# **Preterm Birth, Exasperation to the South Asian Countries**

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# **ABSTRACT**

When a child is born before 37 weeks or 259 days of pregnancy, it is termed as preterm birth. Pre-term birth is prevalent in both developed and developing country. However, difference lies in their survival. In lower and middle income countries, most preterm babies die due to lack of even simple interventions. India ranks top in the world for deaths due to complications of preterm birth. Similarly, other South Asian countries, Pakistan, Bangladesh, Afghanistan and Nepal rank 3<sup>rd</sup>, 6<sup>th</sup>, 9<sup>th</sup> and 20th in the same. The aim of this review paper is to provide a landscape analysis on the burden of pre-term birth and challenges in the context of South Asian region. Databases such as PubMed, Scopus and Google Scholar were searched from 2000 to 2020 and 27 articles are included in the study. It was found that pre-term birth causes huge burden in the form of morbidity, mortality as well as socio economic losses. Preterm birth was associated with increased sepsis, cardiovascular disease, respiratory illness, hypothermia, neurological and gastrointestinal complications. South Asian countries have distinct challenges in eliminating or reducing preterm births which are: poor quality health surveillance data, inadequate trained health workforce, insufficient finance and funding, service delivery and other methodological challenges. Ending pre-term birth is important as it is directly related to Sustainable Development Goal 3. Therefore, there should be increase in priority given to increase financing, quality data gathering, adopting innovative measures as well as joint efforts of all the sectors to control the pre-term birth.

# **KEY WORDS**

Burden, Challenges, Pre-term birth, South Asian countries

# **INTRODUCTION**

A birth before 37 weeks of pregnancy or 259 days of gestation since the first day of last menstrual period of a mother has been defined as a 'Preterm Birth' by the World Health Organization. 1 Based on gestational age, it is further categorized as extremely preterm occurring before 28 weeks; very preterm occurring at 28 to 31 weeks; moderate occurring between 32 to 33 weeks and late preterm occurring at 34 to 37 weeks of gestation.<sup>2</sup> Basically, pre-term birth occurs either as a spontaneous pre-term birth or as a provider-initiated pre-term birth which is done for obstetric or fetal indication or other nonmedically indicated reasons. Spontaneous pre-term birth occurs due to various reasons such as mother's age (less than 20 or more than 35 years), poor maternal nutrition, few or no ante-natal care (ANC) visit, smoking and alcohol consuming habit of mother, maternal psychological ill health, severe anemia during pregnancy, polluted fuel, multiple pregnancies or deliveries, mother's infection or chronic conditions such as diabetes, high blood pressure, thyroid disease and Genetic influence.<sup>3,4</sup>

The global estimated incidence of preterm birth was reported to increase from 9.8 in 2000 to 10.6 in 2014.<sup>5</sup> According to Global Action Report on Preterm birth (2012), there is difference in rate of preterm birth among regions and countries where low-income countries have highest (11.8%) and high income countries have lowest rates (9.3%) comparatively.<sup>6</sup> Out of total preterm births, more than 80% occurred only in Asia and sub-Saharan Africa.<sup>5,6</sup> This high ratecould be possibly due to high fertility rate and high numbers of birth in these regions.<sup>6</sup> However it is difficult to conclude due to lack of quality and representative data in these regions.<sup>3</sup> The WHO report published in 2012

Country	Live births (2010)	Preterm birth rate (2010)	Rank for pre- term birth rate (2010)	Number of preterm births (2010)	Rank for num- ber of preterm births (2010)	Deaths from complications of preterm birth (2010)	Rank for deaths due to complica- tions of preterm birth (2010)
Afghanistan	1,385,200	11.5	66	159,900	18	21,200	9
Bangladesh	3,037,700	14.0	24	424,100	7	36,900	6
Bhutan	14,900	10.2	93	1,500	157	200	123
India	27,200,000	13.0	36	351,9100	1	303,600	1
Maldives	5,300	7.9	132	400	170	<50	168
Nepal	723,500	14.0	20	101,500	29	10,400	20
Pakistan	4,741,500	15.8	8	748,100	4	72,100	3
Sri Lanka	378,400	10.7	81	40,600	60	600	93

Source: Country data and rankings for preterm birth data. 10

showed that 1 million people die each year globally due to associated complications and many survivors bear lifelong problems and disabilities.<sup>3</sup> Although some high and middle income countries have preterm birth as the prime cause of child death, majority of child deaths due to pre-term birth occurs in lower or middle income settings like South Asia.<sup>3,6</sup> According to World Bank Report, most of the countries of South Asian Region lies in lower middle income economy.<sup>7</sup>

South Asia is one of the regions with highest stillbirth and neonatal deaths where one third of the neonatal deaths is attributed to pre-term birth.<sup>8</sup> According to UNICEF, about 1 million (39%) of the world's newborn deaths occurred in the eight South Asian countries in the same year.<sup>9</sup> India, Bangladesh, and Pakistan of South Asia comprise of 23.4%, 4% and 3% respectively of proportion of global preterm births; India lying on the top of the list as shown in table 1.<sup>10</sup>

Albeit preterm birth is prevalent in both developed and underdeveloped countries, the difference lies in their survival. More than 90% of preterm babies born before 28 weeks are able to survive in high income countries, with availability of intensive care, while most preterm babies born even beyond 32 weeks die in low income countries due to lack of even simple interventions such as skilled birth attendance, warmth, breastfeeding support, basic care for infections and breathing difficulties. 6,11 Studies have shown that preterm birth is not only associated with direct neonatal deaths, birth complications, intra-partum related complications neonatal infections, and congenital anomalies, further it has long term repercussions on survivors such as impaired physical and neurological development as well as on family, economic and societal effects.3 Studies have been done on the short term and long term complications of preterm birth in developed countries, but negligible follow up studies have been conducted on short term and long term sequelae of preterm birth in lower and middle income countries.5

The aim of this review paper is to provide a landscape analysis on the burden consequences and challenges of pre-term birth in the context of South Asian region.

#### **METHODS**

In this review, databases such as PubMed, Scopus and Google Scholar were searched for English-language papers from 2000 to September, 2020. A comprehensive search was performed using combinations of key words "preterm birth", "preterm delivery", "prematurity" "respiratory", "cardiovascular", "neurological", "gastrointestinal", "outcomes" "complications", "consequences" "South Asia". The searches were performed using Boolean operators OR, AND between the mentioned keywords Advanced search of search engine was applied. After a review of titles, abstract and full texts were reviewed consequently, 27 articles were included in the study.

# **BURDEN OF PRETERM BIRTH**

# Mortality

Preterm birth complications rank the top most position among causes of deaths of children under five years of age globally in 2016 accounting for 18% of under five years deaths and 35% of newborns deaths. <sup>13</sup> Infants born before 34 weeks of gestation are at 58 times increased risk for neonatal death than infants born at term. Meanwhile, infants born between 34 and 36 weeks of gestation are at 3.2 times increased for neonatal death than infants born at term. <sup>14</sup> Table 1 demonstrates deaths from complications of pre-term birth among South Asian countries.

### Morbidity

Pre-term birth is pre-disposing factor for a number diseases and conditions. Acute and chronic respiratory morbidities are common adverse outcome of preterm birth. In Nepal, birth asphyxia and severe respiratory distress syndrome were reported in 5% and 32% preterm babies respectively. Perinatal asphyxia accounted for 12.3% preterm deaths in India. In Pakistan 23.5% of late preterm presented with respiratory distress syndrome. A study done in Nepal showed sepsis in 37% preterm babies. Sepsis was reported in 50% of severe preterm, 48% of moderate preterm and 29% of late preterm.

9.2% of late preterm in Pakistan.<sup>18</sup> Likewise, preterm birth is found to be the risk factor for cardiovascular disease, hypertension, atrial fibrillation, early heart failure and cardiomyopathies.<sup>18</sup> The lower the gestational age, there is increase in ventricular mass predisposing heart failure.<sup>19</sup> Patent ductus arteriosus was reported in 20% of the severe preterm babies.<sup>15</sup> The restricted vascular bed, narrow and stiff arteries predisposes to arterial hypertension. Preterm infants are at increased risk of developing hypertension in later life.<sup>18</sup>

The major neurological outcomes of preterm birth include cerebral palsy, intraventricular hemorrhage (IVH), intellectual disability, hearing loss and visual impairment with retinopathy of prematurity. 20,21 IVH is reported in 1% of preterm babies and 3.5% of moderately preterm babies. 15 The study done in India showed IVH in 7% of preterm babies.<sup>22</sup> The cerebral palsy predispose behavioral and educational problems in children or adults born preterm.<sup>23</sup> In the same way, the risk of developing hypoglycemia as short term complication is higher in pre-terms. Hypoglycemic brain injury causes small head circumference and poor cognitive abilities. Children born preterm with hypoglycemic brain injury have lower intelligence quotient (IQ).<sup>24</sup> The study done in India showed hypoglycemia in 19.05% of preterm babies.<sup>25</sup> Similar study done in Pakistan reported hypoglycemia in 13.8% late preterm babies.<sup>17</sup> Necrotizing enterocolitis (NEC) is another gastrointestinal complications following preterm birth. It is reported in 4% of newborn babies in Nepal. NEC was developed in 10% of severe preterm, 7% of moderate preterm and 1.5% of late preterm babies. 15 Hypothermia, temperature below 36.5°C to 37.5°C is commonly seen as short term complication of preterm birth. Hypothermia is reported in 6% of late preterm babies.<sup>17</sup> Hypothermia resulted in 69% of preterm mortality causing dehydration, fluid electrolyte imbalance, hypotension, irritability and poor feeding in preterm babies.<sup>26</sup> A study done in Nepal showed that risk of hypothermia was increased in case of pre-term babies.<sup>27</sup>

### Socio-economic burden

Once a preterm is born, a huge investment is done by the family as well as a country to save the child. Similarly, if the child develops any physical or neurological impairment, it is socially challenging for s/he to adapt throughout his/her life.<sup>3</sup>

# Challenges in ending pre-term birth

Ending pre-term birth is important as it is directly related to SDG 3, target 3.2 in eliminating preventable cause of childhood deaths to at least as low as 12 per 1000 live births and under-5 mortality to at least as low as 25 per 1000 live births. Similarly, it is also important in order to achieve UNICEF South Asia target, of reducing newborn deaths from 28 in 2016 to 21 per 1,000 live births by 2021 which will ultimately save 500,000 newborns.<sup>9</sup>

However, there are multiple obstacles and challenges in eliminating or even reducing preterm birth in South Asia. As in most middle income setting, the problems lie in deriving health surveillance data. The most important problem with preterm is measurement in routine Health Information System (HIS).<sup>3</sup> Due to poor routine health surveillance data, it is difficult to estimate the exact numbers of preterm births in a country and the region as a whole. Likewise, other problems are associated with inadequate trained health workforce, insufficient finance and funding. Difficulty in delivering services is another challenge due to hurdle in implementing health care interventions in certain regions of South Asia. For example: Implementing Kangaroo Mother Care was difficult in Asian countries whereas it was easily accepted in African countries.

Similarly there are a set of methodological challenges that hinder in generating data and evidence to end preterm in this region.5 The WHO recommended and most accurate method for determining gestational age is ultrasound. However, due to constraint resources in South Asia, most commonly used method are less accurate ones; E.g.: last menstrual period, symphysis-fundal height measurement, postnatal examination of the baby, or use of birth weight. Similarly, due to non-inclusion of pre-term data in national civil registration and vital statistics provides difficulties in correct estimation of pre-term rates as well as variation in definition among the countries in a region does not provide accurate data. In South Asian region, there is mostly use of facility based (non-population representative) data and due to lack of population based data for estimation of pre-term rate tends to give inaccurate numbers. Lack of integration of local and national level bodies for data assimilation, monitoring and evaluation of pre-term birth and lack of high impact research in the regions are other methodological challenges in eliminating pre-term birth.

# CONCLUSION

Pre-term birth is not only a burden of an individual who is born too soon but is a shared problem as its repercussions are at all of individual, familial, societal, national, regional and global level. Apart from systemic complications and child mortality, there are other deep familial and socioeconomic burdens of pre-term birth. Therefore, priority must be given to eliminate the survival gap for pre-term babies in high incidence regions such as South Asian region and subsequently preventing the preterm birth all over the world. Firstly, we need quality data gathered using standard definition which could be base for research, policy making, planning and implementing programs and interventions to end pre-term birth. Further research is required on the immunological, behavioral and psychological consequences of preterm birth in developing countries. Research is needed to develop effective and affordable interventions to reduce the complications of preterm birth in developing countries. This evidenced-based information will help health care providers in understanding the complications of preterm birth and timely implementation of interventions to reduce complications related to preterm birth. There must be production of skilled work force to take care of pregnant mothers and small pre-term babies. In rural areas, health workers must be motivated with incentives, task sharing and performance based financial protection. There should be increase in priority given to increase financing, adopting

and using innovative measures related to pre-term birth. In doing so, everyone has a role in it; government and policymakers, donors, UN and multilateral organization, civil society, business community, health care workers and associations, academics and researchers as well as common people. Without every sector's efforts, it won't be possible to for us to achieve SDG target by 2030 and eventually we will be overburdened with pre-term birth which is going to cost us dearly.

# **REFERENCES**

- Dbstet A. WHO: recommended definitions, terminology and format for statistical tables related to the perinatal period and use of a new certificate for cause of perinatal deaths. *Acta Obstet Gynecol Scand*. 1977;56(3):247-53. Available from: https://pubmed.ncbi.nlm.nih. gov/560099/
- Moutquin JM. Classification and heterogeneity of preterm birth. BJOG: An International Journal of Obstetrics & Gynaecology. 2003 Apr;110:30-3. Available from: https://obgyn.onlinelibrary.wiley.com/doi/abs/10.1046/j.1471-0528.2003.00021.x
- Howson CP, Kinney MV, McDougall L, Lawn JE. Born too soon: preterm birth matters. Reproductive health. 2013 Nov;10(1):1-9. Available from: https://link.springer.com/article/10.1186/1742-4755-10-S1-S1
- Gurung A, Wrammert J, Sunny AK, Gurung R, Rana N, Basaula YN, et al. Incidence, risk factors and consequences of preterm birth– findings from a multi-centric observational study for 14 months in Nepal. Archives of Public Health. 2020 Dec;78(1):1-9. https://doi. org/10.1186/s13690-020-00446-7
- Chawanpaiboon S, Vogel JP, Moller A-B, Lumbiganon P, Petzold M, Hogan D, et al. Global, regional, and national estimates of levels of preterm birth in 2014: a systematic review and modelling analysis. The Lancet Global Health. 2019;7(1):e37-e46. Available from: https:// www.sciencedirect.com/science/article/pii/S2214109X18304510
- Walani SR. Global burden of preterm birth. *International Journal of Gynecology & Obstetrics*. 2020 Jul;150(1):31-3. Available from: https://doi.org/10.1002/ijgo.13195
- The World Bank. World Bank country and lending groups, country classification.(2021)Available from: https://datahelpdesk.worldbank. org/knowledgebase/articles/906519-world-bank-country-and-lending-groups
- McClure EM, Saleem S, Goudar SS, Dhaded S, Guruprasad G, Kumar Y, et al. The project to understand and research preterm pregnancy outcomes and stillbirths in South Asia (PURPOSe): a protocol of a prospective, cohort study of causes of mortality among preterm births and stillbirths. Reproductive health. 2018 Jun;15(1):45-53. Available from: https://doi.org/10.1186/s12978-018-0528-1
- The United Nations Children's Fund. Save newborn South Asia headline results 2018-2021. 2021. Available from: http://www. unicefrosa-progressreport.org/savenewborns.html
- World Health Organization. Born too Soon: The Global Action Report on Preterm Birth. 2010. Available from: https://www.who.int/pmnch/ media/news/2012/201204\_borntoosoon\_countryranking.pdf
- Blencowe H, Cousens S, Oestergaard MZ, Chou D, Moller AB, Narwal R, et al. National, regional, and worldwide estimates of preterm birth rates in the year 2010 with time trends since 1990 for selected countries: a systematic analysis and implications. The lancet. 2012;379(9832):2162-72. Available from: https://www.sciencedirect. com/science/article/abs/pii/S0140673612608204
- Saigal S, Doyle LW. An overview of mortality and sequelae of preterm birth from infancy to adulthood. *The Lancet*. 2008 Jan 19;371(9608):261-9. Available from: https://www.sciencedirect.com/ science/article/abs/pii/S0140673608601361

- Hug L, Sharrow D, You D. Levels and trends in child mortality: report 2017: The World Bank 2017. Available from: https://www.sciencedirect.com/science/article/pii/S2214109X19301639
- 14. Marchant T, Willey B, Katz J, Clarke S, Kariuki S, Ter Kuile F, et al. Neonatal mortality risk associated with preterm birth in East Africa, adjusted by weight for gestational age: individual participant level meta-analysis. *PLoS Med.* 2012;9(8):e1001292. Available from: https://journals.plos.org/plosmedicine/article?id=10.1371/journal. pmed.1001292
- Shrestha L, Shrestha P. Mortality and Morbidity Pattern of Preterm Babies at Tribhuvan University Teaching Hospital. *Journal of Nepal Paediatric Society.* 2013;33(3). Available from: https://www.researchgate.net/publication/272871254\_Mortality\_and\_Morbidity\_Pattern\_of\_Preterm\_Babies\_at\_a\_Tertiary\_Level\_Hospital\_in\_Nepal
- 16. Jain K, Sankar MJ, Nangia S, Ballambattu VB, Sundaram V, Ramji S, et al. Causes of death in preterm neonates (< 33 weeks) born in tertiary care hospitals in India: analysis of three large prospective multicentric cohorts. *Journal of Perinatology*. 2019;39(1):13-9. Available from: https://www.nature.com/articles/s41372-019-0471-1
- Khowaja WH, Leghari AL, Hussain AS, Ariff S, Khan IA. Frequency and early complications of late preterm infants: A descriptive analysis from two secondary-care hospitals of Karachi. *Cureus*. 2019;11(9).
  Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/ PMC6827863/
- Lewandowski AJ. The preterm heart: a unique cardiomyopathy?: Nature Publishing Group. 2019. Available from: https://www.nature.com/articles/s41390-019-0301-3
- 19. Carr H, Cnattingius S, Granath F, Ludvigsson JF, Bonamy A-KE. Preterm birth and risk of heart failure up to early adulthood. *Journal of the American College of Cardiology.* 2017;69(21):2634-42.Available from: https://www.jacc.org/doi/full/10.1016/j.jacc.2017.03.572
- Vinekar A, Dogra M, Azad RV, Gilbert C, Gopal L, Trese M. The changing scenario of retinopathy of prematurity in middle and low income countries: Unique solutions for unique problems. *Indian journal of* ophthalmology. 2019;67(6):717. Available from: https://www.ncbi. nlm.nih.gov/pmc/articles/PMC6552588/
- Olusanya BO. Screening for neonatal deafness in resource-poor countries: Challenges and solutions. Research and Reports in Neonatology. 2015;5:51-64. Available from: https://www.dovepress. com/screening-for-neonatal-deafness-in-resource-poor-countrieschallenges--peer-reviewed-fulltext-article-RRN
- 22. Mohandas Nair GS, Vijayakumar M, Anjana B. Pattern of growth and neurodevelopmental outcome of preterm babies born ≤ 34 weeks of gestation in a South Indian tertiary care hospital. *International Journal of Contemporary Pediatrics*. 2018;5(2):467. Available from: https://www.ijpediatrics.com/index.php/ijcp/article/view/1380
- Abas O, Abdelaziem F, Kilany A. Clinical spectrum of cerebral palsy and associated disability in South Egypt: A local survey study. *Open access Macedonian journal of medical sciences*. 2017;5(1):37. Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5320905/

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- 24. Sharma A, Davis A, Shekhawat PS. Hypoglycemia in the preterm neonate: etiopathogenesis, diagnosis, management and long-term outcomes. *Translational Pediatrics*. 2017;6(4):335. Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5682372/
- 25. Singh YP, Devi TR, Gangte D, Devi TI, Singh NN, Singh MA. Hypoglycemia in newborn in Manipur. *Journal of Medical Society*. 2014;28(2):108. Available from: https://www.jmedsoc.org/article.asp?issn=0972-4958;year=2014;volume=28;issue=2;spage=108;epage=111;aulast=Singh
- 26. Roychoudhury S, Yusuf K. Thermoregulation: advances in preterm infants. *Neo Reviews*. 2017;18(12):e692-e702. Available from: https://neoreviews.aappublications.org/content/18/12/e692
- 27. Mullany LC, Katz J, Khatry SK, Leclerq SC, Darmstadt GL, Tielsch JM. Neonatal hypothermia and associated risk factors among newborns of southern Nepal. *BMC Med.* 2010, 8: 43-10. Available from: https://bmcmedicine.biomedcentral.com/articles/10.1186/1741-7015-8-43