

"Comprehensive Analysis of Neonatal Jaundice: A Systematic Review"

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ABSTRACT

In the modern world, jaundice is a complex condition that affects everyone¹. Jaundice is really caused by hyperbilirubinemia, or high bilirubin levels, in the body of the new-born. Skin, mucous membranes, and a yellowing of the skin (Kernicterus) are common manifestations of jaundice². Jaundice typically emerges during the first week after delivery and disappears on its own after one to two weeks in both term and preterm infants⁴⁵. Jaundice brought on by bilirubin serum build up. The most frequent morbidity in the new-born period is elevated bilirubin, and 5–10% of all pregnancies necessitate treatment for pathological jaundice. Our comprehensive review's findings revealed that numerous therapies^{15,16}, including the administration of albumin, exchange transfusions, home and hospital phototherapy, immunoglobulin, and tin-mesoporphyrin, have been shown to be helpful and safe in treating jaundice³³.

Keywords: - Neonatal, Jaundice, Bilirubin, Hepatic, Elevated.

I. INTRODUCTION

The most prevalent ailment in new-borns is jaundice. About 10% of infants experience jaundice in the first month of life due to breastfeeding¹. Jaundice can also develop during the first week of a baby's life.²Yellow synthetic pigment called bilirubin has the potential to be hazardous, particularly for newborns³. There is no chemical conjugation of the bilirubin. The French term "jaune," which implies yellow, is where the word "jaundice" first appeared. Jaundice is a sign of elevated bilirubinaemia and suggests that the majority of the high levels of bilirubin may not be in chemically conjugated form.⁴ When the bilirubin level exceeds 2 mg/dL, jaundice will manifest Haem group-manufacturing substrate of bilirubin. It is a component of haemoglobin and haemolytic disease, with jaundice and neonates being the most common population affected. B. Inherited disease spherocytosis, deficiency of glucose-6-phosphate dehydrogenase. New-borns with jaundice

havewhite's eyes and yellow skin (usually the sclera) because bilirubin is deposited in the lining of the skin. Increased blood bilirubin⁷. Urine may be black and stool may be yellow⁸. Jaundice in new-borns is usually not a big problem and develops within the first week of life. Jaundice or persistently high levels of bilirubin can cause brain damage⁹.

On the basis of causes Jaundice can be classified into three types.¹⁰

1.1 Pre hepatic jaundice

Pre hepatic jaundice is a particular kind of jaundice that results from red blood cell destruction⁵.

Itisalsoassociatedwiththesamehemolyticjaundice.A majorfactorcontributingtoredbloodcelldegenerationi sweakredbloodcellmembranes.Degradationofredblood cellsoccursbecausecriticalcellmembranescannot withstandthestress,alsoduetoanincreaseinbilirubin⁵⁰.

1.2 Hepatic jaundice

The primary cause of jaundice is a hepatocyte abnormality in the liver. The liver separates albumin from bilirubin before excreting it through the bile duct. Hepatic jaundice might develop from conjugation and discharge^{17,41}. The main enzyme in conjugation is UDP-glucuronyl transferase. Paediatric Physiology This enzyme causes jaundice, which is brought on by an immature new born, and may lead to a genetic mutation of the UTG1A gene²².

Post hepatic jaundice

The causes of this type of jaundice are fat accumulation and diarrhoea. Post-hepatic obstructive jaundice may be treated with mechanical decompression, but the newborn will also experience difficulties and other symptoms⁴⁶.

1.3 Symptoms

The signs of jaundice include pale-colored faeces, dark urine, newborn white eyes, yellow skin, stomach discomfort, nausea, fatigue, lethargy,

altered muscle tone, loud crying, poor feeding, and convulsions²⁵. The signs of jaundice need to be taken carefully.;¹¹¹⁹ take your kid as soon as you

can to the emergency department or a walk-in clinic²⁰

Type	Symptoms
BREASTFEEDING	Occurs after one day of infancy reduced frequency of breastfeeding There may also be liver problems Not breastfeeding well
PHYSIOLOGIC	<ul style="list-style-type: none"> • Usually occurs in the first week of labour. • Disappears within the first week after birth • may not require treatment
PATHOLOGIC	<ul style="list-style-type: none"> • Jaundice due to a pathological process appears within 24 hours of birth. • Characterized by a rapid rise in bilirubin and/or elevated direct bilirubin levels. • Factors: A) Polycythaemia, sepsis, bruising, Superproduction. B) Intestinal obstruction, poor diet, acidosis, decreased excretion.
BREAST MILK	<ul style="list-style-type: none"> • After 5th day of life, more uncommon • It peaks in the second or third week and lasts for several weeks. • Due to the increased reabsorption of unconjugated bilirubin, there can be no effect on breast milk.

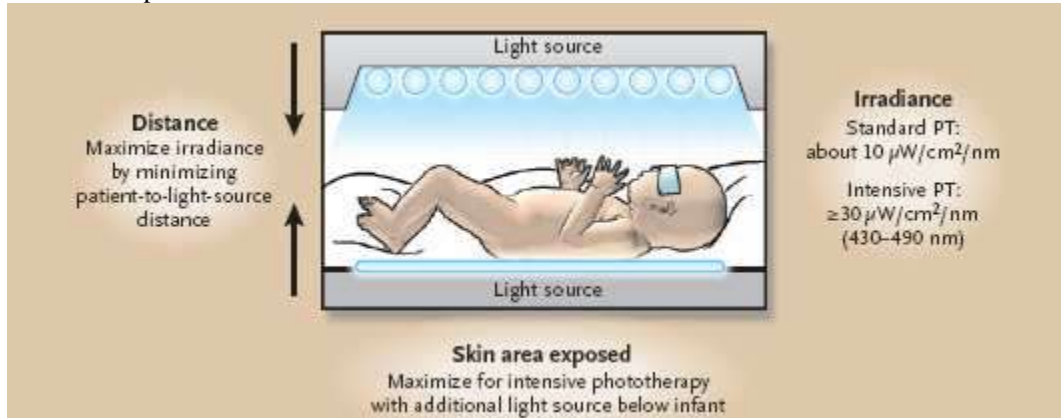
- 2 Causes of pathological jaundice
1. The following are typical causes of jaundice:
2. Hemolysis:
3. This condition is caused by incompatible blood types, such as ABO,²⁰ Rh, and small blood groups, as well as enzyme deficiencies, including 9G6PD deficiency and autoimmune haemolytic anemia.¹⁷
4. Reduced conjugation, similar to immaturity.
5. Increased hepatic circulation, such as inadequate enteral nutrition,²¹ such as inadequate nursing care, or infants not being fed due to gastrointestinal disease or obstruction⁵

6. Leakage of blood:
7. cephalohematoma, severe bruising, etc.
- 3 Treatment

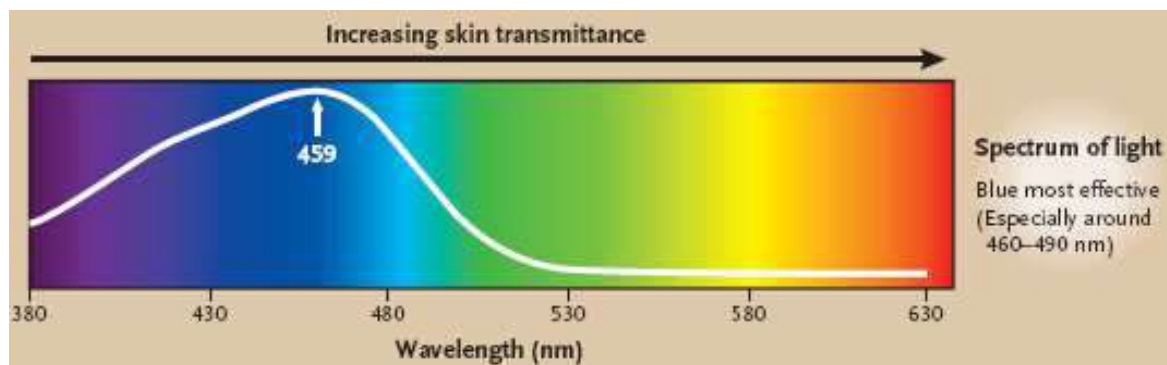
3.1 Phototherapy Principles

phototherapy is composed of radiation,²⁵ a certain light wavelength, and area of exposed skin²⁸. At a depth of about 2 to 3 mm²⁴, phototherapy affects bilirubin that is not chemically conjugated.²³ Without actually conjugating, photo isomerization transforms fat-soluble molecules into water-soluble ones,²⁹ which are then eliminated by the liver.

3.1.1 Structural photoisomerization



Source:www.google.com



Source:www.wikipedia.com

The degradation of bilirubin is largely impacted by blue-green emission light in a relatively small wavelength band (425 – 490 nm).^{35,8}The new birth should maximise the amount of skin exposed to light as feasible when phototherapy is used to lower the level of bilirubin³¹. Using several phototherapy sources might result in a more successful course of treatment⁷. According to medical standards³², the strength of phototherapy is specified in terms of light's wavelength, or nanometers²⁷³³.

3.1.2 Phototherapy Equipment

There are several ways to provide phototherapy³⁴. The approach is determined by the equipment that is available in each institution³⁵. The presently suggested techniques are as follows²⁹³⁶:

1. Phototherapy Lights: The smallest light spectrum is used for maximum bilirubin reduction. The effectiveness depends on the distance to the child, measured with a specific light meter⁵¹.
2. Bili Bassinet: This self-contained appliance combines three phototherapies with a mattress area³⁸. Phototherapy light units can be used in conjunction with phototherapy lamps, but they shouldn't be the only form of treatment³⁹.
3. Bili Blanket: This tiny fibre-optic pad is put beneath the baby³⁷⁴⁰. It is used in conjunction with above phototherapy lamps¹², not as a replacement for them³⁶. The Bili blanket also has the advantage of allowing for uninterrupted treatment during nursing³¹.

3.1.3 Potential Side Effects of Phototherapy

SIDE EFFECT	IMPLICATION	SYMPTOMS	ACTION
Ocular Effects	<ul style="list-style-type: none"> • Eye patches increase risk of eye infections, corneal abrasions, increased intracranial pressure⁴ (if applied too tight) 	<ul style="list-style-type: none"> • Lack of sensory input and stimulation • Use of eye patches for prolonged period 	<ul style="list-style-type: none"> • Remove eye patches often to encourage contact and stimulation, and keep a look out for discomfort or infection indicators¹⁰. • Guarantee eye patches are set up when lights are utilized
Altered Fluid Status	<ul style="list-style-type: none"> • Increased fluid loss may alter uptake of I.M. medications Because of the increase Water loss through evaporation, metabolic rate and possibly Respiratory rate³⁷ 	<ul style="list-style-type: none"> • Increased peripheral blood flow • Expanded torpid water misfortune with open bed or hotter⁴⁸ 	<ul style="list-style-type: none"> • Monitor weight and fluid intake/output
Skin Changes	<ul style="list-style-type: none"> • Due to melanin synthesis • Due to injury to skin mast cells with release of histamine or erythema¹⁶ • From exposure to shortwave emissions from⁵⁰ fluorescent light • Due to decreased hepatic excretion of bilirubin photodegradation by-products 	<ul style="list-style-type: none"> • Tanning • Rashes • Burns • Bronze Baby Syndrome 	<ul style="list-style-type: none"> • Monitor skin condition at least every 4 hours and report changes to team
Altered Activity	<ul style="list-style-type: none"> • May impact parent-infant interaction • May alter fluid and caloric intake 	<ul style="list-style-type: none"> • Lethargy or irritability • Decreased eagerness to feed 	<ul style="list-style-type: none"> • Observe and support parental efforts and concerns
Altered Thermal / Metabolic Function	<ul style="list-style-type: none"> • Influenced by maturity, caloric intake, heat dissipation from 	<ul style="list-style-type: none"> • Increased environmental and body temperature changes • Expanded O₂ 	<ul style="list-style-type: none"> • Monitor temperature, distance from lights

	phototherapy machine, airflow, distance from infant to light.	utilization •Expanded respiratory rate •Expanded skin blood stream	• Ensure adequate food/fluid intake
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3.2 Exchange transfusion

If the new-born exhibits indications of bilirubin encephalopathy regardless⁴¹ of TSB levels or the TSB levels outperform the age-explicit cut-off for trade bonding,⁴² a double volume exchange transfusion (DVET) should be done²⁵.

In new-borns with Rh isoimmunization, there are signs of DVET at birth that include²³:

1. Cord bilirubin is at least 5 mg/dL²¹.
2. If a new-born exhibits signs of hydrops⁴³ or cardiac decompensation while having low PCV (35%), fractional trade bonding with 50 mL/kg of stuffed cells ought to be performed following birth to quickly re-establish the blood's oxygen conveying capacity. Pull and push method employing the umbilical venous channel should be used to execute the ET⁴⁴. Only enough umbilical catheter should be placed to allow for blood flow²⁹.

Table no. 3 :-Type and volume of blood for exchange transfusion.

S. No.	Condition	Type of blood
1	Rh isoimmunization	I am Rh negative and my blood type is "O" or my baby's blood type. Suspended in AB plasma. Cross compatibility between the baby's blood and the mother's ²²
2	ABO incompatibility	Blood type "O" (non-infant) suspended in AB plasma and matched to mother's and infant's blood ³³
3	Other conditions (G6PD deficiency, non-haemolytic, other isoimmunised haemolytic jaundice)	Child's gathering and Rh type Cross coordinated with child's and mother's blood ⁴⁵ .

Blood volume: Twice the baby's blood volume (total volume: 160 to 180 ml/kg)
 Mix one third of plasma with two thirds of packed cells to make blood for DVET preparation³⁴.

3.3 Intravenous immunoglobulins (IVIG)

In cases of isoimmunised haemolytic anaemia (Rh isoimmunization and ABO incompatibility)¹⁰, IVIG lessens haemolysis and the development of jaundice³, reducing the requirement for phototherapy and exchange transfusions.⁴⁷ In all cases of Rh isoimmunization and in some cases of ABO inconsistency with extreme hemolysis,⁴ IVIG (0.5 to 1 gm/kg) is administered²⁸. Necrotizing enterocolitis and intestinal damage can result with IVIG therapy.¹⁶

3.4 IV hydration

IV hydration should be administered to infants who have severe hyperbilirubinemia⁴⁸ and signs of dehydration, such as rapid weight loss¹¹. The demand for exchange transfusion is reduced by

an additional 50 mL/kg of N/3 saline over the course of eight hours²⁷.

3.5 Other agents

Drugs like phenobarbitone, clofibrate, or steroids should not be used in the treatment of jaundiced new-borns⁴⁹ because there is no evidence to support their use in preventing or treating hyperbilirubinemia in neonates^{50,51}.

II. CONCLUSION

Babies between 1-3 days old account for the majority of new-born jaundice cases. More than 2 mg/dl of bilirubin is not acceptable. Neonatal jaundice frequently results from ABO incompatibility, Rh incompatibility, and cephalhematoma. Common signs include yellow

skin, light faeces, stomach pain, and poor appetite are seen in our study. We also looked into the main categories of jaundice. Although phototherapy is the primary pharmacological treatment for newborn jaundice, pharmacological management treatments such as IV immunoglobulins, IV hydration, and exchange transfusion are carried out in severe situations.

REFERENCE: -

- [1]. Jaundice in the Newborn. 2001;68(10):977-980. doi:10.1007/BF02722600
- [2]. Alloimmunization in thalassemia patients: New insight for healthcare. 2017;8:1-6. doi:10.4103/ijpvm.IJPVM
- [3]. Resource Guide for Nurses: Patient Teaching on Newborn Jaundice (Hyperbilirubinemia) Evaluation: Published online 2017:3-5.
- [4]. Study of PCV levels in neonatal jaundice. 2017;3(September):31-34.
- [5]. Maternal and Neonatal Determinants of Neonatal Jaundice – A Case Control Study. 2017;05(03):19659-19665. doi:10.18535/jmscr/v5i3.210
- [6]. Newborn Jaundice. Published online 2021:51-52. doi:10.1542/9781581106640-part02-ch14
- [7]. Association between neonatal jaundice and autism spectrum disorders among children: A meta-analysis. 2020;63(1):8-13. doi:10.3345/kjp.2019.00815
- [8]. Neonatal Hyperbilirubinemia in Low-Income African Countries. 2021;7(1). doi:10.23937/2469-5769/1510073
- [9]. Jaundice in newborns. 1999;4(2):165-166. doi:10.1093/pch/4.2.165
- [10]. Jaundice and Your Newborn. 2022;(October):1-2. doi:10.1542/peo_document197
- [11]. Sex-and age-related differences in bilirubin concentrations and severity of jaundice. 2021;5(November 2020):743-746. doi:10.24911/ijmdc.51-1606747050
- [12]. Target Product Profile: Phototherapy Light - Jaundice Managemet. 2020;(March):1-21.
- [13]. The pathophysiology of neonatal jaundice in urosepsis is complex with mixed bilirubin !!! Published online 2022:68-70. doi:10.15406/jpnc.2022.12.00458
- [14]. Neonatal-Jaundice Identification and Management in Neonates \geq 32 Weeks Gestation File number H15/89163-2 Functional group Clinical/Patient Services-Maternity. Published online 2016.
- [15]. Review Article NEONATAL JAUNDICE - A REVIEW. 2016;5(4):2198-2200.
- [16]. Prevalence and Etiology of Neonatal Jaundice in a Tertiary Care Hospital. 2021;18(2):35-38. doi:10.3126/jngmc.v18i2.38891
- [17]. Hypertension o ov verview. 2019;(March 2018):1-12.
- [18]. Risk factors for neonatal hyperbilirubinemia: a case control study. 2016;6(1):198. doi:10.18203/2320-1770.ijrcog20164657
- [19]. Maternity and Neonatal Clinical Guideline Neonatal stabilisation for retrieval. Published online 2018. www.health.qld.gov.au/qcg
- [20]. Neonatal jaundice. 1947;67(4):104-106.
- [21]. Standard Treatment Guidelines 2022. Published online 2023.
- [22]. Section 5 Gastrointestinal and hepatobiliary.
- [23]. Neonatal jaundice - NICE guidelines. 2005;20(1):47-54.
- [24]. Neonatal Jaundice – Optimizing Phototherapy An initiative of ETAT + Trainers.
- [25]. Recommendations on newborn health: approved by the WHO Guidelines Review Committee. 2017;(May):1-28. https://www.who.int/maternal_child_adolescent/documents/newborn-health-recommendations/en/%0Ahttp://apps.who.int/
- [26]. Document control. 2003;36(6):104. doi:10.4324/9780080958033-21
- [27]. Prevalence and Contributing Factors of Neonatal Jaundice in Neonatal Intensive Care Unit at St Paul's Hospital Millennium Medical College, Addis Ababa, Ethiopia, 2019. 2020;74:2016-2021. doi:10.7176/jhmn/74-03
- [28]. Clinical profile of pathological Jaundice among neonates admitted in the National Referral Hospital, Bhutan. 2021;7(2):13-18. doi:10.47811/bhj.124
- [29]. The epidemiologic study of neonatal jaundice, relation between jaundice and liver and alternative methods to cure jaundice. 2019;16(3):1117-1125. doi:10.37532/fmcp.2019.16(3).1117-1125

- [30]. iMedPub Journals Short note on Neonatal jaundice. Published online 2020:36648. doi:10.36648/2386-5180.8.4.328
- [31]. Clinical Practice Guidelines Screening , Prevention and Management of Neonatal Hyperbilirubinemia National Neonatology Forum , India. 2020;(January).
- [32]. Phototherapy in Neonatal Hyperbilirubinaemia - An Overview. 2021;10(21):1621-1627. doi:10.14260/jemds/2021/337
- [33]. Neonatal jaundice. 2014;(1):1-17.
- [34]. Jaundice in the new-born baby. 1949;162(972):522.
- [35]. Neonatal Hyperbilirubinemia - Early Detection with Umbilical Cord Blood Bilirubin and Umbilical Cord Blood Alkaline Phosphatase Levels - Individual vs Combined Predictive Markers. 2020;05(01):32-38. doi:10.36348/sjm.2020.v05i01.007
- [36]. Management of Hyperbilirubinemia in Newborn Infants 35 or More Weeks of Gestation: American Academy of Pediatrics, 2022. 2023;60(1):63-66.
- [37]. Determinants of neonatal jaundice among neonates admitted to five referral hospitals in Amhara region, Northern Ethiopia: An unmatched case-control study. 2020;4(1):1-9. doi:10.1136/bmjpo-2020-000830
- [38]. Standardization of phototherapy for neonatal hyperbilirubinemia using multiple-wavelength irradiance integration. 2020;61(1):100-105. doi:10.1016/j.pedneo.2019.07.002
- [39]. INFORMATION FOR PARENTS This information is available in other formats on request . Please ask your nursing staff to arrange this.
- [40]. Knowledge, attitudes and practices regarding neonatal jaundice among caregivers in a tertiary health facility in Ghana. 2021;16(6 June):1-20. doi:10.1371/journal.pone.0251846
- [41]. Frequency of Exchange Transfusion in Newborns with Neonatal Hyperbilirubinemia. 2019;8(8):9-13. <https://www.ijmrhs.com/medical-research/frequency-of-exchange-transfusion-in-newborns-with-neonatal-hyperbilirubinemia.pdf>
- [42]. Hyperbilirubinemia in Neonatal Intensive Care Unit: Incidence And Etiology at Fayoum University Hospital. 2019;3(2):8-14. doi:10.21608/fumj.2019.60476
- [43]. The epidemiology of neonatal jaundice. 2021;4:1-14. doi:10.21037/pm-21-4
- [44]. Neonatal Jaundice Risk Factors at a District Hospital in Rwanda. 2020;3(2):204-213. doi:10.4314/rjmhs.v3i2.10
- [45]. Neonatal jaundice causes and management. 2018;5(11):4992. doi:10.18203/2394-6040.ijcmph20184604
- [46]. Child and Adolescent Health Service Phototherapy Phototherapy Treatment Threshold Graphs Click here to access Threshold Graphs Phototherapy Units. :1-6.
- [47]. Section : Paediatrics Clinical Severity of Neonatal Jaundice Especially in ABO and Rh Incompatibility in Northern India. 2020;(6):7-10.
- [48]. Did you know ? Neonatal Jaundice. Published online 2018. <https://resolution.nhs.uk/wp-content/uploads/2018/10/Did-you-know-Neonatal-Jaundice.pdf>
- [49]. Risk factors associated with neonatal jaundice: a cross-sectional study from iran. 2018;6(8):1387-1393. doi:10.3889/oamjms.2018.319
- [50]. Jaundice: Newborn to age 2 months. 2017;38(11):499-510. doi:10.1542/pir.2015-0132
- [51]. Turkish neonatal society guideline to the approach, follow-up, and treatment of neonatal jaundice. 2018;53(4):S172-S179. doi:10.5152/TurkPediatriArs.2018.01816