



Review Article

Epidemiology and determinant factors of neural tube defect: Narrative review

Melese Linger Endalifer¹, Gedefaw Diress²

¹Departments of Midwifery, ²Public Health, Woldia University, Woldia, Amhara, Ethiopia.

E-mail: *Melese Linger Endalifer - melselinger@gmail.com; Gedefaw Diress - gedefawdiress@gmail.com



***Corresponding author:**

Melese Linger Endalifer,
Department of Midwifery,
Woldia University, Woldia 400,
Amhara, Ethiopia.

melselinger@gmail.com

Received : 03 March 2020

Accepted : 27 March 2020

Published : 25 April 2020

DOI

10.25259/SNI_84_2020

Quick Response Code:



ABSTRACT

Background: The epidemiology of neural tube defect (NTD) is face ignorance from the global community. However, the problem is complex and it is a cause for child mortality and morbidity. We provide the latest insights with respect to determinant factors of NTD.

Methods: Google Scholar and PubMed were systematically searched to identify potential research articles concerning the epidemiology and its determinant factors of NTD.

Results: The epidemiology of Neural tube defects increased in some countries. The epidemiology and determinant factors were varies across countries, geographical regions and socioeconomic status of the populations. In general, the determinant factors of NTD were summarized as behavioral, nutrition-related, environmental, medical illness, and health service-related factors.

Conclusion: Birth defect is fatal which affects the new generation; specifically, NTD is the problem of middle- and low-income countries. It is a direct cause for neonatal and perinatal mortality rate globally. Even if little factors identified, yet conducting experimental and clinical trial researches are a better approach to slow down the progress.

Keywords: Determinant factors, Epidemiology, Neural tube defect

BACKGROUND

Neural tube defects (NTDs), including spinal bifida, anencephaly, and encephalocele, occur when part of the neural tube, which forms the spine, spinal cord, skull, and brain, fails to close between 21 and 28 days after conception – before the women realize that they are pregnant.^[1] The physiological development of central nervous system (CNS) develops from ectoderm by folding it to form a tube. The tube has two ends caudally develop to spinal cord and rostral part develop to CNS (forebrain, midbrain, and hindbrain).^[13]

Basically, the exact cause of birth defect is unknown, but nowadays, it is recognized partially, broadly the cause is classified into two that is genetic (preconception) and partially genetic causes (postconception).^[1]

The most common form of NTD is spinal bifida which results functional disabilities, that is, loss to control the bladder and bowel and paralysis of the legs.^[9] Fortification of staple diet and supplementation of folic acid for the women facilitates closure of NTD early.^[2,3,14] On

This is an open-access article distributed under the terms of the Creative Commons Attribution-Non Commercial-Share Alike 4.0 License, which allows others to remix, tweak, and build upon the work non-commercially, as long as the author is credited and the new creations are licensed under the identical terms.

©2020 Published by Scientific Scholar on behalf of Surgical Neurology International

the contrary, excess intake of folate increases incidence of twin pregnancy and cancer as explained in systemic review done in Sweden.^[9] Fortified flour consumption lowers the occurrence of NTD as explored from the experimental study in Zhongyang county, serum folate level among intervention group is higher than in control group.^[2]

At population level, all women of child-bearing age who are capable of becoming pregnant should consume 400 (0.4 mg) g of folic acid daily to facilitate NTD closure as per the WHO recommendation, this might be not applicable at individual level.^[5]

Worldwide 2 billion people were deficient from micronutrients which hamper their physical growth and mental performance. The worst affected countries include Afghanistan, Pakistan, Cambodia, Ethiopia, and West and Central African nations.^[29]

METHOD

Google Scholar and PubMed were systematically searched to identify potential research articles concerning the Epidemiology and its determinant factors of Neural Tube defects. Additionally, by retrieving references from a list of eligible studies was conducted to be confident about the search strategy. We apply Boolean operator that is “AND”, “NOT” and “OR”. Through consideration of the Boolean operator we searched as follows: ((Neural tube defect OR Epidemiology of Neural tube defect) AND (Determinant Factors))

EPIDEMIOLOGY OF NTD

Birth defects means any type of anomaly which occurred congenitally, worldwide 3% of neonates suffered from this defect from all births or 1 in every 33 baby born which is a cause for perinatal and infant mortality in developing and developed countries.^[6]

NTDs cause death, paralysis, or life-long disability – but many consequences of it is preventable. Each year spinal bifida and anencephaly, the two most common forms of neural tube defects, occur in an estimated 3,00,000 newborns worldwide.^[8]

The overall burden of NTD based on live births was 1.67/1000 (IQR = 0.98–3.49) for total NTD burden according to a systemic review conducted elsewhere.^[10] NTD prevalence is higher in Argentina among all congenital anomalies (spinal bifida: 5%, anencephaly: 3%, and encephalocele: 1.4%)^[11] and 1.58/1000 births in bantia.^[12]

The prevalence of spinal bifida 4.5 in India,^[33] 15.1% in Zhengzhou City, China^[36] and a trend analysis done in China 2006–2015 showed that the prevalence of NTD was decreasing^[15] and similarly decreases in Kenya.^[16] The prevalence of NTD in Sudan was 2.8%.^[17] The prevalence of birth defect in central and northwest of Ethiopia is 30.8% and 22.5% in the Amhara region.^[18]

Another finding which is done in Mekelle Ayder Hospital Ethiopia identifies CNS malformation was the most common one with a prevalence of 1.5%;^[19] additionally, a retrospective finding which is done in Addis Ababa referral hospitals identified that spinal bifida and hydrocephalus are a main cause of childhood mortality in the area.^[7] A retrospective descriptive study conducted among Central and Northwest Ethiopia revealed that the prevalence of NTD is 30.8%.^[18]

Almost all states plan Sustainable Development Goal 3 that is building good health and well-being in all community, but such type of international consensus may be affected through unrecognizable hidden and preventable problems.^[20]

DETERMINANT FACTORS OF NTD

In general, the determinant factors of NTD were summarized as behavioral, nutrition-related, environmental, medical illness, and health service-related factors.

Women who have exposure to second-hand smoking have strong positive association to NTD.^[21] Taking foliate tablet increases serum level of folic acid.^[2,22] In contrary, taking excessive amount of folic acid hampers brain development and change child behavior.^[25] The use of trimethoprim, a folic acid antagonist, at the preconception period increased the risk of exencephaly in the mouse. A study conducted in primate animals (mouse) showed that; the use of trimethoprim, a folic acid antagonist, at the preconception period increased the risk of exencephaly.^[23]

Fever, hyperthermia in early pregnancy,^[4] certain medications (e.g., valproic acid), alcohol, tobacco, certain chronic diseases (e.g., diabetes and obesity), iodine, iron, and vitamin B12 deficiency have an association with NTDs.^[24,26,27]

People living near to industry, especially in coal mine, living near to main road, production of fruit, maternal occupational exposure, distance to factories, living in higher altitude, soil type, number of doctors, use of chemical fertilizers,^[28,30,31] per capita income, and living in faults were correlated with NTD as reported study conducted in China.^[32] A case–control study in Iran revealed that maternal history of abortion and obesity^[26] has a significant association with NTD.^[34] The occurrence of NTD affected by biological differences; high in female as compared to male.^[35]

In general, NTD has a high correlation with, prenatal screening, nutritional, environmental, occupational exposure, and genetic factors.^[37]

CONCLUSION

Birth defect is fatal which affects the new generation; specifically, NTD is the problem of middle- and low-income countries. It is a direct cause for neonatal and perinatal

mortality rate globally. Even if little factors identified, yet conducting experimental and clinical trial researches are a better approach to slow down the progress.

Declaration of patient consent

Patient's consent not required as there are no patient's in this study.

Financial support and sponsorship

Publication of this article was made possible by The James I. and Carolyn R. Ausman Educational Foundation.

Conflicts of interest

There are no conflicts of interest.

REFERENCES

- Abdullah NL, Gunasekaran R, Mohd-Zin SW, Lim BH, Maniam P, Mohd-Salleh AS, *et al.* Cranial neural tube defect after trimethoprim exposure. *BMC Res Notes* 2018;11:475.
- Allagh KP, Shamanna BR, Murthy GV, Ness AR, Doyle P, Neogi SB, *et al.* Birth prevalence of neural tube defects and orofacial clefts in india: A systematic review and meta-analysis. *PLoS One* 2015;10:e0118961.
- Barua S, Chadman KK, Kuizon S, Buenaventura D, Stapley NW, Ruocco F, *et al.* Increasing maternal or post-weaning folic acid alters gene expression and moderately changes behavior in the offspring. *PLoS One* 2014;9:e101674.
- Botto LD, Moore CA, Khoury MJ, Erickson JD. Neural-tube defects. *N Engl J Med* 1999;341:1509-19.
- Bourouba R, Houcher B, Akar N. Risk factors of neural tube defects: A reality of batna region in algeria. *Egypt J Med Hum Genet* 2017;19:225-9.
- Center for Disease Control. Global Situation of Birth Defects and Initiatives for Prevention. United States: Center for Disease Control; 2016.
- Christianson A, Howson CP, Modell B. March of Dimes. Global Report on Birth Defects the Hidden Toll of Dying and Disabled Children. White Plains, New York: March of Dimes Birth Defects Foundation; 2006.
- Desposito F, Cunniff C, Frias JL, Panny SR, Trotter TL, Wappne RS. Folic acid for the prevention of neural tube defects. *Am Acad Pediatr* 1999;104:325-7.
- Flores AL, Vellozzi C, Valencia D, Sniezek J. Global burden of neural tube defects, risk factors, and prevention. *Indian J Community Health* 2014;26:3-5.
- Githuku JN, Azofeifa A, Valencia D, Ao T, Hamner H, Amwayi S, *et al.* Assessing the prevalence of spina bifida and encephalocele in a kenyan hospital from 2005-2010: Implications for a neural tube defects surveillance system. *Pan Afr Med J* 2014;18:60.
- Golalipour MJ, Qorbani M, Mirfazeli A. Risk factors of neural tube defects in northern iran. *Iran Red Crescent Med J* 2014;16:e7940.
- Groisman B, Gili J, Giménez L, Poletta F, Bidondo MP, Barbero P, *et al.* Geographic clusters of congenital anomalies in argentina. *J Community Genet* 2017;8:1-7.
- Hoyt AT, Canfield MA, Romitti PA, Botto LD, Anderka MT, Krikov SV, *et al.* Associations between maternal periconceptional exposure to secondhand tobacco smoke and major birth defects. *Am J Obstet Gynecol* 2016;215:613.
- Kondo A, Kamihira O, Ozawa H. Neural tube defects: Prevalence, etiology and prevention. *Int J Urol* 2009;16:49-57.
- Langlois PH, Hoyt AT, Lupo PJ, Lawson CC, Waters MA, Desrosiers TA, *et al.* Maternal occupational exposure to polycyclic aromatic hydrocarbons and risk of neural tube defect-affected pregnancies. *Birth Defects Res A Clin Mol Teratol* 2012;94:693-700.
- Liao Y, Wang J, Li X, Guo Y, Zheng X. Identifying environmental risk factors for human neural tube defects before and after folic acid supplementation. *BMC Public Health* 2009;9:391.
- Liu J, Xie J, Li Z, Greene ND, Ren A. Sex differences in the prevalence of neural tube defects and preventive effects of folic acid (fa) supplementation among five counties in northern china: Results from a population-based birth defect surveillance programme. *BMJ Open* 2018;8:e022565.
- Lo A, Polšek D, Sidhu S. Estimating the burden of neural tube defects in low-and middle-income countries. *J Glob Health* 2014;4:010402.
- Makelarski JA. Selected Environmental Exposures and Risk of Neural Tube Defects. Iowa: University of Iowa; 2010.
- Mekonen HK, Nigatu B, Lamers WH. Birth weight by gestational age and congenital malformations in Northern Ethiopia. *BMC Pregnancy Childbirth* 2015;15:76.
- Meskelu M. Effects of maternal folic acid supplementation on the development of neural tube and cardiovascular system of the off springs in human and animal models. AAU Inst Repository; 2014. SDG Sustainable Development Goal; 2017. Available from: <http://www.un.org/sustainabledevelopment>.
- Molloy AM, Kirke PN, Troendle JE, Burke H, Sutton M, Brody LC, *et al.* Maternal Vitamin B12 status and risk of neural tube defects in a population with high neural tube defect prevalence and no folic acid fortification. *Pediatrics* 2009;123:917-23.
- Omer IM, Abdullah OM, Mohammed IN, Abbasher LA. Research: Prevalence of neural tube defects Khartoum, Sudan August 2014-July 2015. *BMC Res Notes* 2016;9:495.
- Rana M, Bisht SS, Rana AJ, Upadhyay J. Neural tube defects, its etiology: Environmental exposures and genes, possible risk factors. *J Pharm Sci Res* 2017;9:131-8.
- Salih MA, Murshid WR, Seidahmed MZ. Epidemiology, prenatal management, and prevention of neural tube defects. *Saudi Med J* 2014;35:S15-28.
- SDG Sustainable Development Goal; 2017. Available from: <http://www.un.org/sustainabledevelopment>.
- Sheet F. World Birth Defects Day 2016. Geneva: Neural Tube Defects, World Health Organization; 2016.
- Sorri G, Mesfin E. Patterns of neural tube defects at two teaching hospitals in addis ababa, Ethiopia a three years retrospective study. *Ethiop Med J* 2015;53:119-26.
- Swedish Council on Health Technology Assessment. Benefits and Risks of Fortifying Flour With Folic Acid to Reduce the

- Risk of Neural Tube Defects. Sweden: Swedish Council on Health Technology Assessment; 2007.
30. Talebian A, Soltani B, Sehat M, Zahedi A, Noorian A, Talebian M. Incidence and risk factors of neural tube defects in Kashan, central Iran. *Iran J Child Neurol* 2015;9:50-6.
 31. Taye M, Afework M, Fantaye W, Diro E, Worku A. Magnitude of birth defects in central and northwest ethiopia from 2010-2014: A descriptive retrospective study. *PLoS One* 2014;11:e0161998.
 32. Verity C, Firth H, French-Constant C. Congenital abnormalities of the central nervous system. *J Neurol Neurosurg Psychiatry* 2003;74:i3-8.
 33. Wang H, De Steur H, Chen G, Zhang X, Pei L, Gellynck X, Zheng X. Effectiveness of folic acid fortified flour for prevention of neural tube defects in a high risk region. *Nutrients* 2016;8:152.
 34. World Health Organization. *Bulletin of the World Health Organization*. Vol. 82. Geneva: World Health Organization; 2004.
 35. Wolff T. Folic acid supplementation for the prevention of neural tube defects: An update of the evidence for the U.S. Preventive services task force. *Ann Intern Med* 2009;150:632-9.
 36. Wu JW, Xing YR, Wen YB, Feng JJ, Wang XX, Li TF, *et al*. Prevalence of spina bifida occulta and its relationship with overactive bladder in middle-aged and elderly Chinese people. *Int Neurourol J* 2016;20:151-8.
 37. Zhang TN, Jiang CZ, Gong TT, Li J, Li LL, Chen Y, *et al*. Time trends in the prevalence and epidemiological characteristics of neural tube defects in Liaoning Province, China, 2006-2015: A population-based study. *Oncotarget* 2017;8:17092-104.

How to cite this article: Endalifer ML, Diress G. Epidemiology and determinant factors of neural tube defect: Narrative review. *Surg Neurol Int* 2020;11:81.