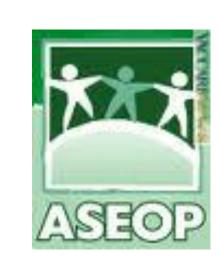




ASEOP
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# Background

Exposure to pesticides used in agriculture has been suggested to be a risk factor of childhood leukemia, though definitive evidence on this relation and the exact identification of the specific pesticides involved are still lacking.

# Methods

We carried out a population-based case-control study in the Modena and Reggio Emilia provinces of Emilia-Romagna region, Northern Italy. We included the 111 childhood leukemia cases newly-diagnosed from 1998 to 2011, and 444 controls matched by age and sex to the cases. Each children's house was georeferred and positioned on the provincial territorial cartography.

#### Study population

#### Cases: 111

All cases of leukemia (0-14 years) diagnosed in period 1998-2011 in Modena and Reggio Emilia provinces.

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#### Controls: 444

4 children for each case randomly extracted from the general population and matched for:

- Province of residenceYear of birth
- Gender

**Table 1**. Main classes of pesticides frequently used in different crops in Modena and Reggio Emilia farms.

Crops	Classes of pesticides Phosphorganics, Triazoles, Phenylalanines			
Vineyards				
Orchards	Phosphorganics, Triazoles, Phenylalanines, Neonicotinoids			
Arable	2,4D, MCPA, Dicamba, Azatrine, Pyrethroids Neonicotinoids, Pyrethroids, Phenylalanines			
Vegetable				

### Exposure assessment

Through remote sensing mapping and Geographical Information System analysis, we assessed land use in the 100-meter radius around children's homes. In particular, we examined four types of crops (vineyards, orchards, vegetable and arable) characterized by the use of specific pesticides potentially involved in childhood leukemia etiology. Exposure to air pollutants from motorized traffic (benzene and  $PM_{10}$ ) was assessed for each subject in study using detailed emission and dispersion modeling.

**Figure 2**: Geo-positioning on the provincial territorial cartography of children's homes.

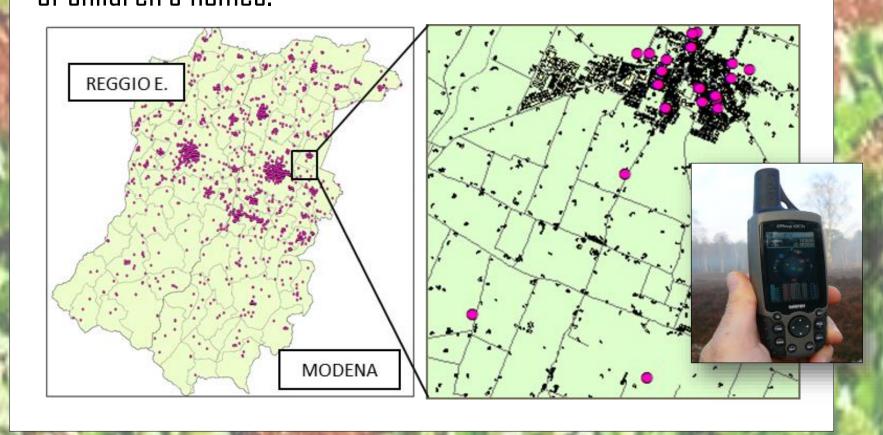


Figure 3: Soil analysis.

Types of crops:

Arable

Orchards

Vineyard

Vegetable

Figure 1: Geographical Information System: layers overlapping.

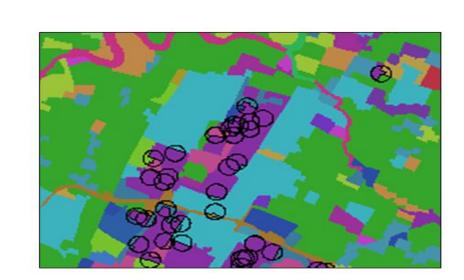
Layer C: Geo-positioning of cases and controls

Layer B: Land use cartography: types of crops

Layer A: Provincial territorial cartography

**Figure 4**: 100-meter radius buffers around children's houses and land use cartography with different types of crops.





**Table 2**. Adjusted odds ratios (95% confidence intervals) of leukemia for children with >95% of arable soil (crops such as corn, wheat, soya, sugar beet) in the 100-meter circular buffer around the child's home building).

	Arable soil					
			≥95%			
	Cases/ controls	OR	Cases/ controls	OR (95%CI)	P-trend <sup>8</sup>	
All subjects	80/326	1.00	3/6	2.14 (0.53-8.65)	0.285	
Age <5 year Age ≥5 year	36/150 44/176	1.00 1.00	2/2 1/4	5.64 (0.76-41.73) 0.90 (0.10-8.15)	0.090 0.926	

<sup>1</sup>Adjusted for annual average atmospheric concentration ( $\mu$ g/m³) of benzene; estimates not possible for subject diagnosed after 2009 or residing in mountain municipalities

<sup>2</sup>Urban and residential use <95% used as reference category <sup>3</sup>P for linear trend based on continuous values of land use

# Results

We computed the odds ratios (OR) of the disease and their 95% confidence intervals (CI) according to type and percentage of land use around child's home and adjusting for air pollution from vehicular traffic.

Adjusted OR (95%Cl) of leukemia for children with >95% of arable soil (crops such as corn, wheat, soya, sugar beet) around their residence, characterized by use of Dicamba, 2.4-D, MCPA, diazines and pyrethroids, was 2.1 (0.5-8.7). Risk further increased among children aged less than 5 years (OR= 5.6 (0.8-41.7).

No other association between specific crops and disease risk emerged.

## Conclusions

The excess childhood leukemia risk observed among children residing in rural areas close to arable soils seems to indicate a role of passive pesticide exposure in disease etiology.

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