

Epidemiology of Brain/Nervous System Tumors in Children

Parviz Ghadirian, Ph.D.,^{1,4} Kazem Fathie, M.D., Ph.D.,¹ Jean-Pierre Thouez, Ph.D.¹

¹Department of Nutrition, Faculty of Medicine, University of Montreal, and Centre de recherche, CHUM-Hotel-Dieu, Montreal, Quebec, Canada - Hospital Sainte-Justine, Research Center, Montreal, Quebec, Canada

⁴Chairman of the American Academy of Neurological and Orthopedic Surgeons

¹Department of Geography, Faculty of Arts and Science, University of Montreal, and Centre de recherche, CHUM-Hôtel-Dieu, Montreal, Quebec, Canada

Introduction

In the United States cancer is relatively less frequent among children. It has been estimated that¹ only 1 in 540 children will develop cancer. The relative rate of cancer occurring before the age of 15 years and among children less than 5 years of age is around 40%.² Although this disease is uncommon among young individuals, the mortality due to it is high, accounting for more than 11% of all causes of death among children.³ In other words, childhood cancer is the second leading cause of death, after accidents. The age-adjusted rate for all cancers combined among white children aged less than 15 years in the United States is 1.8% per 100,000 children per year. The most common malignancy in children is leukemia which represents almost one-third of all cancer sites.⁵ Malignant neoplasms of the brain and central nervous system (CNS) are the next most common cancer in children, at about one-fifth of all cancer sites. In other words, 1 out of 5 cancers in children involve brain tumors. In the United States, it appears that the major types of tumors in the CNS are astrocytomas (21%), medulloblastomas (19%), gliomas (18%), glioblastomas (14%), astroblastomas (13%) and, finally, ependymomas.⁶

In general, because of early detection and progress with therapeutic methods, mortality due to malignant diseases as a whole during childhood has been decreasing. Although mortality from all cancers in

England, a significant rise (1.8% on average) in the annual incidence rate for all CNS cancers, particularly for neuroectodermal tumors (3.1% elevation) was observed in 1974-1995.⁸ These increases are not explained by an increment in the proportion of histologically-verified tumors.

Genetic Factors

Susceptibility to certain types of childhood cancers may also follow an autosomal dominant pattern of inheritance. For example, it has been estimated that 40% of retinoblastomas are due in part to an autosomal dominant germ cell mutation.⁹ Neurofibromatosis is another autosomal dominant syndrome associated with the occurrence of cancer in childhood.¹⁰

Socio-economic Status

It appears there is no significant association between socioeconomic status, ethnicity and CNS tumors.⁸

Age

In general, there is no apparent age-dependency of brain and CNS cancer sites, but a specific age pattern averages for some CNS malignancies. For example, neuroblastoma is more common among children aged less than 1 year, and is rare after 5 years of age.¹¹ The highest incidence rate of astrocytomas belongs to 3 years old children. Its rate decreases afterward, while medulloblastoma occurs mostly at 5 years, and glioblastoma is more common among 7 year-old.¹²

In general, the incidence of primary cancer and CNS tumors seems to be increasing in the United States among all age groups. In a large cohort of children less than 15 years old,¹³ a modest rise in

Infectious diseases

In a case-control study in Greece, a significant association was found between influenza in pregnant women and tumor occurrence in the index child. In other words, the risk increased more than 3-fold among children from mothers who had influenza during pregnancy.¹⁴

Gender

The lowest male:female ratio (0.3) for all brain and CNS tumors in the world comes from Mali (Bamako), and the highest ratio (1.8) from China (Tianjin). In North America, The lowest male:female ratio (0.3) for all brain and CNS tumors in the world comes from Mali (Bamako), and the highest ratio (1.8) from China (Tianjin). In North America, the lowest male:female ratio is seen among Los Angeles the highest ratio (1.3) among the non-black population of the Greater Delaware Valley in the United States.¹⁵

Contact with Domestic and Farm Animals

It has been suggested that domestic and farm animals may increase the risk of brain cancer in children. In a large in the United States in 1984-91, it was found that childhood brain tumors were more common among children mothers had been exposed to pigs (OR = 3.8) and horses (OR = 2.2) during the index pregnancy.¹⁶ It seems that children diagnosed with primitive neuroectodermas have a higher risk for childhood brain tumors with personal and maternal prenatal exposure to swine (OR = 4.0 for child and 11.9 for mother) or poultry (OR = 3.0 for child and 4.0 for mother). A non-significant increased risk for childhood brain tumors has also been found for children mothers who had worked on livestock farms compared with controls.¹⁶

Smoking

A large body of evidence implicates cigarette-smoking in the etiology of childhood cancer, particularly brain tumors. Most of these studies concentrated on the effect of maternal smoking. In a case-central study in Shanghai¹⁷ paternal preconception smoking was related to a significantly elevated risk of childhood cancers, particularly brain tumors (OR = 2.7); the second most common cancer in children after leukemia, yet its etiology remains unknown. N-nitroso compound precursor is one of several dozens of toxic compounds downstream of tobacco smoke. Fetuses and infants have incompletely-formed blood-brain barriers that may allow the passage of carcinogenic tobacco metabolites into the CNS and initiate the formation of neural tumors.¹⁸

Other factors

In a nested case-central study utilizing data from a national birth registry in Sweden,¹⁹ the risk for brain tumors (all types) was elevated significantly when the mother of the index case had been exposed to oral contraceptives prior to conception (OR = 1.3) or Penthrane (OR = 1.5) during delivery as well as after treatment for neonatal distress (OR = 1.6) and, finally neonatal infections (OR = 2.4).

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4To whom correspondence should be address:
 Epidemiology Research Unit, Research Centre-
 CHUM-Hbtel-Dieu
 3850 St-Urbain
 Montreal (Quebec) H2W IT8
 Tel. (514) 843-2742; Fax: (514) 843-2715
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 Nevada

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