Abstract
The overall objective of this study is to examine the literature investigating the associations between traffic-related air pollution (TRAP) and the incidence and prevalence of childhood asthma throughout a critical literature review. The study examines and demonstrates the association between TRAP and childhood asthma, the literature associated with it, and the gaps in the current state of research. The exposure assessment methods currently in use in the literature are also overviewed and critically discussed for strengths and limitations.

Keywords
air pollution, childhood asthma, exposure assessment, exposure surrogates

1 Introduction
Air pollution is a commonly recognized external cost of the use of motor vehicles (Barabas, 2013). In many urban areas nowadays, road traffic is the principal source of outdoor air pollution which has been recently linked to around 4 million global preventable deaths per annum (Solvang Jensen, 1999; Colville et al., 2001; European Environment Agency, 2007; Health Effects Institute, 2010; World Health Organization, 2014a). Despite many air quality improvements in recent decades, air pollution continues to pose a significant threat to human health and well-being worldwide (World Health Organization, 2006; Lim et al., 2013; Vidal, 2014). In particular, traffic-related air pollution is a key component to the ambient air pollution mix in present urban areas, and has been linked to a wide spectrum of global disease (Health Effects Institute, 2010).

In this study, the term traffic-related air pollution (TRAP) refers to the concentrations of primary and secondary air pollutants, elevated above background levels due to motor vehicles emissions. Traffic-related primary pollutants include carbon monoxide (CO), carbon dioxide (CO2), nitrogen oxides (NOx), ammonia (NH3), particulate matter (PM), and hydrocarbons (HC). Traffic-related secondary pollutants are those formed mainly by the various chemical reactions which take place in the atmosphere such as ozone (O3), secondary particulates and secondary NOx.

In particular, recent research provides evidence indicating that significant associations exist between residential traffic-related pollutant levels and asthma occurrences (Brauer et al., 2007; Salam et al., 2008).

With childhood asthma currently being a condition that has a profound impact on the quality of life for a large number of the population, especially in childhood; when individuals are most vulnerable, working towards establishing a clearer understanding of the interactions between a common exposure such as that to TRAP and the onset/ prevalence of the disease is clearly a priority. Such an understanding may be offering a partial explanation of the relatively rapid changes in prevalence, in times when traffic air pollution and certain fleets became more dominant.
This study aims at examining the question of whether early-life exposure to TRAP can cause asthma to develop in children and highlight the current limitations in the evidence base.

2 Traffic-related air pollution and childhood asthma

A sharp rise in the prevalence and incidence of asthma has been reported in many parts of the world, and particularly in the more developed and industrialized societies (Mutius, 1998; Beasley, 2004; Ferguson et al., 2004; Das, 2006; Anderson et al., 2007). These sharp increases in prevalence remain unexplained fully (Aubier, 2000; Bracken et al., 2002; London, 2007; Clark et al., 2010) and the prevailing views on the link between asthma and air pollution are conflicting. Although there is ample evidence that (traffic-related) air pollution can exacerbate asthma and trigger symptoms in persons who already have the condition, the question of whether it is an independent risk factor for the onset of asthma in children remains unresolved.

Different studies that investigated this matter produce conflicting results making consensus difficult to reach (Gasana et al., 2012; Gowers et al., 2012; Lindgren et al., 2013). In these studies and in others, however, the effect of TRAP exposures was repeatedly shown as spatially heterogeneous, where it is mostly marked amongst those living within 50 to 75 meters of a busy road or a highway, whilst diminishing with increasing distance from the road (Venn et al., 2001; McConnell et al., 2006; Perez et al., 2012; Price et al., 2012). These observations are particularly relevant as TRAP is known to be spatially heterogeneous in space and time. Concentrations of common TRAP vary significantly over short distances as short as tens of meters.

In the investigated research articles, some authors reported an increase in the odds of incident asthma with increased exposure to different traffic-related air pollutants (Gauderman et al., 2002; Nicolai et al., 2003; Jerrett et al., 2008; Gehring et al., 2010), whilst others reported null association (Sahsuvaroglu et al., 2009; McConnell et al., 2010; Mölter et al., 2014). It is however possible that the different designs and exposure assessment methods in the different studies had an important role to play in these conflicting findings. The different studies employed different surrogates and/or air pollution measurements to represent exposure to TRAP levels and to then investigate the effects of this exposure. The collected studies differ in many ways as follows:

(1) the exposure assessment methods employed and the pollutants or traffic-related metrics investigated, (2) the definitions and ascertainment of asthma as an outcome, in (3) the study’s design and (4) the study’s sample’s size; all of which are factors that can explain some of the variability and inconsistency of the results from different studies.

3 Exposure assessment methods: strengths and limitations

Because it is neither practical nor feasible to measure the personal exposure of individuals to TRAP at all points and at all times, different surrogates for the human exposure have been customarily employed in different epidemiological studies as a reasonable compromise for actual measurements. This part of the paper synthesizes the reviewed papers in reference to the exposure assessment methods employed whilst drawing attention to the main gaps found in the current state of research.

In the reviewed literature a wide variety of exposure metrics were employed to study the effects of air pollution on children’s asthma. These exposure surrogates included:

- Proximity to a ‘freeway’ or a ‘major road’;
- Land use regression models;
- Dispersion models;
- Mathematical and statistical interpolation;
- Averaged measurements from fixed monitoring sites; and
- Hybrid models

Many of the exposure surrogates that were used in the literature cannot comprehensively represent the misaligned spatio-temporal nature of common vehicular air pollutants.

3.1 Proximity to a ‘freeway’ or a ‘major road’

In the studies using this metric, subgroups of people living close to ‘major’ roads or the nearest freeway are compared to those who live at further distances. The distance to the road(s) of interest is typically objectively estimated using GIS. Although this exposure metric is intuitive and simple, it assumes a road of a certain type or size corresponds to a certain amount of traffic, and fails to take into account the compounded effects of proximity to multiple roads. As this metric cannot take meteorological conditions into account, it has a limited directional dependence, and automatically assumes that all pollutants disperse in a similar manner, which is inaccurate.

Six out of the sixteen collected studies used this metric, either in isolation or alongside other criteria. In five of these studies, there was little evidence for an effect of major roadway proximity alone, where weak local and no associations between proximity to major roads and asthma were reported (Gauderman et al., 2005; McConnell et al., 2010). The exception of this was in the study by Gauderman et al. (2005) where the residential distance from the nearest freeway was the distance investigated. In their study, simple distance to a freeway was as strongly associated with a life history of asthma as was measured ambient NO\textsubscript{2}. On the other hand, Venn and co-workers found that the increased odds of childhood wheeze occurrence in relation to road proximity were local and only visible within 150 meters from the road. Similarly, McConnell et al. (2006) and follow-up work in McConnell et al. (2010) showed high risks of asthma in children residing at less than 75 meters from a major road, with susceptibility found to
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have significantly increased in long-term residents with no parental history of asthma. The authors also observed that these risks decreased to reach background rates at 150–200 meters from the road as illustrated in Fig. 1. Finally, Lindgren et al. (2013) and Sahsuvaroglu et al. (2009) found no association between this exposure metric and the occurrence of asthma.

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These weak, localized and absent effects can be possibly explained by one of three reasons. Firstly, an effect of the exposure could be absent. Secondly, an existing effect may have been concealed because of the crudity of the exposure metric employed. Proximity to major roads assumes that a classified or a numbered road corresponds to a certain amount of traffic, usually with a relatively wide range. It furthermore lacks important information on traffic density, the vehicular mix and fails to take into account the completely different nature of dispersion for different pollutants. The compounded influence of proximity to other roads is also often neglected when using such a metric (Brook, 2012), and the unusual concentrations of traffic-related air pollution in street canyons cannot be accounted for. Finally, studies finding a localized effect when using this metric may have implications on which pollutant(s) is more likely to be responsible for the observed effect(s), as the composition of fresh vehicular exhaust is different than that of the aging exhaust, and the different dispersion manners for different pollutants can cause them to vary significantly in concentrations between the roadside and further. Here, ultra-fine particles are pollutants which attracts attention as their concentrations are relatively high nearby roadsides, with very steep decreases with increasing distances from the source.

3.2 Land use regression models

Land Use Regression models (LUR) were the second exposure metric that has been used in four out of the sixteen collected studies and in other recent major air pollution and health research projects such as the European Study of Cohorts for Air Pollution Effects (ESCAPE) (Eeftens et al., 2012; Beelen et al., 2013).

LUR models seek to predict air pollution concentrations at sites of interest based on previous monitoring data and the surrounding land use and traffic characteristics (Jerrett et al., 2004). LUR have recently gained considerable interest and have become more widely used to estimate the exposure of cohort studies participants to air pollution (Ryan and LeMasters, 2007). This increased popularity of this method is possibly due to the facts that LUR models require relatively little time to establish, input data that is easy to collate, have relatively low costs and have the capability of showing changes in the spatial patterns of pollutants over large distances.

Although LUR have the capacity to show changes in the spatial patterns of pollutants, they can only reflect the predictors that were used in establishing the model, suffer from varying uncertainties amongst different pollutants (Brook, 2012), and the true contribution of traffic to the regression is not always known or reported (Health Effects Institute, 2010). Similarly, to the ‘proximity to major roads’ metric, LUR models cannot take into account the effects of terrain, topography and meteorological conditions on air pollution concentrations which can greatly influence exposure experiences. As the models are also based on actual measurements made for the pollutants of interest, the model’s outputs are sensitive to the locations of the sampling sites, and to a lesser extent, the number of air sampling sites (Ryan and LeMasters, 2007). Finally, as the used predictors can have a high contribution to the final regression, the meaningfulness of these predictors is an essential point, especially when moving to study areas with much different land use and topography (Jerrett et al., 2004).

In work undertaken by Gehring et al. (2010), the authors used predictor variables that are related to motor vehicles traffic as the model’s predictors, such as the traffic intensity in the area of investigation and the population density which can have a role in driving trips from and to a zone. On the other hand, Sahsuvaroglu et al. (2009) used seven predictors to set up their model as follows: traffic density, open land use within 500 meters, industrial land use within 200 meters, presence of a highway within 50 meters, presence within 1000 meters from downtown industrial core, presence downwind from a highway, and distance to the lake. Work by Clark et al., 2010 relied on LUR models developed by Henderson and co-workers who used fifty-five variables describing each sampling site as predictors of the LUR model in a GIS (Henderson et al., 2007). Even with this high number of predictive variables used for the models development, adjusted R2 values ranged from 0.39 to 0.62. Finally, the LUR models developed in work by Mölter et al. (2014) employed eight covariates including traffic intensity, emissions, land use and physical geography as predictor variables for the final models. The R2 values of their models ranged from 0.56 to 0.86.

In all of these studies it could be argued that these predictors are still not allowing for some important factors that are of vital influence on the traffic-PM and traffic-NOx concentrations,
which these models predicted. Some of these factors include the vehicular mix and fleet’s composition and speeds, the emission factors for the main classes of the vehicles composing the fleet and the street characteristics and meteorological conditions which can greatly impact the dispersion processes.

As for different pollutants’ uncertainties, good agreement between the long-term measured and predicted NO₂ and PM$_{2.5}$ was shown in LUR models (Montagne et al., 2013), even from nitrogen dioxide samples collected 12 years apart (Cesaroni et al., 2012). It is important to note though that agreement studies are usually based on averaged measurements similar to those initially used to develop the LUR models, and averaged measurements have their own inherent limitation in assessing the exposure. There is also another important question of whether NO₂, the pollutant most studies by LUR models, is a reasonable proxy for traffic-related air pollution.

### 3.3 Dispersion modelling

Dispersion modelling is a technique which involves the construction of a dynamic model that utilizes emission rates from different sources such as motor vehicles, alongside meteorology and boundary layer conditions to simulate the dispersion process, and to predict the following ambient concentrations of air pollutants (Briggs et al., 1997; Health Effects Institute, 2010). Estimates derived from dispersion models are continuous exposure metrics, which incorporate traffic flows, meteorology and atmospheric chemistry when making the final ambient pollution levels predictions. The models generally rely on Gaussian plume equations to predict the pollution concentrations, in which steady-state meteorological conditions are assumed. Gaussian plume models also assume that there are no chemical or removal processes taking place (Vardoulakis et al., 2003).

In recent years, dispersion models have been jointly used with GIS. The models can make air pollution predictions for relatively large areas, and can assess severe episodic short-term and long-term exposures at receptor points of interest. The outputs of dispersion models have the highest temporal and spatial resolution in relation to all other exposure metrics found in use in the reviewed literature. Furthermore, many of the commonly used dispersion models such as INDIC AIRVRO and Atmospheric Dispersion Modelling System (ADMS) can also take into account a street canyon contribution in the most polluted segments using their built-in street canyon model, namely; the Danish Operational Street Pollution Model (OSPM).

Five out of the sixteen collected studies were found to have used dispersion models to assess the exposure, either in isolation (Nordling et al., 2008; Gruzieva et al., 2013), or alongside other metrics (Gauderman et al., 2005; McConnell et al., 2010; Lindgren et al., 2013). In these studies, the use of dispersion models to assess the exposure is advantageous over the previously described metrics as it allows a more accurate assessment of the impact of variable sources’ emissions loads on ambient air quality, and provides outputs that are characterized by a relatively high temporal and spatial resolution. However, although such models can better reflect the different dispersion manner for different pollutants and take into account meteorology, emission rates and some atmospheric chemistry, their output’s quality is highly dependent on that of the input. The validity of the dispersion modelling results is at the mercy of the emission factors used to calculate inputted emission rates (Barrat, 2013). Emission factors and consequently emissions’ estimates remain an essential source of uncertainty in dispersion modelling; especially when considering the real world divergence of the NO₂ emission factors for certain growing sub fleets such as diesel cars. The models must therefore be calibrated correctly in order to realize their advantages (Health Effects Institute, 2010), and cross validated with monitoring data where possible to test their performances.

### 3.4 Interpolation techniques

Alongside dispersion models, interpolation techniques can also be used to map and predict ambient air pollution concentrations. Interpolation models can assign pollution concentrations to locations of interest based on some surrounding measured values, and various underlying mathematical formulas or statistical models (ERSI, 2008).

The three most commonly used interpolation tools in air pollution and health research are the Inverse Distance Weighting (IDW), the Splines and Kriging (Ranade et al., ND). These three tools were used in the study by Sahsuvaroglu and co-workers to assess the exposure, where no associations were found with incident asthma. Their results may be explained by either the absence of the effect, or the by inherent limitations of these metrics which are described next. The Spline tool works by fitting a mathematical function across all input data values to create a prediction surface. Spline works best for surfaces that vary mildly (Ranade et al., ND), and does not perform well when large variations along the surface can occur within short distances (U.S. Environmental Protection Agency, 2004). As previously discussed, concentrations of common vehicular pollutants are spatially misaligned and can significantly fluctuate even over no more than a few tens of meters (Briggs et al., 1997; Sharma et al., 2005; Bell, 2009), therefore, results emerging from the use of this method should be treated with caution.

On the other hand, the IDW is a simple deterministic interpolation method based on the principle that sample values closer to the prediction location have more influence on prediction value than sample values which are further away. The main shortcoming of this approach is its “bull’s eye” effect where the highest values will be assigned to points that are near the sampled locations (Ranade et al., ND). The other problem with the IDW tool is that the range of the predicted values is at the mercy of the input. The quality is highly dependent on that of the input. The validity of the dispersion modelling results is at the mercy of the emission factors used to calculate inputted emission rates (Barrat, 2013). Emission factors and consequently emissions’ estimates remain an essential source of uncertainty in dispersion modelling; especially when considering the real world divergence of the NO₂ emission factors for certain growing sub fleets such as diesel cars. The models must therefore be calibrated correctly in order to realize their advantages (Health Effects Institute, 2010), and cross validated with monitoring data where possible to test their performances.
Protection Agency, 2004). This would become an issue when no measurements were made within a certain area(s); arguably causing a biased spatial heterogeneity when investigating the cause and effect, and an existing pattern is likely to be concealed where no representative measurements were made.

Finally, kriging is the geostatistical technique most commonly used in the air pollution field (Jerrett et al., 2004). Kriging uses a tool known as the semivariogram for making predictions. The semivariogram model and its underlying statistical assumptions are an ad hoc approach, especially for making predictions of air pollution surfaces. This was discussed in details elsewhere, after kriging in GIS was previously used in a case study in Sheffield.

### 3.5 Averaged measurements

Averaged measurements of different pollutants from continuous regulatory monitoring stations or from fixed sited measurement equipment such as NO\textsubscript{2}, Palmes tubes were another exposure surrogate used in the literature reviewed.

Three studies out of the sixteen collected studies used continuous measurements obtained from fixed regulatory air quality monitoring stations to reflect the population exposure to traffic-related air pollutants (Hwang et al., 2005; Clark et al., 2010; McConnell et al., 2010). Four other studies have undertaken their own measurements campaigns and used averaged and temporally adjusted measurements to reflect the populations’ exposure, either in isolation or alongside other metrics (Brauer et al., 2002; Nicolai et al., 2003; Gauderman et al., 2005; Jerrett et al., 2008,).

Although continuous monitoring stations are cost effective and have a very high and continuous temporal resolution, they have important shortcomings. Because of the distinct spatially heterogeneous nature of TRAP, the fact that air quality monitoring stations cannot be present at all locations of interest, and that their locations are usually based on regulatory rather than scientific purposes (Bell, 2009), this approach is problematic when assessing this exposure and can lead to misclassifications. Problems can arise when using this method either because the exposure can be overestimated when considering populations residing in industrialized areas (Mölter et al., 2014), or it can be underestimated when specifically investigating TRAP. Indeed, fixed monitoring stations were found to significantly underestimate the exposure concentration levels experienced by individuals exposed in urban environments, especially when in or near transport microenvironments (Kaur et al., 2007). Moreover, the use of monitoring stations to assign the exposure for large populations will most certainly conceal people’s differences because of a mismatch between the data used to estimate exposure and the actual subjects’ locations.

Secondly, because of the temporally heterogeneous nature of traffic-related air pollution, averaged measurements from fixed sited measurement equipment, even if spatially well-distributed, will fail to pick up the variations in pollutants concentrations in real time. It is also likely to underestimate occasional exposure as severe episodic pollution levels can be concealed in the averaged values if they don’t occur often enough.

### 3.6 Hybrid models

Hybrid models were also employed in some studies. These models are largely based on the above mentioned basic exposure metrics combined with each other or with other measurements such as vehicles counts, traffic flows etc. The limitations of such metrics can be explained by the inherent limitations of its composing elements.

Against this background, it is clear that the development of models to assess exposure to TRAP is a priority area for future research. Evaluating the health effects of exposure to TRAP is subtle to the method of the exposure estimation. Many of the exposure surrogates that were used in the literature are crude, and do not relate well to the misaligned spatio-temporal nature of common vehicular air pollutants.

### 4 Early-life exposure

In the vast majority of the studies confirming a link between traffic-related air pollution and the onset of asthma, the effects are stronger and highlighted in the early years of the child’s life. Some of the papers reviewed demonstrate that a critical exposure window could exist for children before their third year and whilst they are still in utero (Brauer et al., 2002; Gauderman et al., 2005; Nordling et al., 2008; Clark et al., 2010; Gruzieva et al., 2013). These findings are in line with other research, showing that exposure to air pollution before three years of age can have long-term consequences on the lung and the lung development and suggesting that the timing of the exposure is possibly a crucial factor in the inception of asthma.

### 5 Identified critical pollutants

In line with growing evidence, no safe threshold for TRAP can be drawn from the reviewed studies, which below zero risk would occur. Numerous studies are demonstrating that the effects of air pollution on the respiratory health are evident, even at concentration levels that are officially considered safe. For instance, after adjusting to potential confounders, a 5 parts per billion increase in the average NO\textsubscript{2} exposure during the first year of a child’s life was found to be associated with a 1.17 odds ratio for physician-diagnosed asthma (Nishimura et al., 2013).

From the pool of pollutants that were investigated throughout the collected studies, exposure to NO\textsubscript{2} showed the most consistent and significant associations (described by the odds ratios) with the incidence and prevalence of childhood asthma, followed by exposure to PM\textsubscript{10} and PM\textsubscript{2.5}. Other unregulated pollutants such as NH\textsubscript{3} and UFPs – with the latter being arguably most toxic because of its physicochemical properties were not described in any of the studies.
6 Conclusion and Future Work

There has been increasing evidence to suggest that TRAP exposures are associated with the onset and prevalence of childhood asthma. However, an analysis of the literature suggests that the development of more refined exposure models to assess exposure to TRAP should be a priority area for future research.

Evaluating the health effects of exposure to TRAP is affected by the method of the exposure estimation and many of the exposure surrogates that were used in the literature continue to be a limitation. More refined exposure models are needed, and will arguably produce the most robust associations when investigating the potential health effects of TRAP. No study was found to assess the effect of repeated short-term severe exposures and future studies may gain new insight from investigating the effect exposure to episodic severe concentrations of TRAP, alongside continual long-term exposures. At present, there are no available studies on the particular effect of long-term exposure, or short-term repeated exposures to ultra-fine particles (UFP) on the incidence or the prevalence of asthma in children. This is in part a result of the high costs and logistic difficulties in measuring levels of UFPs for large populations on the one hand. On the other hand, an incomplete understanding of the sources and vehicles’ emissions trends of UFPs contribute to additional methodological difficulties when trying to model and predict UFPs levels.

Current dispersion models are not designed to predict the UFPs ambient concentrations, especially when associated with traffic sources. Recent research, however, demonstrates that UFPs warrant special examination in the context of asthma and other respiratory disease, and that it is the fraction of particulate matter where most of the mixture’s toxicity seems to reside. Future studies could benefit from investigating associations with UFPs which can be modelled based on other explanatory variables at the local scale such as traffic NOx. As it stands, the impact of TRAP on asthma prevalence is potentially significant due to the large number of children exposed. Policy decision-making should address these impacts by reducing childhood exposures through relevant transport and land use policies.

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