children can be reduced by reshaping the built
environment(Rosenthal et al. 2007). Alert systems or early warning systems have been
developed (Díaz et al. 2006; Kalkstein et al. 1996; Metzger et al. 2009; Nicholls et al. 2008;
Pascal et al. 2006) and could perhaps be adjusted to see if their locally-relevant "threshold
temperatures" used to issue alert levels could be revised to apply to pediatric mortality or
morbidity, and give parents and caregivers advance warning to take precautionary measures.

358 Knowledge gaps

Existing research gives evidence that children are more likely to be associated with heat-359 related morbidity. Nonetheless, there are still substantial knowledge gaps. Several key 360 361 methodological challenges should be addressed in future research such as: 1. What is the definition of heat wave from a children's health perspective? 2. How is children's exposure to 362 heat wave best measured? 3. What are the most common health consequences of heat waves 363 among children? 4. What are the impacts of heat waves on years of life lost or disability-364 adjusted life years among children? 5. What are the differences in vulnerability to heat waves 365 among children of different ages? 6. How does vulnerability of children, particularly the very 366 367 young, vary between high income and low or middle income countries? 7. What are the best ways to project children's disease burden from heat waves under different climate change 368 scenarios? 8. What are the most effective modifiable risk factors of a public health response 369

- to the impact of heat waves on children's health? And how those risk factors differ by
- developmental stage of the child? And consequently, 9. What are effective heat wave
- 372 mitigation and adaptation strategies from a children's health perspective?
- 373

374 Conclusion

- 375 It is still inconsistent about whether heat waves significantly increase children's mortality.
- However, more heat-related deaths among infants are reported during heat wave periods.
- 377 Additionally, children are more likely to be attacked by respiratory disease, renal disease,
- 378 electrolyte imbalance and fever during persistent hot episodes.
- 379

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Study ^a	Location	Design	Time period	Heat wave definition	Outcome variable	Adjusted for potential confounders	Results
Basagaña (2011)	Catalonia, Spain	Case- crossover	May 15- October 15, 1983-2006	Maximum temperature over 95th percentile of the period 1983-2006	Total and cause- specific mortality	Yes	In children aged under one year old, the effect of heat waves was observed on the same day and was detected only for conditions originating in the perinatal period (RR:1.53; 95% CI: 1.16–2.02)
Fouillet (2006)	France	Descriptive study	2000-2003	NA	Total mortality	No	During heat waves, significant excess mortality was observed for male children aged less than one year (Observed mortality/expected mortality: 1.3; 95% CI:1.0-1.6)
Huang (2010)	Shanghai, China	Descriptive study	June 15- September 15, 2003	Consecutive periods of at least three days during which the daily maximum temperature is ≥35.0 °C	Total and cause- specific mortality	Yes	No significant mortality increase was found in children aged 0-4 years during heat waves (RR: 0.67; 95% CI: 0.24, 1.87)

Table 1. Characteristics of studies about heat waves and children's health

Study ^a	Location	Design	Time period	Heat wave definition	Outcome variable	Adjusted for potential confounders	Results
Hutter (2007)	Vienna, Austria	Time-series	1998-2004	Consecutive periods of at least three days during which the daily maximum temperature is \geq 30.0 °C and daily minimum temperature is \geq 25°C	Mortality	Yes	The relative risk of death on a heat wave day was highest in children under one year old (RR=1.25; 95% CI:0.82-1.90)
Knowlton (2009)	58 counties of California, USA	Descriptive study	July 8, 2006- August 22, 2006	Climatologic definition	Hospitalizations and emergency department visits	No	In heat waves, emergency department visits for electrolyte imbalance increased rapidly among 0-4 year age children (RR: 1.19; 95% CI:1.10–1.30)
Kovats (2004)	London, UK	Time-series	April 1, 1994- March 31, 2000	Maximum daily temperatures exceeded 30 °C.	Emergency hospital admissions	Yes	Hospital admissions increased during hot weather among children under five years old, but not among elderly and adults (Percent change: 0.24; 95% CI: 0.02 – 0.46)

Table 1. Characteristics of studies about heat waves and children's health (Continued)

Study ^a	Location	Design	Time period	Heat wave definition	Outcome variable	Adjusted for potential confounders	Results
Kysely (2009)	South Korea	Descriptive study	1991-2005	Consecutive periods of at least three days during which the daily heat index is ≥33.0 °C	Mortality	No	The relative increase in mortality was larger in children aged 0-14 years (183 excess deaths; 95% CI: 133 to 234) than in any other age group
Leonardi (2006)	England	Time-series	December 19, 2001- May 23, 2004	Mean temperature above 19 °C for the July episode and above 23 °C for the August episode	Calls to National Health Service ^b (fever, vomiting, difficulty breathing, heat- /sun-stroke)	Yes	In heat waves, the largest fever call rise was seen for children 0-4 years in Greater London and South East regions: 2.5 % increase in the proportion of fever calls (95 % CI 1.8, 3.3) for every 10°C increase in mean temperature
Nitschke (2007)	Adelaide, Australia	Case-series	1993-2006	Maximum temperature reached 35°C or above for three consecutive days	Hospital admissions, ambulances, and mortality	No	Hospital admissions for respiratory diseases decreased during heat waves among children aged 0-4 years (IRR:0.86; 95% CI: 0.76-0.97). No significant rise was found in mortality during heat waves among children aged 0-4 years (IRR:1.19; 95% CI: 0.82-1.71) and children aged 5-14 years (IRR:1.15; 95% CI: 0.58-2.29).

Table 1. Characteristics of studies about heat waves and children's health (Continued)

Study ^a	Location	Design	Time period	Heat wave definition	Outcome variable	Adjusted for potential confounders	Results
Nitschke (2011)	Adelaide, Australia	Case-series	2008-2009	Maximum temperature reached 35°C or above for three consecutive days or more with 35°C marking the 95th percentile for maximum daily temperature for the period 1993-2009	Hospital admissions, ambulance call outs, emergency department presentations and mortality	No	 During heat waves, there were: 1). Significant rise of renal hospital admission in 5-14 year age group (RR: 2.64; 95% CI:1.47-4.73); 2). Significant rise of emergency department presentations in 0-4 (RR:1.02; 95% CI: 0.92-1.13) and 5-14 (RR: 1.04; 95% CI: 0.95-1.14) year old groups; 3). Significant rise of mortality in 0-4 year old group (RR: 3.23; 95% CI:1.30-7.99)
Rooney (1998)	England, Wales, and Greater London	Descriptive study	1995	NA	Mortality	No	Mortality among children aged 0-15 years increased during 1995 heat waves in England and Wales (Percent change in deaths: 4.6%), and Greater London (Percent change in deaths: 13.0%).
Son (2012)	Seven cities in South Korea	Time-series	January 1, 2001- December 31, 2007	Mean temperature reached at or above the 98th percentile for the warm season for ≥ 2 consecutive days	Mortality	Yes	No significant percent change was found in children aged 0-14 years during heat waves in Seoul.

Table 1. Characteristics of studies about heat waves and children's health (Continued)

a These studies are ordered by the name of the first author.

b A nurse-led helpline which provides health-related information and advice and directs callers to the appropriate health service and self care.

Abbreviations: CI, confidence interval; IRR, incidence relative risk; NA, not applicable; RR, relative risk.

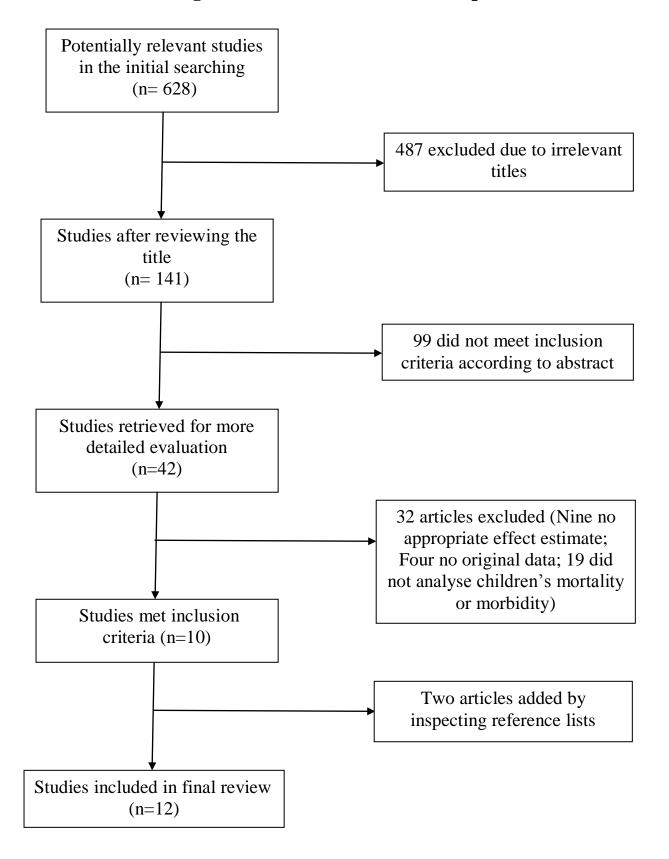


Figure 1 The literature selection process