

# Neonatal IMCI Evidence-based Interventions

IN THE CONTEXT OF THE CARE CONTINUUM FOR MOTHERS, NEWBORNS AND INFANTS



**Pan American  
Health  
Organization**



Regional Office of the  
World Health Organization



**Texas Children's Hospital®**



# Neonatal IMCI Evidence-based Interventions

IN THE CONTEXT OF THE CARE CONTINUUM FOR MOTHERS, NEWBORNS AND INFANTS

Also published in Spanish as:  
AEIPI Neonatal: Intervenciones basadas en evidencia. Segunda edición.  
© Organización Panamericana de la Salud, 2010  
ISBN 978-927533135-4

**PAHO HQ Library Cataloguing-in-Publication Data**

Pan American Health Organization  
Neonatal IMCI Evidence-based Interventions  
Washington, D.C.: PAHO, © 2011

ISBN: 978-92-75-11630-2

I. Title

1. INFANT CARE - standards
2. CHILD HEALTH SERVICES
3. PERINATAL CARE - standards
4. MATERNAL AND CHILD HEALTH
5. PRENATAL NUTRITION
6. EVIDENCE-BASED MEDICINE - methods
7. COMPREHENSIVE HEALTH CARE

NLM WS 420

The Pan American Health Organization welcomes requests for permission to reproduce or translate its publications, in part or in full. Applications and inquiries should be addressed to Editorial Services, Area of Knowledge Management and Communications (KMC), Pan American Health Organization, Washington, D.C., U.S.A. The Family and Community Health Area/Healthy Life Course Project will be glad to provide the latest information on any changes made to the text, plans for new editions, and reprints and translations already available.

© Pan American Health Organization, 2011. All rights reserved.

Publications of the Pan American Health Organization enjoy copyright protection in accordance with the provisions of Protocol 2 of the Universal Copyright Convention. All rights are reserved.

The designations employed and the presentation of the material in this publication do not imply the expression of any opinion whatsoever on the part of the Secretariat of the Pan American Health Organization concerning the status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries.

The mention of specific companies or of certain manufacturers' products does not imply that they are endorsed or recommended by the Pan American Health Organization in preference to others of a similar nature that are not mentioned. Errors and omissions excepted, the names of proprietary products are distinguished by initial capital letters.

All reasonable precautions have been taken by the Pan American Health Organization to verify the information contained in this publication. However, the published material is being distributed without warranty of any kind, either expressed or implied. The responsibility for the interpretation and use of the material lies with the reader. In no event shall the Pan American Health Organization be liable for damages arising from its use.

# Neonatal IMCI Evidence-based Interventions

IN THE CONTEXT OF THE CARE CONTINUUM FOR MOTHERS, NEWBORNS AND INFANTS



**Pan American  
Health  
Organization**



Regional Office of the  
World Health Organization



**Texas Children's Hospital**®



Family and Community Health Area  
Healthy Life Course Project

Washington DC, 2011

In 2000, the neonatal component began to be incorporated in the IMCI. Generic materials were developed, and adaptations were made in more than 12 of the region's countries.



# Contents

<b>Foreword .....</b>	<b>xi</b>
<b>Preface .....</b>	<b>xiii</b>
<b>Introduction .....</b>	<b>1</b>
Quality of care in neonatology .....	1
Evidence-based medicine .....	2
Evidence-based medicine in perinatology and neonatology .....	2
<b>I. Prepregnancy interventions .....</b>	<b>3</b>
1. Using body mass index (BMI) to monitor nutritional status prior to pregnancy ....	3
2. Folic acid (FA) and multivitamin (MV) supplementation .....	4
3. Preventing and treating anemia with iron .....	5
4. Micronutrients and reproductive health in women .....	5
5. Detecting and treating sexually transmitted infections (STIs) before pregnancy	6
6. Deparasitization in areas of high prevalence .....	6
7. Detecting and treating chronic diseases (cardiovascular, nutritional, endocrine)	6
8. Detecting, preventing, and managing domestic violence .....	7
9. Detecting, preventing, and managing alcohol and tobacco consumption .....	7
10. Detecting, preventing, and managing depression .....	7
11. Detecting and preventing cervical cancer .....	7
12. Complete immunization series .....	8
13. Detecting and treating periodontal disease .....	8
14. Preventing pregnancy in adolescents .....	8
<b>II. Interventions during pregnancy .....</b>	<b>11</b>
1. Prenatal check-up .....	11
2. Preventing fetal death .....	12
3. Preventing preterm birth .....	12
4. Managing premature rupture of membranes .....	15
5. Using the CLAP/WR perinatal clinical record (PCR) and perinatal card .....	16
6. Weight gain during pregnancy .....	16
7. Reducing unnecessary (medically not indicated) caesarean section .....	18
8. Immunization with tetanus toxoid, taking previous vaccination status into account .....	19
9. Preventing influenza with vaccination .....	19
10. Syphilis screening and case management .....	20
11. Detecting and treating symptomatic and asymptomatic bacteriuria .....	21
12. Preventing and treating anemia with iron .....	21
13. Preventing preeclampsia and eclampsia with calcium (in populations with diets low in calcium) .....	21
14. Preventing preeclampsia and eclampsia with aspirin .....	22
15. Detecting and treating sexually transmitted infections .....	23
16. Antiretrovirals and other measures to reduce the risk of vertical transmission of HIV .....	23
17. Detecting and treating diabetes .....	23
18. Deparasitization in areas of high prevalence using albendazole .....	24
19. Detecting and treating group B streptococcus .....	24
20. Nifedipine during preterm labor .....	24
21. Prenatal corticosteroids (betamethasone, dexamethasone) to foster	

lung maturation .....	24
22. Detecting, preventing, and treating Rh isoimmunization .....	25
23. Detecting and treating periodontal disease .....	26
24. Detecting, preventing, and managing domestic violence and abuse, and depression.....	26
25. Detecting and preventing exposure to tobacco .....	26
26. Detecting and preventing exposure to alcohol .....	27
27. Detecting and preventing exposure to drugs .....	27
<b>III. Interventions during delivery .....</b>	<b>29</b>
1. Clean and safe delivery practices (with skilled personnel) .....	29
2. Accompanying the mother during labor and delivery .....	29
3. Erythromycin in PROM before delivery in first 37 weeks of gestation .....	29
4. Monitoring labor with partograms .....	30
5. Caesarean section in breech presentation with LBW infants .....	30
6. Antibiotic prophylaxis in caesarean section .....	30
7. Modified active management of the third stage of labor .....	30
8. Late clamping of the umbilical cord .....	30
<b>IV. Maternal interventions after delivery .....</b>	<b>33</b>
1. Early skin-to-skin contact .....	33
2. Mother-newborn rooming-in .....	33
3. Detecting and managing puerperal endometritis .....	33
4. Detecting, preventing, and managing postpartum depression .....	34
<b>V. Neonatal interventions after delivery .....</b>	<b>35</b>
1. Neonatal resuscitation with ambient air .....	35
2. Immediate care for the healthy newborn .....	36
3. Care of the umbilical cord .....	36
4. Skin care at birth .....	36
5. Breastfeeding in the first hour .....	37
6. Kangaroo mother program (skin-to-skin contact) .....	37
7. Stabilizing and transporting the neonate .....	38
8. Treating neonatal sepsis with a single daily dose of gentamicin .....	38
9. Treating respiratory distress syndrome (RDS) with natural surfactant .....	38
10. Neonatal screening to detect congenital hypothyroidism, galactosemia, phenylketonuria, and cystic fibrosis .....	39
11. Detecting and preventing tuberculosis and hepatitis B .....	40
12. Detecting and treating retinopathy of prematurity in the newborn (ROP) .....	40
13. Early home visits for newborn care .....	41
14. Supine sleeping position and sudden infant death syndrome (SIDS) .....	41
15. Exclusive breastfeeding .....	41
<b>VI. Summary of interventions and their management .....</b>	<b>43</b>
<b>VII. Bibliographic references .....</b>	<b>52</b>
<b>VIII. Annexes .....</b>	<b>81</b>

# Acronyms

<b>BMI</b>	Body Mass Index
<b>BMT</b>	Betamethasone
<b>CC</b>	Cervical cancer
<b>CDC</b>	Centers for Disease Control (Atlanta, USA).
<b>CLAP/WR</b>	Latin American Center for Perinatology/Women and Reproductive Health
<b>DMT</b>	Dexamethasone
<b>GA</b>	Gestational Age
<b>HIV</b>	Human immunodeficiency virus
<b>HPV</b>	Human papillomavirus
<b>IM</b>	Intramuscular
<b>IMCI</b>	Integrated Management of Childhood Illness
<b>IOM</b>	Institute of Medicine
<b>IUGR</b>	Intrauterine growth retardation/restriction
<b>IV</b>	Intravenous
<b>LAC</b>	Latin America and the Caribbean
<b>LBW</b>	Low birthweight
<b>LGA</b>	Large for gestational age
<b>SGA</b>	Small for gestational age
<b>PAHO</b>	Pan American Health Organization
<b>PCR</b>	Perinatal clinical record, developed by CLAP/WR
<b>PPROM</b>	Preterm premature rupture of membranes
<b>PROM</b>	Premature rupture of membranes
<b>RDS</b>	Respiratory distress syndrome
<b>ROP</b>	Retinopathy of prematurity
<b>RPR</b>	Rapid plasma reagins
<b>STI</b>	Sexually transmitted infection
<b>VDRL test</b>	Venereal Disease Research Laboratory test
<b>WHO</b>	World Health Organization
<b>ZDV</b>	Zidovudine

Neonatal mortality is the basic indicator used to measure the development of newborns and the quality of care that they receive in a particular geographical area or set of health care facilities.



# Technical group responsible for preparing this document

## **Dr. Yehuda Benguigui**

Senior Adviser  
Healthy Life Course Project  
Pan American Health Organization/World Health Organization

## **Dr. Gerardo Cabrera-Meza**

Pediatrician, Neonatologist  
Texas Children's Hospital  
Baylor College of Medicine

## **Dr. Martha Beltrán**

Pediatrician, Intensive Care Specialist  
Clínica del Country  
Bogotá, Colombia

## **Dr. Miguel Dávila D.**

Pediatrician  
PAHO/WHO consultant  
Lima, Peru

## **Dr. Francisco Martínez-Guillén**

Pediatrician, Neonatologist  
PAHO/WHO Regional Consultant  
Managua, Nicaragua

## **Dr. Pedro Marte**

Perinatologist, Neonatologist  
Altagracia Hospital  
Santo Domingo, Dominican Republic

## **Dr. Rolando Cerezo Mulet**

Pediatrician, Neonatologist  
PAHO/WHO consultant  
INCAP, Guatemala

# Technical review group

## **Dr. Ricardo Fescina**

Director  
Latin American Center for Perinatology/  
Women and Reproductive Health  
(CLAP/WR-PAHO/WHO)  
Montevideo, Uruguay

## **Dr. Fernando Amado**

PAHO/WHO consultant  
Guatemala

## **Dr. Yolanda Cifuentes Cifuentes**

Division of Perinatology and Neonatology,  
Department of Pediatrics. Medical School,  
National University of Colombia  
Bogotá, Colombia

## **Dr. Francisco Chew Beb**

Pediatrician, Gastroenterologist  
Guatemala, Guatemala

## **Dr. Dilberth Cordero Valdivia**

Pediatrician, Ministry of Health  
La Paz, Bolivia

## **Dr. Bremen De Mucio**

Regional Adviser on Reproductive Health  
(CLAP/WR) PAHO/WHO  
Montevideo, Uruguay

## **Dr. José Luis Díaz Rossello**

(CLAP/WR) PAHO/WHO  
Montevideo, Uruguay

## **Dr. Pablo Durán**

Neonatology Adviser  
(CLAP/WR-PAHO/WHO)  
Montevideo, Uruguay

## **Dr. Daniel Frade**

National Consultant  
Sexual, Reproductive, and Maternal Health  
PAHO/WHO, Guatemala

## **Dr. Clara Gálviz**

University Educator  
President, Colombian Neonatology  
Association  
Colombia

## **Dr. Miguel Angel Hinojosa**

Pediatrician, Neonatologist  
Neonatology Unit, Dr. Enrique Garcés  
Hospital  
Quito, Ecuador

## **Dr. Emilton López Pimentel**

Perinatologist, Neonatologist  
State Secretariat of Health  
Bali, Dominican Republic

## **Peg Marshal, CNM, EdD, MPH**

Senior Technical Advisor for MCH and ID  
United States Agency for International  
Development (USAID)  
Washington, D.C. USA

## **Dr. Gerardo Martínez**

(CLAP/WR) PAHO/WHO  
Montevideo, Uruguay

## **Goldy Mazia, MD, MPH**

Adviser on neonatal health  
MCHIP  
Washington, D.C.

## **Dr. Martha Mejía**

PAHO/WHO consultant  
La Paz, Bolivia

## **Dr. Berta Pooley**

Save the Children, Bolivia

## **Dr. Oswaldo Revelo Castro**

Pediatrician-Neonatologist  
Chief, Medical Division  
Benjamin Bloom National Children's  
Hospital  
San Salvador, El Salvador.

## **Dr. Dalton Humberto Solís Montiel**

Pediatrician, Neonatologist  
Coordinator of Education and Research,  
Dr. Francisco de Ycaza Bustamante  
Children's Hospital  
Guayaquil, Ecuador

## **Dr. Nicolas Ramos Rodriguez**

Pediatrician, Neonatologist  
El Bosque University  
Bogotá, Colombia

## **Dr. Bernardo Sanchez**

PAHO/WHO consultant  
Asunción, Paraguay

## **Dr. Oscar Suriel**

Pediatrician, PAHO/WHO consultant  
Quito, Ecuador

## **Dr. María Beatriz Ugaz Estrada**

Pediatrician, Neonatologist  
Lima, Peru

## **Dr. Hernando Villamizar Gómez**

President  
Latin American Pediatrics Association  
(ALAPE)  
Colombia

# Acknowledgments:

The work of Dr. Rolando Cerezo M.  
Pediatrician, Neonatologist, and INCAP/PAHO  
consultant in Guatemala, who assumed the  
responsibility of reviewing the entire text, as well  
as overseeing the final changes and copy editing,  
is much appreciated.

Material translated into English with support  
from USAID.

It has been pointed out that mothers who do not get prenatal check-ups early and periodically, or whose check-ups are deficient, are at greater risk of perinatal, maternal, and neonatal morbidity and mortality.



# Foreword

In 2005, *The Lancet* published a series of articles on neonatal survival and health, calling the attention of the international community to the inadmissible death of millions of newborns despite the availability of effective and low-cost interventions. In Latin America and the Caribbean, neonatal mortality accounts for 58% of the deaths of children under five, and over 70% of deaths among those less than 1 year old. Seventeen percent of the region's 11.2 million annual births take place without trained staff, and mortality is highest in the most disadvantaged areas, creating a real problem of equity in the region and in individual countries. Infant mortality in indigenous and rural populations is nearly twice what it is in Hispanicized and urban populations. Populations in the poorest percentiles have less access to health services, fewer opportunities for prenatal check-up, and less access to drugs and state-of-the-art treatment. The result is higher mortality.

Given this scenario, effective, evidence-based interventions should be mustered to reduce mortality in the first 28 weeks of life, focusing on the family and the community, with emphasis on the poorest and most disadvantaged. This should take place in the context of primary health care that targets the communities' principal health problems by providing the promotion, prevention, treatment, and rehabilitation that they call for.

Such a process – unfolding with a life cycle orientation in the context of a continuum of care for mothers, newborns, and children, and with knowledge of the countries' potentials and constraints – can help the countries more rapidly reduce mortality in children under 5 to meet the United Nations Millennium Development Goals for 2015.

The Pan American Health Organization (PAHO), along with bilateral agencies, scientific associations, nongovernmental organizations, and other partners such as the Global Partnership, has taken this situation to heart. On the occasion of World Health Day in 2005, these organizations met in Washington D.C. to mark the emergence of an Interagency Strategic Consensus. That consensus has spawned the Neonatal Alliance, an entity that promotes newborn health in Latin America and the Caribbean in the context of the care continuum, promoting interagency initiatives, policies, and evidence-based programs that target vulnerable populations.

It was on the basis of the document prepared by the Strategic Consensus that PAHO submitted a resolution on neonatal health to its Directing Council in September 2006, at the 58th session of the Regional Committee. The resolution was adopted by all of the region's Ministers of Health in 2008 as a Strategy and Plan of Action to address newborn health, which the countries draw on in obtaining National Neonatal Health Plans that meet their local needs."

PAHO promoted and spread this work through the Integrated Management of Childhood Illness (IMCI) initiative and its neonatal IMCI component. IMCI is the result of efforts undertaken in 2000 by a group of professionals in the Region of the Americas. It has been adapted and used as a national strategy in more than 14 of the Region's countries.

Since science evolves rapidly, and far more rapidly in the neonatal field than in others, many evidence-based interventions that have now proven to reduce morbidity and mortality at low cost were not included in the first neonatal IMCI documents.

Therefore, the efforts of different institutions that collaborate with PAHO were brought together in the form of a group of experts in neonatology with broad IMCI experience. The group met to develop a generic document that presents all the evidence-based, cost-effective interventions that can be conducted at different care levels.

As a part of packages tailored to their national situations and to the level of care provided in their health systems, the region's countries can use these interventions, which have been scientifically shown to diminish maternal and neonatal morbidity and mortality, and which are conceived in the framework of the life cycle, the family and the community.

Our challenge is to ensure that these effective interventions, carried out in the framework of public health strategies, and focusing on priority areas and vulnerable populations, in fact contribute to reducing neonatal mortality, and hence infant mortality and mortality in children under 5 in the Region of the Americas.

**Dr. Gina Tambini**

Area Manager

Family and Community Health

Pan American Health Organization/World Health Organization

Washington, D.C.

# Preface

In September 2000, the United Nations Millennium Development Goals set clear standards for progress in the health and well-being of populations across the globe. Three of the eight goals established targeted significant challenges to global health: to combat HIV/AIDS and malaria; to reduce child mortality, and specifically decrease by two-thirds the mortality rate among children under 5; and to improve women's health by lowering the maternal mortality rate worldwide by three-quarters by 2015.

At Texas Children's Hospital and Baylor College of Medicine, we are participating in efforts to achieve the Millennium Development Goals and all endeavors that seek to effectively, comprehensively improve the health of a number of populations throughout the globe. We demonstrate our dedication to these goals through our wide range of programs targeting in global child health.

We are fortunate to have multi-lateral partnerships that over the past 15 years have allowed us to create the Baylor International Pediatric AIDS Initiative (BIPAI) at Texas Children's Hospital ([www.baylorails.org](http://www.baylorails.org)). In partnership with governments, international agencies, and private foundations, the BIPAI Children's Clinical Centers of Excellence Network has catalyzed the treatment of children with HIV/AIDS and the training of health professionals in Eastern Europe and across sub-Saharan Africa. We are proud that this Network now provides care and treatment to more than 100,000 children and is responsible for the ongoing training of health professionals in more than 13 countries.

Of the many lessons we have learned from our work in HIV/AIDS are two fundamental principles that guide all of our work internationally: 1) Initiatives must be evidence-based and ultimately target improvements in care and treatment; 2) Interventions must build local capacity and promote sustainable results at the local level.

Recognizing that evidence-based care for mothers and newborns has immense potential to close the care and treatment gap that leads to almost 4 million newborn deaths annually and accounts for 37% of all deaths in children under 5, Texas Children's saw an opportunity to collaborate in ways similar to BIPAI and achieve better outcomes for children in their first days of life.

In partnership with the Pan American Health Organization/World Health Organization, Texas Children's Hospital launched the WHO Collaborating Centre for Perinatal-Neonatal Health in August 2008. The goals of this partnership are simple:

- 1) **Create locally sustainable and highly-effective** operational interventions that translate clinical best practices into action, and have a **measurable positive impact** on maternal and child health
- 2) **Create globally replicable models** for maternal and child interventions aimed at advancing child health and reducing child mortality worldwide

The overarching philosophy of the WHO Collaborating Centre for Perinatal-Neonatal Health is to put knowledge into action. Much of the knowledge necessary for clinically-effective, evidence-based work in maternal and child health already exists. The challenge lies in assuring that the knowledge is adapted to be locally relevant and that it is used by the providers of care in the local communities.

With this in mind, Texas Children's works within three areas of focus as a WHO Collaborating Centre: capacity building across the health professional continuum through education and training; the development of tools and products targeting the improvement and dissemination of best practices; and broad-based quality improvement initiatives.

In this context, Texas Children's supports and encourages the broader dissemination of evidence-based guidelines for newborn care. And, we specifically support the distribution of these guidelines for newborn care in documents in English and Spanish throughout the Western Hemisphere and to other WHO regions. In the future, we look toward providing these crucial documents in more languages and disseminating them to areas in which the content is needed most.

We hope this document will prove useful at the point of patient care and in policy setting, and, we express our sincere gratitude to the Pan American Health Organization/World

Health Organization for their partnership. It is through these types of meaningful, solutions-focused collaborations that we can resolve the most critical challenges facing the health of vulnerable populations.

**Mark W. Kline, MD**

Physician-in-Chief  
Texas Children's Hospital  
Professor and Chairman  
Department of Pediatrics  
Baylor College of Medicine

**Stephen E. Welty, MD**

Chief of Neonatology  
Texas Children's Hospital  
Director  
WHO Collaborating Centre for Perinatal-Neonatal Health

# Introduction

The majority of infant and child health indicators in the Region of the Americas have improved substantially in most of the countries. However, neonatal health has been neglected by health authorities. Thus, there has been great emphasis on reducing postneonatal mortality and very little on reducing neonatal mortality. As a result, over 70% of infant mortality today is neonatal mortality.

In 2000, the neonatal component began to be incorporated in the Integrated Management of Childhood Illness (IMCI). Generic materials were developed, and adaptations were made in more than 12 of the region's countries.

The materials have been updated continually in their different adaptations as knowledge advances. The advances are evident in The Lancet's publication of material on neonatal health (1), which includes new scientific evidence demonstrating that many of the cost-effective interventions that can reduce neonatal morbidity and mortality remain unimplemented or are implemented poorly.

There is much discussion today on the importance of the quality of neonatal care, and on its short-, medium- and long-term consequences. Quality care means not only providing optimal care for every newborn, but also changing health workers' attitudes and practices so that they take advantage of modern evidence-based medical practices that have a direct impact on neonatal outcomes.

## **QUALITY OF CARE IN NEONATOLOGY**

Neonatal mortality is the basic indicator used to measure the development of newborns and the quality of care that they receive in a particular geographical area or set of health care facilities.

Infant and neonatal mortality rates vary in the different countries as a function of economic level, health conditions, and the quality of medical care. Neonatal mortality differs from postneonatal mortality in that it is less dependent on environmental and health conditions, and more dependent on organizational quality and the quality of perinatal and neonatal care, as well as on the high-technology medical treatment that occasional cases require.

The principal factors responsible for reducing neonatal mortality in those countries with low neonatal mortality rates – in addition to socioeconomic improvement and parental education – are good pregnancy monitoring and skilled childbirth assistance, the regionalization of perinatal care, the provision of timely neonatal intensive care, and good immediate care of newborns, including, when necessary, proper and immediate resuscitation.

High quality care in neonatology means proper care for every healthy newborn and more specialized care for those who need it, with quality protocols, an enabling environment, and tools for measuring short- and medium-term neonatal outcomes as well as satisfaction on the part of parents and care providers (2).

Neonatal management protocols are more easily adapted if they are:

- based on procedures of low complexity;
- compatible with existing standards, easy to follow, and ideally not dependent on the acquisition of new knowledge or skills (3);
- up-to-date and based on scientific evidence (4).

Projects targeting hospital staff and physicians working in the community, with a focus on quality and improvement, have been effective in enhancing preventive services for newborns while in the hospital. These preventive services include: immunization with hepatitis B vaccine, support/promotion of breastfeeding, risk assessment for hyperbilirubinemia, metabolic and auditory screening, attention to newborns' sleeping position, preventive respiratory support, nasal CPAP, hand hygiene, aseptic procedures in handling the umbilical cord, parenteral feeding, and planning for outpatient follow-up, among others things (5).

Neonatal monitoring and the comparative evaluation of data have provided a solid basis for lasting improvement of care. This type of evaluation has methodological limitations, and its findings vary, reflecting a complex mix of factors that often make practical application difficult. Discrepancies tend to arise in comparing research, evidence, information from monitoring, standards, and practices (6, 7).

Systematic review of the literature is a way of keeping up to date with developments in daily clinical practice. Traditional practices of health professionals – such as reading educational materials, or attending conferences, courses, and workshops – cannot replace them. These activities have but minimal impact on care quality. Another factor is that, if clinicians fail to make practical use of scientific evidence on policies and protocols, it is not solely because they are unfamiliar with recommendations or disagree with them. The more important impediment lies in local obstacles to implementing the recommendations.

## **EVIDENCE-BASED MEDICINE (EBM)**

According to the definition proposed by Sackett in 1996 (8), "EBM consists of the conscious, explicit and judicious use of the best tests available when making decisions on comprehensive care for each patient." The focus is on the patient, not merely on the disease or process that is present. Thus, from the time of precise diagnosis to the selection of the best thera-

peutic option, the patient's rights, principles, and preferences must be identified and taken into consideration.

Textbook information can be questionable as time passes. The quantity of publications and information sources is enormous, and our time, knowledge, and ways of locating, reading, interpreting, and assimilating what is published are less than sufficient. The principal value of EBM is as a tool to quantify and identify the most valid data or studies, to clarify the best way of understanding and using the evidence, and to facilitate translating it into practice.

## **EVIDENCE-BASED MEDICINE IN PERINATOLOGY AND NEONATOLOGY**

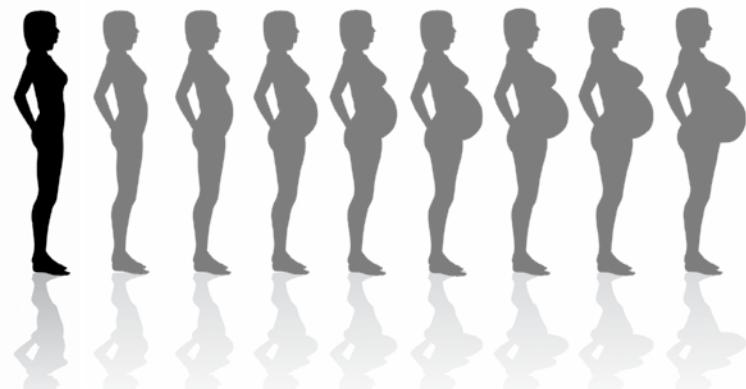
Much perinatal and neonatal medical evidence today has been subjected to major, systematic review, and provides data for practical application. Systematic reviews provide information on original work by selecting work, evaluating it critically, and summarizing it, following an explicit, formal, and rigorous methodology. Systematic reviews and meta-analyses are generically known as synthesis resources. The Cochrane Library offers systematic reviews of the Cochrane groups, and PubMed Clinical Queries makes it possible to locate systematic reviews in the MEDLINE database. Other sources that can be consulted are the WHO Reproductive Health Library and the publications of the Latin American Center for Perinatology (CLAP/WR).

The interventions presented in this document were selected on the basis of current scientific evidence and on an evaluation of their utility and potential for reducing neonatal morbidity. They can be included in packages designed for the levels of care that different health systems offer.

### **Dr. Yehuda Benguiui**

Asesor Principal Salud del Recién Nacido  
Proyecto del Curso de Vida Saludable  
Organización Panamericana de la Salud/  
Organización Mundial de la Salud

# I. Prepregnancy interventions



Preconceptional care is recognized as a critical component in the care of women of childbearing age. It is defined as a set of interventions designed to identify and reduce risk factors. The relevant risk factors are a function of demographic and medical variables that relate directly or indirectly to causes of malformations or to poor perinatal outcomes (9-14).

Preconception counseling optimizes perinatal outcomes by detecting pathologies, medications, habits, and behaviors that could be hazardous for the future mother or fetus. This counseling is important because:

- Most of the relevant risk factors are present before conception.
- The countries have high rates of adolescent and unwanted pregnancies.
- Some interventions are more effective if carried out before pregnancy.
- Some interventions are not feasible during pregnancy.

At the same time, it is important to encourage healthy lifestyles in couples by offering **health promotion**. This means giving them the tools they need to improve their health and have greater control over it. The idea of health promotion was conceived in 1990 by the World Health Organization (WHO), which defined it as encompassing all of a population's activities, as well as its health services, health authorities, and other social and productive sectors, insofar as all of these bear on improving individual and collective health.

## 1. Using body mass index (BMI) to monitor nutritional status prior to pregnancy

### Antecedentes

Various anthropometric indicators of nutritional status have been used in the last few years. Body mass index prior to pregnancy has recently been considered a good indicator and tool for diagnosing, monitoring, and following up on women's nutritional status from the preconception period through the pregnancy. The Institute of Medicine (IOM) of the American Academy of Sciences has now adopted BMI as a tool, and WHO recommends its broad use for nutritional evaluation of nonpregnant adults in the context of prepregnancy nutritional assessment (15).

Any deviation from adequate nutritional status should be rectified before conception, through nutritional support measures to lower weight (by promoting dietary changes and physical activity).

<b>Calculation</b>	Weight (kg)/height(m) <sup>2</sup>
<b>Interpretation</b>	<p>Underweight: BMI &lt; 18.5</p> <p>Normal: BMI 18.5–24.9</p> <p>Overweight: BMI 25.0–29.9</p> <p>Obese: BMI ≥ 30.0</p>
<b>Evidence</b>	<p>A body mass index of &lt; 18.5 is associated with malnutrition. Before pregnancy, it is a predictor of low birthweight (LBW) and intrauterine growth retardation (IUGR).</p> <p>Recent evidence points to prepregnancy BMI as an independent predictor of many adverse effects for the pregnant woman and fetus in both the short and long term. The greatest risk factor, which increases the incidence of congenital malformations (CMFs) by as much as a factor of 6 over a period of a decade, is a threefold increase in the prevalence of preconceptional diabetes preceded by increased obesity (20–23).</p>

## 2. Folic acid (FA) and multivitamin (MV) supplementation

**Background** Folic acid is one of the most important B-complex vitamins before and during the pregnancy, for studies have shown that adequate intake diminishes the risk that a baby will suffer from neural tube defects (NTDs). The neural tube is the structure from which the brain and spinal cord develop during the first 3 months of gestation. If it does not mature and close properly (28 days after conception) certain defects such as spina bifida, encephalocele and anencephaly can appear. Anencephaly is incompatible with life, and the symptoms associated with spina bifida include infantile paralysis, lack of sphincter control, and learning disabilities.

Women at high risk for NTDs and other congenital malformations (CMFs) are those with a family history of NTDs, those who are taking antiepileptics or methotrexate, those with diabetes, obese women with a BMI ≥ 0 kg/m<sup>2</sup>, and mothers with sickle cell anemia or thalassemia. Women with poor compliance with medical instructions (for FA and/or MV supplementation) and low consumption of FA-rich foods, and those consuming teratogens (alcohol, tobacco), require counseling on healthy lifestyles, FA-rich diets, and 4 mg of FA daily plus a MV supplement in order to prevent NTDs and other CMFs.

**Administration** To prevent neural tube defects (NTDs) (in children without prior problems), as well as other congenital malformations (CMFs), in cases of mothers not at risk for NTDs and other CMFs, administer:

- From three months before pregnancy to three months after birth, or as long as breastfeeding.
- [TN: Those two bulleted instructions are either contradictory or need some further explanation]
- It is generally administered in combination with ferrous sulfate.

**Dose** Folic acid, 400 µg/day (0.4 mg/day) orally.

In women at high risk for NTDs (malnutrition, obesity, history of previous children with malformations), dosage should be increased to 4–5 mg a day orally. Multivitamins should also be administered during the period the medication is taken.

To prevent occurrence and recurrence (in children with previous problems) where mothers are at HIGH risk for NTDs and other CMFs:

Administer 4–5 mg/day of folic acid plus multivitamins from at least 3 months before pregnancy until 3 months after birth. Between the 3rd month of the pregnancy and the 3rd postpartum month (or as long as breastfeeding), administer 0.4–1 mg of folic acid daily plus multivitamins orally.

**Evidence** The administration of folic acid alone reduces the incidence of neural tube defects by 72% (42%–87%). Intake of FA for a year or more before conception significantly reduces the risk of spontaneous preterm birth by 70% between weeks 20 and 28 and by 50% between weeks 28 and 32.

Periconceptional administration of multivitamins reduces the risk of preterm birth before the 34th week by 71% (statistically significant), and in non-obese women it reduces births of babies who are small for their gestational age by 46%.

Recent studies show periconceptional multivitamins plus folic acid significantly (46%) reducing not only NTDs but also other CMFs: cleft lip and palate, cardiac defects, defects of the limbs and urinary tract, and hydrocephalus. The multivitamin mix in question includes: A, 4000 IU; B1, 1.6 mg; B2, 1.8 mg; nicotinamide, 19 mg; B6, 2.6 mg; calcium pantothenate, 10 mg; biotin, 0.2 mg; B12, 4.0 mcg; C, 100 mg; D, 500 IU; E, 15 mg; folic acid, 0.8 mg. Also included are the minerals calcium, 125 mg; phosphorus, 125 mg; magnesium, 100 mg; and iron, 60 mg, as well as three trace elements: copper, 1 mg; manganese, 1 mg; and zinc, 7.5 mg.

Other benefits reported for children from periconceptional administration of multivitamins are significantly reduced risk of cerebral tumor (27%), neuroblastoma (47%), and leukemia (39%).

A random double-blind study ( $n = 31,290$  pregnant women) compared the administration of multiple micronutrients (MMNs) – iron; AF; vitamins A, D, E, C, B1, B6, and B12; niacin, zinc, copper, selenium, and iodine – with giving only FA + iron (FAFe) during pregnancy. For 34% of the women, administration began in the first trimester, while 43% began the regimen in the second trimester. All continued until the third postpartum month. The MMN group was found to have reduced early infant mortality (< 3 months of age) by 18%, RR 0.82 (95%, CI 0.70-0.95) (24-41).

### **3. Preventing and treating anemia with iron**

#### **Background**

The prevalence of anemia in preschool children and adolescent girls remains high in the developing countries. WHO has estimated the prevalence of anemia in preschool children in the region of Latin America and the Caribbean (LAC) to be 39.5%, and the prevalence among nonpregnant women to be 23.5%. Assuming that the average absorption of medicinal iron is 10%, the daily requirements of elemental iron would be 10 mg for children, men, and postmenopausal women; 20 mg for nonpregnant women; and 30 mg for pregnant women.

#### **Diagnosis**

Hb < 12 g/dL (adjust as a function of altitude).

#### **Treatment**

Hb < 7 g/dL: 120 mg of elemental iron/day orally.

Hb 7-12 g/dL: 60 mg of elemental iron/day orally.

Administer between meals or at bedtime to avoid alkalinizing the food and to take advantage of the high nighttime levels of gastric acidity.

A rise of 2 g/dL or more of the Hb, and/or a hematocrit that has normalized after 2-3 months of iron therapy with iron, is a good response to treatment, but treatment should continue for another 3-6 months to replace iron reserves in the bone marrow. Iron-rich foods should also be recommended: red meats (liver is the greatest source), eggs (yolks), fish, vegetables (lentils and beans), poultry, raisins, and whole-grain bread.

Vitamin C (citrus juice, tomato juice) facilitates iron absorption, while coffee, tea, soft drinks, milk, calcium, magnesium, quinolones, tetracyclines, and antacids impede it. The best iron supplement is ferrous sulfate.

#### **Evidence**

Iron deficiency anemia in adolescent girls is associated with an elevated risk of anemia during pregnancy. It has also been associated with alterations in cognitive functions and memory, reduced school performance, and immunosuppression with increased infection rates (42-51).

### **4. Micronutrients and reproductive health in women**

#### **Background**

Proper nutritional status in women before, during, and after pregnancy is an important element of

reproductive health. It helps maintain health, reduces the risk of adverse outcomes during pregnancy, and reduces the prevalence of birth defects and chronic disease in the children after they are born. Pregnancy creates a special metabolic demand for high-quality nutrients. A good selection of food can provide most of the recommended levels of nutrients. Beyond food intake, however, good nutrition depends on a woman's economic environment, social and cultural conditions, and personal habits.

<b>Diagnosis</b>	Diagnosis is based on ascertaining the woman's food consumption, eating habits, and socioeconomic status. BMI is useful in determining nutritional status.
<b>Treatment</b>	Appropriate balanced diet plus multivitamin supplements.
<b>Evidence</b>	Nutritional imbalances can produce detrimental effects in pregnant woman (hypertension, anemia, complications during delivery) and can cause harm to the fetus (congenital malformations, premature delivery, intrauterine growth restriction). It can also be detrimental to the composition of the mother's milk (52-55).

## 5. Detecting and treating sexually transmitted infections (STIs) before pregnancy

<b>Background</b>	Most STIs affect both men and women, but the health problems involved can be more serious in women. STI in a pregnant woman can create severe health problems for both mother and baby.
<b>Diagnosis</b>	Chlamydia ( <i>Chlamydia tracomatis</i> ), gonorrhea ( <i>Neisseria gonorrhoeae</i> ), herpes virus, HIV/AIDS, syphilis, human papillomavirus (HPV).
<b>Evidence</b>	In women, chlamydia and gonorrhea can induce pelvic inflammatory disease, which can lead to infertility or problems with a pregnancy. In men, gonorrhea can cause epididymitis and infertility. HIV kills or damages the cells of the organism's immune system, leading to severe infections and death. It is transmitted during pregnancy, labor, and breastfeeding. Vaccination for HPV can diminish cancer of the cervix, vulva, vagina, and anus. In men, HPV can cause cancer of the anus and of the penis (56-66).

## 6. Deparasitization in areas of high prevalence

<b>Background</b>	In areas of endemic intestinal helminthiasis, routine deparasitization considerably reduces the prevalence of anemia in women and its possible complications.
<b>Diagnosis</b>	Stool test (ascaris, oxyurids, uncinaria, Trichuris).
<b>Treatment</b>	Albendazole, 400 mg orally, single dose, or Mebendazole, 100 mg twice a day orally for 3 days, or Mebendazole 500 mg orally, single dose.
<b>Evidence</b>	Deparasitization of women of childbearing age reduces anemia and its complications and is associated with lower risk of anemia during pregnancy (67-68).

## 7. Detecting and treating chronic diseases (cardiovascular, nutritional, endocrine)

<b>Background</b>	The prevalence of overweight in women between the ages of 15 and 49 in some Latin America and Caribbean countries is 25.1%, and the prevalence of obesity averages 8.5%. Obesity is a risk factor for several chronic noncommunicable diseases such as diabetes, hypertension, cancer, hypothyroidism, and lupus, which not only are detrimental to the health of women of childbearing age, but can lead to fetal death.
<b>Diagnosis</b>	Overweight: BMI 25.0–29.9 Obesity: BMI $\geq 30 \text{ kg/m}^2$
<b>Evidence</b>	Psychiatric and emotional disturbances, poor school performance and dropout, prolonged treatments, cardiovascular disease (69-79).

## 8. Detecting, preventing, and managing domestic violence

<b>Background</b>	Violence against women by their partners or ex-partners is widespread in the world, occurring in all social groups regardless of economic status, cultural level, or any other variable. Although it continues to be difficult to quantify, since not all cases are known to people other than the couple, it is assumed that many women to suffer or have suffered from this type of violence. Studies conducted in developing countries point to abuse rates of around 20%.
<b>Diagnosis</b>	Repeated injuries, low self-esteem, depression, personality disturbances, poor school performance, neglect of personal hygiene.
<b>Evidence</b>	Violence diminishes motivation and morale, causes physical and psychological damage, depression, and post-traumatic stress. (80-89).

## 9. Detecting, preventing, and managing alcohol and tobacco consumption

<b>Background</b>	Consumption of alcohol at a very early age is a reliable predictor of future problems related to alcohol consumption, while drinking in adolescents is associated with high-risk sexual behavior and is predictive of early motherhood and a smoking habit.
<b>Diagnosis</b>	Antisocial personality, depression, smoking, poor school performance.
<b>Evidence</b>	Studies suggest that several genes that code parts of receptors for the neurotransmitter gamma-aminobutyric acid (GABA) are involved in the development of alcohol or nicotine dependency. Other studies have identified further genes that may be associated with alcohol or nicotine dependency. Findings to date indicate that both common and drug-specific genetic influences play a role in the development of alcohol and nicotine dependency.  Findings of delayed reproductive functions in alcoholic women are compatible with the occurrence of alcohol-related reproductive dysfunctions (90-95).

## 10. Detecting, preventing, and managing depression

<b>Background</b>	Depression ranks as the fourth most important disease in young people's estimated disease burden. It is a common problem, with prevalence figures as high as 8%. Strong associations link depression, suicide, and smoking in adolescents.
<b>Diagnosis</b>	Poor school performance, social dysfunctions, abuse of psychotropics, attempted suicide, poor family relations, and scant use of health services.
<b>Evidence</b>	Depressive symptoms are common in adolescents, and their course is difficult to predict. Most adolescents with minimal symptoms maintain their state and seem to be at low risk, but adolescents with moderate or serious symptoms of depression should be monitored and reassessed – above all if pregnancy is a possibility.  Several risk factors for developing depressive symptoms during pregnancy have been identified. Strong associations have been established with: maternal anxiety, stress due to major life events (death in the family, divorce), personal history of depression, and lack of support from one's intimate partner. There is moderate association with: lack of social support, domestic violence, and unwanted pregnancy (96-103).

## 11. Detecting and preventing cervical cancer

<b>Background</b>	Cervical cancer (CC) is one of the most common cancers in women at the world level. A number of strategies have been recommended to limit the spread of human papillomavirus (HPV) 16 and 18, which is one of the principal risk factors for cervical cancer, causing 70% of all cervical cancers in the world. The strategies include promoting lifestyles and behaviors that minimize risk, promoting the use of condoms in sexual relations, reducing the number of sexual partners, and implementing strategies for lower-risk sexual activities.
-------------------	---

<b>Diagnosis</b>	Cytology and colposcopy are highly sensitive as detectors of CC, but their specificity is quite low. Screening tests for HPV are both highly sensitive and highly specific. Combining HPV testing with colposcopy produces a sensitivity of 97.2% and raises negative predictive value to 92.3%.
<b>Prevention</b>	Vaccination against human papillomavirus (HPV) is advised for all adolescents 11-12 years of age, and for women ages 13-26 not previously vaccinated. Minimum age for vaccination is 9 years.
	Dose 1 upon detection.
	Dose 2 two months after dose 1.
	Dose 3 six months after the first.
	Boosters are not recommended. The vaccine can be administered with other vaccines.
<b>Evidence</b>	Early detection of CC by screening all women of childbearing age, as well as vaccination for human papillomavirus, reduces the number of women requiring colposcopy, biopsy, and treatment of precancerous cervical injuries ( <i>104-114</i> ).

## 12. Complete immunization series

<b>Background</b>	Childhood vaccination recommendations have increased over the years. Many care providers, patients, and families have concerns about the safety and efficacy of vaccines. Health systems and pediatricians currently face numerous challenges in improving immunization rates among children and adolescents. The strategies that can be used include fostering coverage by counseling parents with clear information about the risks and benefits of vaccines, and taking advantage of routine checkups as an opportunity for immunization.
<b>Prevention</b>	Complete the vaccination series following the country's ministry of health standards.
<b>Evidence</b>	Immunization to prevent communicable diseases has been one of the most effective interventions for diminishing morbidity and mortality in the countries of the Region of the Americas. Women of childbearing age should complete the entire vaccination series before their next pregnancy to diminish the risk of intrauterine infection, which is most often fatal to the fetus ( <i>115-122</i> ).

## 13. Detecting and treating periodontal disease

<b>Background</b>	Caries is a disease of the hard tissue of the teeth. It is caused by an imbalance over time in the interactions between cariogenic bacteria in the dental plaque and fermentable carbohydrates (principally sugars). It is also a well-established fact that many health conditions have oral manifestations.
<b>Diagnosis</b>	Dental checkups (semi-annual).
<b>Prevention</b>	Regular tooth brushing with toothpaste that contains fluoride in concentrations of 1000 ppm or greater.
<b>Evidence</b>	Periodontal disease is associated with adverse health conditions such as rheumatic fever, bacterial endocarditis, and systemic diseases, and can affect eating and food selection. Oral and facial pain from the teeth, temporomandibular joint disorders, and oral infections affect social interactions and daily behavior. The consequences of oral disorders can be significant for our society's young people, not only physically and socially, but also economically ( <i>123-131</i> ).

## 14. Preventing pregnancy in adolescents

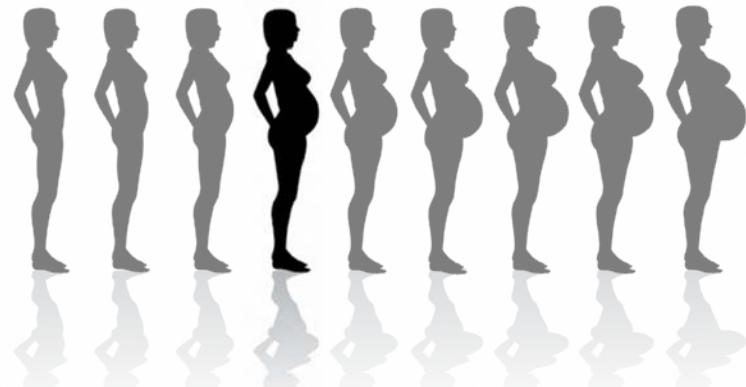
<b>Background</b>	Teenage pregnancy is a major public health challenge in both the developed and developing countries. Numerous preventive strategies have been used by countries around the world to address this problem, including health education, developing other abilities (sports, languages, etc.), and improving access to contraceptives.
-------------------	---

<b>Prevention</b>	Two affordable options have been proposed for those who wish to reduce negative health outcomes associated with adolescent pregnancy. The first consists of expanding the range of life choices for the young female population, in an attempt to improve their social and economic circumstances. The second is to ensure that their intentions to get pregnant are understood, so as to provide the appropriate services and obtain the best possible outcome for mother and child.  Educational interventions that include service providers and parents, as well as programs on the use of contraceptives, have been effective in diminishing unwanted pregnancy among adolescents.
<b>Evidence</b>	Adverse outcomes have been identified in mothers, principally those under 15 (high rates of caesarean section, puerperal infection, and intrapartum complications), as well as in fetuses (preterm birth, low birthweight, and neonates who are small for their gestational age) (132–138).

According to the new WHO model, five check-ups are sufficient for pregnant women without complications, that is, for 75-80% of pregnant women.



# III. Interventions during pregnancy



Maternal mortality is one of the major health problems in many Latin America and the Caribbean countries, and has major impact on the lives of children and their families. It is very closely linked to health determinants such as poverty, access to health services, education, and cultural barriers, and with maternal factors such as parity, adolescent pregnancy, short birth interval, and families' and communities' ignorance of warning signs.

Pregnancy usually proceeds without complications, or with minor complications that do not merit intervention. However, a percentage merit certain interventions to ensure the continued health and safety of fetus and mother through the pregnancy, e.g., prenatal check-ups and interventions in order to prevent maternal complications that could cause fetal, maternal, or neonatal death.

The interventions proposed are based on scientific evidence and real cost, and have been shown to reduce maternal, fetal, and neonatal mortality. They can be implemented at any level of complexity of health services. Complexity of implementation and high economic cost are less likely to be barriers than is acceptance by the care providers who work with pregnant women.

## 1. Prenatal check-up

**Background** It has been pointed out that mothers who do not get prenatal check-ups early and periodically, or whose check-ups are deficient, are at greater risk of perinatal, maternal, and neonatal morbidity and mortality.

**Frequency** According to the new WHO model, five check-ups are sufficient for pregnant women without complications, that is, for 75–80% of pregnant women. The activities included in the basic component fall into three general areas:

- Ascertaining socioeconomic status, and detecting pathologies that increase the possibility of specific adverse outcomes;

- Carrying out proven therapeutic interventions; and

- Educating pregnant women to recognize danger signs and emergencies during pregnancy, and teaching them how to deal with them.

<b>Evidence</b>	Prenatal check-ups provide an opportunity to increase access to safe delivery in an environment that provides care for obstetric emergencies as well as skilled institutional neonatal care. With prenatal check-ups, fewer complications have been reported during pregnancy and childbirth: less preeclampsia, urinary tract infection, postpartum anemia, and maternal mortality, as well as lower incidence of LBW (139-151).
-----------------	---

## 2. Preventing fetal death

<b>Background</b>	Fetal death, defined as occurring when any infant at birth has shown no sign of life during the perinatal period, accounts for at least 60% of all perinatal deaths and 75% of all potentially preventable miscarriages. The causes of fetal death include birth defects, isoimmunization, toxemia, syphilis, hemorrhage before delivery, maternal illness or trauma, and unexplained causes.
<b>Risk factors</b>	<i>Maternal factors:</i> multiparity, advanced age, obesity, periodontal disease, smoking. Social factors: education, racial group, access to quality care.
<b>Prevention</b>	Adequate access to obstetric care, particularly during labor and delivery, and better screening for and treatment of syphilis and malaria in endemic areas, can dramatically reduce fetal mortality in developing countries (152-170).

## 3. Preventing preterm birth

<b>Background</b>	A preterm birth is one that occurs during the first 37 weeks of gestation. Percentages of premature births vary from country to country, but are generally between 6% and 9% in the less developed countries, and above 9% in the more developed countries.
-------------------	---

The rate of preterm birth (< 37 weeks of GA) in 2008 was 10% at the world level, 12% in Africa, 11% in North America, 8% in Latin America and the Caribbean, and 6% in Europe. In Central America specifically, it was between 9% and 10%. Preterm birth rates are increasing throughout the world due to the rising number of late-preterm births (34 to 36 weeks of gestation).

According to WHO, preterm newborns account for more neonatal mortality than do low birthweight babies: for nearly 75% of perinatal mortality and 50% of long-term neurological sequelae. Preterm babies are subject to acute complications as well as long-term problems of various kinds that generally entail high economic costs. They include respiratory, gastrointestinal, immunological, central nervous system, hearing, vision, cognitive, and social/emotional problems.

In approximately one half of preterm births, the cause is unknown. Three types of risk factors that can contribute to spontaneous preterm birth have been identified:

Social stress and race: Poverty, low educational level, being single or adolescent, and lack of adequate prenatal check-ups increase the probability of preterm birth and low birthweight. In the United States, adjusting for other risk factors, blacks are twice as likely as whites to experience preterm birth, and their recurrence rate is four times as great.

*Infection and inflammation:* Chorioamnionitis is the most obvious example of an infection that triggers labor and preterm delivery. There is microbiological evidence that infection may account for up to 25%-30% of preterm births, with rates of bacterial colonization as high as 79% in babies born at 23 weeks, and 11% among those born at 31-34 weeks of gestation. In most labors and preterm deliveries, there are no clinical manifestations of intrauterine infection, even when there is histological evidence of inflammation in the decidua, fetal membranes, and umbilical cord. For example, *mycoplasma* and *ureaplasma* are microorganisms frequently cultivated from fetuses, but pregnant women at risk of preterm delivery due to vaginal colonization by them seldom receive erythromycin treatment. New mechanisms of chronic infection and inflammatory response have been identified in urinary tract infections. Bacteria and fungi tend to form communities or biofilms within a self-developed polymer matrix. The CDC estimates that

these biofilms play a role in up to 65% of human bacterial infections, such as in periodontitis, urinary tract infection, endocarditis, and other chronic infections.

*Genetic factors.* A history of preterm births significantly increases a mother's risk of future preterm births, just as a history of post-term birth increases the risk of future post-term births.

#### Risk factors

Many maternal and fetal characteristics have been associated with preterm birth, including demographic characteristics and nutritional status of the mother ( $BMI < 18.5$ ), a history of premature birth, syphilis, birth defects, certain psychological characteristics, detrimental habits (tobacco, alcohol, drugs), infections, uterine contractions, and cervical size, as well as genetic and biological markers.

Preterm births can be spontaneous or a result of medical indications. Spontaneous preterm births occur as a result of preterm labor or preterm premature rupture of membranes (< 37 weeks of GA), the latter accounting for 25%-30% of preterm births. Medically indicated preterm births (25%-30% of the total) occur because of health problems in the mother or fetus: preeclampsia, placenta previa, or limited fetal growth. In 40%-50% of cases, the etiology of the preterm birth is unknown. Babies born prematurely when spontaneous preterm labor begins without apparent maternal pathology more frequently show deficient intrauterine growth.

Medically indicated preterm births share with spontaneous preterm births several **maternal risk factors** that increase the probability of preterm birth. Prepregnancy BMI and weight gain during pregnancy below the levels recommended by the Institute of medicine (IOM) increase risk by a factor of 14, while levels above those recommended increase risk by a factor of 3. Urinary tract infections – whether asymptomatic (asymptomatic bacteriuria) or symptomatic – and African-American ethnic identify both double the risk of preterm birth. Multiple gestation increases risk by a factor of 6, a history of preterm birth by a factor of 4, and vaginal bleeding (placenta previa, abruption, and unexplained persistent bleeding) by a factor of 3.

The **risk of a recurrence of preterm birth** rises with the number of previous preterm births, and rises in an inverse relation with the gestational age of previous preterm babies. Chronic hypertension increases risk by a factor of 7, and preeclampsia by a factor of 13. Asthma, heart disease, pre-gestational and gestational diabetes mellitus, hyperthyroidism, and systemic lupus erythematosus also increase risk. Other maternal risk factors are periodontal disease, non-existent or inadequate prenatal check-ups, age < 18 or > 35, exhausting work, high stress level, anemia, tobacco, cervical injuries or uterine anomalies, and use of technology for reproductive assistance. **Fetal risk factors:** red blood cell isoimmunization, twin-to-twin transfusion, and congenital malformations.

#### Prevention

Preventing preterm births should be a national public health priority, with programs of evidence-based multidisciplinary interventions, education for health care professionals and users on prevention, a strong national system of vital statistics, careful research on risk factors, and access to quality prepregnancy and perinatal health care.

*Promote healthy lifestyles, prevent and treat the risk factors associated with preterm birth, both spontaneous and indicated, from pre-pregnancy through pregnancy, childbirth and the neonatal period.*

The problem of preterm birth and the mothers involved should be treated at three levels: primary (preventing and reducing risk in the population), secondary (identifying and treating women who are at greater risk), and tertiary (providing treatment to reduce morbidity and mortality after preterm delivery begins). In the last 30 years, morbidity and mortality among premature babies has been reduced by improving tertiary obstetric and neonatal intervention. Little has been done through primary and secondary intervention to reduce the spontaneous preterm birth rate.

A combination of approaches can help reverse the rising rates of preterm birth: emphasizing perinatal quality initiatives, increasing evidence-based interventions, using tools that improve predictions regarding labor, encouraging the spread of effective community programs, promoting commitment to

more equitable conditions, enhancing preconception health promotion, developing effective continuing education programs for health workers, and producing health education materials with content that is culturally acceptable to patients.

### **Evidence-based interventions**

- *Treat risk factors and prevent pathologies starting before conception:* poor nutrition (malnutrition or obesity as measured by BMI and evaluated according to WHO standards, along with weight increase during pregnancy that is below or above IOM recommendations).
- *Treat and prevent pathologies during pregnancy.*
- *Space births.* Achieve a birth interval between 2 and 5 years. One meta-analysis of outcomes comparing birth intervals of under 6 months with intervals of 18-23 months found a 40% increase in preterm births, a 61% increase in incidence of LBW, and 26% more neonates with intrauterine growth retardation when the interval was smaller. Preterm births increased by 20% when the birth interval was greater than 60 months. In countries with high birth rates, promoting the spacing of births could reduce deaths by one third and infant mortality by 10%.
- *Achieve the weight gain recommended by the IOM during pregnancy, as measured by BMI.* With any preconceptual BMI, a low (0.12-0.22 kg/week) or very low (<0.12 kg/week) weight increase during pregnancy increases the risk of preterm birth (< 32 weeks of gestational age) by a factor of 14 in women with low BMIs, and by as much as a factor of 5 in the obese. Given any preconceptual BMI and high weight gain during pregnancy (>0.79 kg/week), the risk of preterm birth rises by a factor of 4, while a BMI indicating great obesity increases the risk by a factor of 8.
- *Administer periconceptional folic acid.* This prevents preterm birth and neural tube defects. A regimen of folic acid plus multivitamins prevents preterm birth before 34 weeks of GA, as well as various malformations besides NTD.
- *Monitor infection.* Conduct HIV tests of all pregnant woman. Check for syphilis in the first and third trimester of pregnancy, or before delivery or discharge from the care facility. Conducting RPR before discharge for all women with a history of miscarriage, fetal death, or newborns infected with syphilis diminishes miscarriages, preterm births and perinatal mortality.
- *Detect and treat asymptomatic urinary tract infections in pregnant women.* This reduces the incidence of low birthweight and preterm birth by one third.
- *Treat periodontal disease in pregnant women.* This reduces preterm birth by 45-65%.
- *Progesterone* for women with a history of spontaneous preterm birth significantly reduces the incidence of < 37-week and < 34-week births. In women with short cervical canals (< 15 mm), progesterone has been shown to diminish preterm births < 34 weeks of GA, as well as neonatal sepsis.
- *Erythromycin or another antibiotic* (ampicillin, benzylpenicillin, or clindamycin) in cases of premature rupture of membranes at under 37 weeks of gestational age significantly reduces births within 48 hours, maternal and neonatal infections during the first seven days, and the need for oxygen and surfactant.
- *Nifedipine* as an inhibitor of uterine contraction significantly reduces births within 7 days of treatment, as well as reducing adverse effects and neonatal morbidity (RDS, cerebral hemorrhage, and necrotizing enterocolitis) in newborns of GA< 34 weeks.
- *Corticoids (dexamethasone or betamethasone)* before birth significantly reduce neonatal mortality, RDS, cerebral hemorrhage, necrotizing enterocolitis, sepsis, admissions to intensive care, and mechanical ventilation.
- *Calcium* for pregnant women at high risk of preeclampsia significantly diminishes preterm birth, low birthweight and neonatal mortality.

- *Aspirin* for pregnant women at risk for preeclampsia significantly diminishes preterm birth, the incidence of newborns who are small for their gestational age, and perinatal mortality.
- *Avoid elective births (vaginal or by caesarean section) before 39 weeks of gestation.* There is sufficient evidence that serious neonatal and infant morbidity and mortality are lower when gestational age is optimal (39–41 weeks) and when birthweight is optimal (3,500–4,250 grams), while morbidity and mortality increase when births are scheduled before 39 weeks of gestation in pregnancies without complications.

<b>Evidence</b>	Premature births are responsible for 75% of perinatal mortality and more than half of long-term morbidity. Those premature babies who survive are at high risk of developing problems of neurological development as well as gastrointestinal and respiratory complications (171–198).
-----------------	--

## 4. Managing premature rupture of membranes

<b>Background</b>	Premature rupture of membranes is the rupture of the amniotic membranes before labor has begun. When the fetus has not reached term (< 37 weeks of GA), the rupture is called preterm premature rupture of membranes (PPROM).
-------------------	---

<b>Diagnosis</b>	Discharge of amniotic fluid with a characteristic odor. When the rupture is not recent, a vaginal compress can be placed on the vulva, and the appearance and odor of the compress examined in an hour. A sterilized speculum can also be used for vaginal examination. The fluid can be seen flowing from the cervix or accumulating in the posterior fornix. Ask the woman to cough, which can produce a spurt of fluid.
------------------	--

<b>Treatment</b>	<b>In preterm premature rupture of membranes (PPROM):</b> <i>In the absence of labor (&lt; 37 weeks of GA), chorioamnionitis, preeclampsia, fetal stress, or any other contraindication to continuing the pregnancy:</i>
	<ul style="list-style-type: none"> <li>■ Give erythromycin, 250 mg orally every 6 hours for 7 days or until the birth.</li> <li>■ Refer to a hospital where preterm newborns receive adequate care.</li> <li>■ If GA &gt; 26 weeks or GA &lt; 36 weeks, administer prenatal corticoids.</li> </ul>

*In the presence of labor, with gestation < 37 weeks, administer the antibiotic as soon as possible after the preterm labor begins, and until delivery, in order to reduce the risk of transmitting group B streptococcus to the fetus (the cause of up to 50% of early sepsis in Latin America and the Caribbean):*

- Penicillin G, initial dose 5 million units IV, followed by 2.5 million units IV every 4 hours orally; or
- Ampicillin, initial dose 2g IV, followed by 1g IV every 4 hours; or
- In women allergic to penicillin, clindamycin, 900 mg IV every 8 hours until delivery.

To maximize the antibiotic's prophylactic efficacy, the first dose should be administered at least 2 hours before the birth.

Prophylactic antibiotic treatment for streptococcus B is unnecessary for women with preterm premature rupture of membranes unless labor is in progress.

If there are signs of infection (fever, malodorous vaginal secretion):

Treat it as amnionitis, beginning antibiotics and continuing until delivery: ampicillin, 2g IV every 6 hours + gentamicin, 5 mg/kg bodyweight IV every 24 hours.

- Refer to hospital.

<b>Evidence</b>	The administration of antibiotics in PPROM reduces the risk of RDS, early neonatal infection, and neonatal mortality (199–208).
-----------------	---

## 5. Using the CLAP/WR perinatal clinical record (PCR) and perinatal card

**Background** The PCR is an instrument designed by CLAP/WR, and published in 1983 to contribute to decision-making on the individual clinical management of pregnant women during prenatal check-ups, childbirth, and puerperium, and management of the neonate from birth to time of discharge.

The perinatal card makes it possible to integrate the health team's actions during pregnancy, childbirth, and puerperium, preventing the problem of lack of data, and helping to improve care quality.

**Frequency** At prenatal check-up, delivery, puerperium, and the neonatal stage.

**Evidence** Processing the data from the PCR enhances perinatal care providers' capacity for self-evaluation, makes personnel aware of the importance of documenting actions and observations thoroughly, and gives perinatal care facilities a fast and easily used tool for operational research (209–210).

## 6. Weight gain during pregnancy

**Background** One of the most important factors affecting weight gain during pregnancy and its impact on the health of mother and infant is weight at the beginning of pregnancy. The best available measure of prepregnancy weight is body mass index (BMI), which has been incorporated in the new guidelines developed by the World Health Organization.

Epidemiological and experimental studies have shown that environmental influences early in life can “program” susceptibility to future disease, and that these effects can be transmitted to subsequent generations through nongenetic (epigenetic) mechanisms, with profound implications for human health. A mother’s (but not father’s) birthweight and weight and height at 20 months of age are positively associated with birthweight in the following generation.

Recent research has found that newborns who are small for their gestational age (SGA) or who have low birthweight (LBW) are at greater risk of central obesity, insulin resistance, metabolic syndrome, type 2 diabetes, hypertension, and coronary disease in the future. The association is accentuated by rapid weight gain in childhood. High birthweight has also been associated with later obesity and its consequences.

**Interpretation** The table on the following page shows the new recommendations for total weight gain during pregnancy, as well as recommended gains by trimester, as a function of prepregnancy weight:

BMI <sup>a</sup> before pregnancy		Recommended weight gain in pregnant women*		
Category	BMI (Kg/m <sup>2</sup> )	Over entire pregnancy, pounds (kilograms)	Trimesters 2 & 3, average: pounds/week. (range)	Over entire pregnancy in multiple pregnancies (pounds)
Low Weight	< 18.5	28 – 40	1 (1.0 – 1.3)	
Normal Weight	18.5 – < 25	25 – 35	1 (0.8 – 1.0)	37 – 45
Overweight	25.0 – 30.0	15 – 25	0,6 (0.5 – 0.7)	31 – 50
Obese (all categories)	≥ 30.0	11 – 20	0.5 (0.4 – 0.6)	25 – 42

Source: IOM (Institute of Medicine), 2009. Weight Gain During Pregnancy: Reexamining the Guidelines. Washington, DC: The National Academies Press. IOM, [www.iom.edu](http://www.iom.edu).

\* Calculation assumes a weight gain of 0.5-2 kg (1.1-4.4 lbs.) in the first trimester of pregnancy, based on Siega Riz et al., 1994; A, 1995; Carmichel et al., 1997.

<sup>a</sup> BMI: Body mass index; WHO classification of BMI (weight in kg / square of height in m). Obesity (BMI): class I, 30 - < 35; class II, 35 - < 40; class III, ≥ 40 kg/m<sup>2</sup>.

## Evidence

Achieving the recommended weight gain means more children with optimal birthweights (3500-4250 g) and fewer premature births, fewer newborns who are small for their gestational age (SGA) or large for their gestational age (LGA), and fewer with low birthweight (LBW) or suboptimal weight (2500-3500 g).

Neonatal and infant mortality is lower when gestational age at birth is 39-41 weeks and weight is between 3500 g and 4250 g.

Prepregnancy BMI and weight gain in pregnancy above or below the IOM recommendations are risk factors associated with unfavorable short- and long-term maternal, perinatal, neonatal, childhood and adult outcomes. The short-term outcomes include *perinatal and neonatal death*, *preterm birth*, newborns who are small or large for their gestational age, CMFs (of the neural tube and others), dystocia in delivery, hypertension, etc. Long-term outcomes include changes in adiposity and lean mass, with cardiac and metabolic consequences.

Low preconception BMI and weight gain during pregnancy below what is recommended by the IOM are each associated with greater risk of newborn with SGA newborn LBW and preterm birth, and the risk is multiplied when both are present. Similarly, preconception BMI and weight gain during pregnancy above the levels recommended by the IOM are associated with LGA and macrosomia among newborns.

Pregnant women who are obese or overweight should be instructed about the statistically significant increase in risk of complications such as maternal mortality, hypertension and gestational diabetes, preeclampsia, serious depressive disorders and less production of milk, heart and pulmonary diseases and obstructive sleep apnea, caesarean section, and venous thromboembolism, as well as risks for the fetus and newborn such as congenital malformations, macrosomia, LGA, obstetric trauma, miscarriage, and perinatal and neonatal mortality.

Obesity in pregnant women is associated directly or indirectly with a 52% increase in maternal mortality when BMI > 25, and with an increase of 27% when BMI ≥ 30.

There is strong evidence for an association between weight gain during pregnancy and the following outcomes: preterm birth, low birthweight, macrosomia, and newborns who are large or small for their gestational age. There also is evidence of increased adverse outcomes during labor and delivery (211-236).

## 7. Reducing unnecessary (medically not indicated) caesarean section

### Background

Births by caesarean section have increased progressively throughout the world. In the United States between 1996 and 2007, the incidence rose 65% (from 20.7% of deliveries to 32%). One of the reasons for the increase in preterm birth is the prevalence of medically unnecessary caesareans. Late preterm births (34-36 weeks of GA) account for approximately 75% of all preterm births, and the risk of death or complications in the neonatal period for these infants is 6 times as high as it is for full-term babies, while their risk during the first year of life is 3 times higher.

In a study in Utah (n= 343,322 late preterm births), late preterm SGA infants were 44 times as likely to die during the prenatal period, and 22 times as likely during the first year of life, as newborns of adequate gestational age.

A WHO study in Latin America in 120 of 410 randomly selected institutions in 24 geographical regions of 8 countries (n= 97,095 births) found the median rate of caesarean section to be 33%, with the highest rates (51-57%) in private hospitals. Of these deliveries, 49% were elective caesareans, 46% intrapartum operations, and 5% emergency caesareans in the absence of labor. In some countries, caesarean sections in private hospitals account for as much as 82% of births. The most frequent indication cited for caesarean section overall was previous caesarean 31% (16% in cases without complications in the current pregnancy and 15% with complications); in 26% of cases, the indication was cephalopelvic disproportion; in 20%, dystocia or arrested labor; in 11%, fetal stress, problematic breech presentation, or other reasons; and in 11%, preeclampsia/eclampsia.

### Interventions

#### Reducing caesarean section

Interventions that involve health workers in monitoring and feedback mechanisms, quality improvements, and multifaceted strategies proved effective in reducing rates at statistically significant levels (reductions of 20-30%). Identifying barriers to change was the most effective approach to reducing caesarean section.

Some interventions have demonstrated that they can diminish the rate of caesarean section, particularly those in which health workers analyze and modify their practices. This requires identifying the barriers to change that need to be addressed.

- ***Avoid elective vaginal and caesarean section births before the full 39 weeks of GA.*** Optimal gestational age at birth is 39-41 weeks, and optimal weight 3500-4250 g.
- Provide written guidelines with evidence-based indications for caesarean section in all facilities 24 hours a day, and make them available to all care staff.
- Establish and implement clinical standards.
- Hold weekly and monthly conferences to discuss the indications for caesarean section, in which a trained team provides analysis and feedback on successful interventions.
- Conduct periodic reviews of the records of 100 cases of labor and delivery, in order to evaluate the quality of care. Provide feedback through a final meeting, as well as a written summary of conclusions and recommendations.
- Promote vaginal birth for women who have had a caesarean.
- Hold conferences for feedback.
- Conduct monthly audits to review outcomes.
- Ensure ongoing community-based care, with emphasis on continuity provided by a team rather than relying on a single resource (midwives).

#### Interventions in labor that diminish the risk of caesarean section

- Continuous support during labor.
- Induction of labor in pregnancies without complications and more than 41 weeks of gestation.

- Partogram with monitoring of progress every 4 hours in full-term single pregnancies with spontaneous labor.
- Availability of obstetricians trained in vaginal birth procedures.
- Participation of consulting obstetricians in decisions to conduct caesarean section.

**Evidence**

Comparison of newborns delivered by elective caesarean at weeks 37 and 38 with those born at 39 weeks of GA shows that those born before 39 weeks are at statistically significant greater risk of neonatal death and morbidity (resuscitation, sepsis, RDS, and need for respiratory support (assisted ventilation), Apgar  $\leq$  6 at 5 minutes, admission to ICU > 5 days). These problems occur in 1 out of 5 neonates born at 37 weeks of gestation, 1 out of 8 born at 38 weeks, and 1 of 10 born at 39 weeks.

Comparison of children born by elective caesarean with those where vaginal birth is attempted shows that, at a gestational age of 36 weeks, respiratory distress syndrome (RDS) or hyaline membrane disease is 7 to 19 times more frequent among those delivered by elective caesarean, while it is between 6 and 32 times as frequent in cases of elective caesarean as it is in cases of full-term vaginal birth (37–38 weeks of GA).

The increase in preterm births in Latin America is due principally to unnecessary caesarean section, in part as a result of excessive medicalization and improper use of ultrasound (237–249).

## **8. Immunization with tetanus toxoid, taking previous vaccination status into account**

**Background:**

In 1989, the World Health Assembly adopted the goal of eliminating the transmission of neonatal tetanus by 1995. This target was later moved to 2005. Neonatal tetanus occurs throughout the world, but is most frequent in developing countries where maternity services are limited and tetanus vaccination coverage is low among women of childbearing age and pregnant women, and where unsafe practices are used in handling the newborn's umbilical stump.

**Infectious agent**

The tetanus bacillus (*Clostridium tetani*) is a gram-positive rod-shaped anaerobic microorganism and produces an endotoxin that affects the fetus.

**Transmission**

Infection is due to poor hygiene during delivery, when the umbilical cord is cut in unsanitary conditions, or when the umbilical stump is handled incorrectly (salves, dressings, plasters).

**Incubation**

The incubation period is 6 days, as an average, with a range of 3 to 28 days.

**Immunity**

In mothers, and in children from 6 weeks of age on, considerable immunity can be achieved with three doses of tetanus toxoid adsorbed, administered at least 4-week intervals (0.5 mL per dose, IM). Three doses of tetanus toxoid provide at least 5 years of immunity, while 5 doses provide lifetime immunity. Pregnant women are revaccinated only if the last dose was 10 years or more ago.

**Control**

The principal strategy of the program to eliminate neonatal tetanus is to immunize women of childbearing age with tetanus toxoid. This prevents both mothers and neonates from contracting the disease. If a mother has not been immunized completely, she should receive tetanus toxoid during pregnancy in order to complete the five doses.

**Evidence**

In populations where the incidence of tetanus is high, it can reduce neonatal mortality by 35–58% and reduce the incidence of neonatal tetanus by 88–100% (250–258).

## **9. Preventing influenza by vaccination**

**Background**

The Advisory Committee for Immunization Practices of the Centers for Disease Control and Prevention (CDC) recommends ordinary flu vaccination for all women who are or will be pregnant during the flu season. During seasonal flu epidemics, during previous pandemics, and with the current 2009 type A flu (H1N1) pandemic, pregnancy puts otherwise healthy women at greater risk for serious complications from influenza, including death.

**Safety**

The influenza vaccine for the seasonal virus or pandemic influenza, which is made from inactivated

viruses, can be administered safely and effectively during any trimester of pregnancy. No study to date has demonstrated major risk of maternal complications or adverse fetal outcomes associated with inactivated flu vaccine. Nor is there scientific evidence that vaccines with thiomersal produce adverse outcomes in children of women who received influenza vaccines during pregnancy.

When there is evidence that H1N1 is in circulation, the recommendation is to administer 75 mg twice daily for 5 days, beginning within 48 hours of onset of symptoms (fever, cough, sore throat, joint or muscular pains), without waiting for laboratory confirmation of influenza A H1N1-2009.

If the pregnant woman is at high risk for morbidity, hospitalization, and death from influenza A H1N1, it is important to vaccinate her as soon as inactivated H1N1-2009 influenza A and seasonal influenza vaccine is available, whatever trimester of pregnancy she is in.

#### Evidence

Clinical trials and data from the CDC show relatively high mortality in pregnant women from H1N1-2009 infection. The current recommendations call for initiating antiviral therapy in pregnant women as soon as possible, even before waiting for test findings confirming H1N1-2009.

Mortality in children from seasonal influenza in previous years was most frequent in the < 6-month age group, for which a vaccine is still not available. In the children of women vaccinated for influenza, 63% effectiveness has been achieved up to sixth months old, with a reduction of 29% in febrile respiratory illnesses in children, along with a 36% reduction in mothers (259–274).

## 10. Syphilis screening and case management

#### Background

Syphilis before delivery is a health problem in the developing countries, and the developed countries are seeing a trend of reappearance. All pregnant women should receive adequate prenatal care, including a compulsory syphilis test: the VDRL test or rapid plasma reagins (RPR). Infection by syphilis should be borne in mind where the mother's history or clinical signs in the fetus are suggestive, especially in cases of hepatosplenomegaly or fetal dropsy.

#### Infectious agent

*Treponema pallidum*

#### Transmission

The infection is transmitted from mother to child during fetal development or at birth. Almost half of all babies infected with syphilis in utero die shortly before or after birth.

#### Control

The principal strategy for eliminating congenital syphilis is compulsory case detection and management during pregnancy. Every pregnant woman should have two RPR examinations, VDRL tests, or fast treponemal tests in the first and third trimesters, but if this does not occur in the third trimester, it should be done before delivery or before discharge.

#### Treatment

Penicillin continues to be effective as a treatment for syphilis in pregnancy that prevents congenital syphilis. It consists of a single dose of benzathine penicillin, 2.4 million units IM, after ascertaining that the woman is not allergic to penicillin. If allergic, the patient should be desensitized and treated with penicillin if the physician is qualified to do so. Otherwise, she should be referred to a higher level of care.

If treatment is early in the pregnancy, it effectively prevents infection of the fetus. Even in women with long-standing syphilis, whose treatment would consist of three weekly doses of penicillin, a single dose of the drug will prevent fetal infection. The outcomes of these women's pregnancies are comparable to those of women without syphilis.

Counseling and voluntary HIV screening should be offered to all women with positive findings of syphilis.

#### Evidence

Reduction in fetal mortality, miscarriage, and perinatal mortality as a function of prevalence (275–292).

## 11. Detecting and treating symptomatic and asymptomatic bacteriuria

### Background

Urinary tract infections are a common complication during pregnancy. Asymptomatic bacteriuria occurs in 2%-10% of pregnancies. If it is not treated, over 30% of the mothers can develop pyelonephritis, and a series of complications affecting both mother and fetus can be triggered. A daily intake of 100 mg of ascorbic acid can reduce the incidence of bacteriuria.

### Diagnosis

Detection with reactive strips.

Urine culture (> 100,000 colonies/mL).

### Treatment

The antibiotic of choice is nitrofurantoin, 100 mg taken orally twice daily with food; other options if available: Cephalexin 500 mg taken orally twice daily; or cefadroxil 500 mg twice daily. Duration of treatment should be 7-10 days.

Persistent asymptomatic bacteriuria or asymptomatic bacteriuria that develops symptomatology, recurrent urinary tract infection (UTI) (i.e., three infections in the last 12 months or two in the last 6 months), and recurrent UTIs or concomitant renal lithiasis should be treated orally for 10-14 days with one of the following: nitrofurantoin, 100 mg 4 times daily; cephalexin, 1g every 6-8 hours; or cefadroxil, 1g twice daily. This should be followed by prophylaxis during the rest of the pregnancy in the form of 100 mg of nitrofurantoin, 125 mg of cephalexin, or 250 mg of cefaclor at bedtime.

### Evidence

Bacteriuria during pregnancy is strongly associated with premature birth, LBW, sepsis, and shock. Affected mothers may develop pyelonephritis, hypertension, or preeclampsia, and maternal and/or fetal death is a possibility (293-308).

## 12. Preventing and treating anemia with iron during pregnancy

### Background

Anemia is a lack of red blood cells or a reduction of their capacity to carry oxygen or iron. Some enzymes from tissues that require iron can be necessary for the functioning of nerve and muscle cells. The fetus depends on the blood of the mother, and anemia can lead to deficient fetal growth, premature birth, and low birthweight. The causes are associated with deficient iron intake, folate deficiency, hemorrhage (vaginal or caused by parasites) and, in some areas, malaria.

### Diagnosis

Hb < 11 g/dL.

### Treatment

120 mg of elemental iron daily, taken orally, for severe anemia (severe pallor or Hb < 7 g/dL) and 60 mg of elemental iron daily for moderate anemia (moderate pallor or Hb 7 - < 12 g/dL) or from week 12 of the pregnancy, throughout, and during the first three months postpartum.

### Evidence

Iron deficiency anemia in pregnant women causes maternal and perinatal mortality. It is also highly correlated with prematurity and LBW. Studies indicate that iron supplements during pregnancy improve perinatal outcomes (309-318).

## 13. Preventing preeclampsia and eclampsia with calcium (in populations with diets low in calcium)

### Background

Gestational hypertension is a very frequent complication, present in around 10% of pregnancies, and it affects perinatal outcome. In the Region of the Americas, it accounts for 25% of all maternal deaths. Through a series of mechanisms, calcium supplements can prevent hypertension and help prevent premature labor.

*High risk of preeclampsia:*

- Age of mother ≥ 40 years.
- History of previous preeclampsia, antiphospholipid antibodies, hypertension, renal disease, preexisting diabetes, BMI > 29 kg/m<sup>2</sup>, mother or sisters with history of preeclampsia.

- Current pregnancy: multiple pregnancy, birth interval  $\geq$  10 years, systolic blood pressure (SBP)  $\geq$  130 mmHg, or diastolic blood pressure (DBP)  $\geq$  80 mmHg. A DBP of 110 mm Hg or 100 mm Hg during the first 20 weeks of gestation has greater predictive value for preeclampsia when the pregnant woman has chronic hypertension. Periodontal disease and urinary tract infection are also risk factors.

Moderate risk of preeclampsia:

- Primipara, mild gestational hypertension (without proteinuria), multiple pregnancy, adolescent age, family history of severe preeclampsia.

Low risk for preeclampsia:

- Women who are without any pathology, do not take drugs or vitamins, and do not have the risk factors described above.

#### **Diagnosis**

One of the following indicators of risk:

- Severe headache.
- Blurred vision, loss of consciousness, and/or convulsions.
- Systolic blood pressure  $>$  140 mm/Hg and/or diastolic  $>$  90 mm/Hg.
- Significant proteinuria.
- Less than 3 portions of dairy products consumed daily.
- BMI  $<$  18.5 or  $>$  25.0 in the first trimester.

#### **Treatment**

For pregnant women at moderate or high risk for preeclampsia, 1.5 g/day, and if the patient's calcium intake is low, 2 g/day of elemental calcium orally, begun before 24 weeks of gestation – ideally before 20 weeks have passed – and continuing until birth.

#### **Evidence**

There is strong evidence that calcium supplementation benefits both women who are exposed to a high risk of gestational hypertension, and those belonging to communities with low calcium intake. Fewer children are born weighing  $<$  2500 g to women treated with calcium (319–336).

## **14. Preventing preeclampsia and eclampsia with aspirin**

#### **Background**

Preeclampsia is associated with deficient production of the vasodilator prostacyclin, and with excessive production of thromboxane, which is a vasoconstrictor and stimulates the aggregation of blood platelets. Antiplatelet agents such as aspirin in low doses can prevent or delay the appearance of preeclampsia.

#### **Diagnosis**

One of the following signs:

- Severe headache.
- Blurred vision, loss of consciousness, and/or convulsions.
- Systolic blood pressure  $>$  140 mm/Hg and/or diastolic  $>$  90 mm/Hg.
- Less than 3 portions of dairy products consumed daily.
- BMI  $<$  18.5 or  $>$  25.0 in the first trimester.

#### **Treatment**

Acetylsalicylic (aspirin): for pregnant women at moderate risk for preeclampsia, 75–100 mg/day; for those at high risk, up to 100–150 mg/day orally at bedtime, beginning in week 12 or 13 of gestation and continuing until birth.

#### **Evidence**

This treatment reduces preeclampsia in high-risk groups by 17%, and up to 36% if the dose taken is over 75 mg of aspirin/day. It is associated with an 8% reduction in preterm births, a 14% reduction in perinatal death, and a 10% reduction in IUGR (337–346).

## 15. Detecting and treating sexually transmitted infections

<b>Background</b>	Pregnant women with STIs such as chlamydia or gonorrhea at the end of the pregnancy can transmit it to the baby, which can cause problems at birth (ophthalmia neonatorum). Children of mothers who have active genital herpes infections during or near the time of delivery can be infected. This can be serious, and sometimes fatal, to the newborn. Vertical transmission of HIV/AIDS can be reduced considerably by the appropriate measures.
<b>Diagnosis</b>	Chlamydia ( <i>Chlamydia tracomatis</i> ), gonorrhea ( <i>Neisseria gonorrhoeae</i> ), vaginosis, herpes virus, HIV/AIDS, hepatitis B, syphilis ( <i>Treponema pallidum</i> ).
<b>Evidence</b>	Diseases like chlamydia, gonorrhea, syphilis, trichomoniasis, and bacterial vaginosis can be treated and cured with antibiotics during pregnancy. There is no cure for viral STIs such as genital herpes and HIV, but antiviral drugs can reduce symptoms in pregnant women. If a woman has active sores from genital herpes, HIV, or hepatitis B at the time of the delivery, a caesarean can be performed to protect the newborn against the infection, and treatment can be provided for mother and newborn. Women whose hepatitis B screening was negative can receive hepatitis B vaccine while pregnant (347–363).

## 16. Antiretrovirals and other measures to reduce the risk of vertical transmission of HIV

<b>Background</b>	Antiretroviral drugs (ARVs) reduce viral replication and can cut down mother-to-child transmission of HIV by diminishing the pregnant woman's plasma viral load. Alternatively, they can be used for postexposure prophylaxis in the newborn. In the absence of intervention, HIV infection is transmitted from mother to child in as many as 48% of cases.
<b>Treatment</b>	Zidovudine: <i>During pregnancy:</i> 100 mg orally 5 times a day, beginning between weeks 14 and 34, and continuing throughout pregnancy. During an elective caesarean section: 2 mg/kg IV followed by continuous infusion of 1 mg/kg/hour until delivery. <i>After elective caesarean section:</i> 2 mg/kg of syrup to the newborn every 6 hours for 6 weeks, beginning 8–12 hours after the delivery.
<b>Evidence</b>	The use of antiretrovirals significantly reduces vertical transmission of HIV from mother to child, as do other interventions such as elective caesarean and formula in place of mother's milk for all babies exposed (364–369).

## 17. Detecting and treating diabetes

<b>Background</b>	Hyperglycemia and inadequate weight gain during pregnancy are associated with adverse outcomes for fetuses and neonates, increasing the possibility of neonatal hypoglycemia, macrosomia, and birth defects. Gestational diabetes can occur even when there are no symptoms or risk factors. For this reason, pregnant women are often advised to eat a healthy diet and avoid a sedentary lifestyle. Gestational diabetes usually disappears after birth. However, women who have had gestational diabetes run a greater risk of developing diabetes many years later.
<b>Diagnosis</b>	A family history of diabetes  Fasting blood glucose greater than 105 mg/dL  Overweight: BMI 25.0–29.9  Obesity: BMI ≥ 30.0
<b>Evidence</b>	Diabetes during pregnancy is associated with fetal death, with macrosomia at birth, and with trauma, respiratory problems, hypoglycemia, hyperbilirubinemia, and metabolic problems. There is strong evidence of an association between birth defects and poor control of blood glucose levels during pregnancy (370–389).

## 18. Deparasitization in areas of high prevalence using albendazole

<b>Background</b>	In areas of endemic intestinal helminthiasis, routine therapy with albendazole considerably reduces the prevalence of anemia in pregnant women, as well as its possible complications.
<b>Diagnosis</b>	Stool examination (for ascaris, oxyurids, uncinaria, trichuris).
<b>Treatment</b>	Albendazole, 400 mg orally, single dose, in both the 2nd and 3rd quarters
<b>Evidence</b>	Deparasitization during pregnancy reduces maternal anemia and its complications, as well as low birthweight, and there is no evidence that albendazole has teratogenic effects on the fetus (390–395).

## 19. Detecting and treating group B streptococcus

<b>Background</b>	Group B streptococcus ( <i>Streptococcus agalactiae</i> ) is a gram-positive bacterium that lives in the vaginal or rectal areas of between 10% and 35% of healthy adult women. A woman with group B streptococcus in her vagina or rectum is said to be “colonized” by the organism. For every 100 women giving birth who are colonized with group B streptococcus, 1-2 infants are infected with it at the time of delivery, and can become ill or die from sepsis.
<b>Diagnosis</b>	Rectal and vaginal culture between weeks 35 and 37 of gestation.
<b>Prophylaxis</b>	All women with one or more of the following risk factors should receive intrapartum prophylaxis: delivery at < 37 weeks of gestation, intrapartum temperature > 38° C, or rupture of membranes ≥ 8 hours ago.
<b>Treatment</b>	Penicillin G: 5 million units IV (initial dose), then 2.5 million units IV every 4 hours up to delivery; or ampicillin, 2g IV (initial dose), then 1g IV every 6 hours for 48 hours. If the woman is allergic to penicillin, give cefazolin 2g IV (initial dose), then 1g IV every 8 hours up to delivery.
<b>Evidence</b>	Prophylactic therapy begun at least 4 hours before delivery reduces the incidence of early infection by 86–89%, and cuts neonatal mortality due to group B streptococcus (396–408).

## 20. Nifedipine during preterm labor

<b>Background</b>	Premature birth is one of the greatest causes of morbidity and mortality in the developing countries, representing between 11% and 21% of all births. Different tocolytics have been used to prevent it. The use of magnesium sulfate as a tocolytic has been shown not to affect the risk of birth within 48 hours of treatment, while those exposed to it are at greater risk of fetal, neonatal, and postneonatal death. It has recently been demonstrated that nifedipine is an effective, safe, and low-cost tocolytic drug to prevent preterm birth.
<b>Diagnosis</b>	Signs of labor before 37 weeks of gestation.
<b>Treatment</b>	<i>Initial dose:</i> Oral nifedipine, 10 mg. If uterine activity continues, administer 10 mg every 20 minutes orally 2 more times more, for a total of 3 doses. <i>Maintenance dose:</i> Nifedipine, 10–20 mg orally every 4–8 hours, depending on how the pregnant woman responds, for 2–3 days. <i>If the pregnant woman's blood pressure is below 110/70 mm of Hg, do not use nifedipine.</i>
<b>Evidence</b>	Nifedipine has proven more effective and safer than other tocolytics to address the risk of preterm delivery, and is very beneficial given orally. It prevents RDS, intraventricular hemorrhage, and jaundice (409–417).

## 21. Prenatal corticosteroids (betamethasone, dexamethasone) to foster lung maturation

<b>Background</b>	Premature birth is statistically the major factor in neonatal and infant mortality. Respiratory distress syndrome (hyaline membrane disease) is a severe complication of prematurity, and is the principal cause of neonatal morbidity and mortality in the countries. In order to encourage lung maturation in births taking place up to 35 weeks of gestation, steroids have been used with very good results.
-------------------	--

<b>Diagnosis</b>	Signs of labor or threat of preterm delivery (PTD) between weeks 26 and 36 weeks of gestation. Act if there are suspicions or signs of PTD. Do not wait to confirm the diagnosis, and <i>always treat, unless birth is imminent (within 60 minutes)</i> .
<b>Treatment</b>	Betamethasone: 12 mg IM every 24 hours for 2 days  Dexamethasone: 6 mg IM every 12 hours for 4 days
<b>Evidence</b>	The administration of betamethasone or dexamethasone before birth has been shown to significantly diminish RDS by 36%-50%, and neonatal mortality by 37%-40%, without side effects for surviving babies. Neonates whose gestational age is 32 weeks and whose mothers have completed a prenatal course of steroids experience an 82% drop in mortality with the treatment. It also diminishes the risk of intraventricular hemorrhage by 40%-70%, as well as patent <i>ductus arteriosus</i> , and the evidence suggests that it can also protect against neurological sequelae. A single dose provides more benefit than multiple doses. Multiple doses at intervals of 14 days do not improve preterm birth outcomes, and are associated with reduced weight, height and cephalic circumference at birth.  A recent meta-analysis of 44 of 1412 initially selected studies found that the impact of prenatal steroids in reducing mortality is greater in middle-income countries than in higher-income ones. Neonatal mortality was significantly ( $p=0.008$ ) reduced – between 53% and 65% in middle-income countries, and between 31% and 35% in high-income countries – while morbidity fell by 37%-51% in preterm babies with less than 36 weeks of gestation. Treatment between weeks 31 and 36 of gestation reduced neonatal mortality by over 30% in babies born before 30 weeks. The effect was small, and after 36 weeks there is no evidence that it reduces neonatal mortality (418-441).

## 22. Detecting, preventing, and treating Rh isoimmunization

<b>Background</b>	Rh isoimmunization is an autoimmune disease that originates in the perinatal period and is characterized by fetal hemolytic anemia as a result of incompatibility between the mother's and fetus's blood types. This occurs when an RH-negative woman is carrying an Rh-positive fetus and has proven capable of forming antibodies in response to the fetus's Rh factor antigen. Initial sensitization can occur not only because of pregnancy, but in cases of miscarriage, ectopic pregnancy, amniocentesis, etc. The severity of the condition increases with subsequent pregnancies, and each succeeding baby is more likely to be affected.
<b>Diagnosis</b>	Diagnosis of this intrauterine disease is based on an indirect Coombs test in Rh-negative women to ascertain whether the Rh factor is present at the beginning of the pregnancy. The test tells whether a woman already has Rh antibodies. Periodic titration of antibodies during the pregnancy makes it possible to monitor their level.
<b>Amniocentesis</b>	to assess the amount of bilirubin in the amniotic fluid should be conducted in cases where the indirect Coombs titres increase throughout the pregnancy, or when sonography shows that ascites or fetal hepatosplenomegaly has made an appearance.  In special cases, cordocentesis is needed to provide blood directly from the umbilical cord in order to evaluate the degree of fetal anemia.
	Diagnosis in the neonate is confirmed by a direct Coombs test, which actually shows the presence of the antibodies.  Currently, Doppler ultrasound measurement of the peak systolic velocity of blood in the fetus's cerebral artery is a quite sensitive and specific non-invasive tool to draw on in managing isoimmunization during pregnancy.
<b>Prophylaxis</b>	In all cases of nonsensitized Rh-negative women who have had an RH-positive child, administer a 300-microgram IM or IV dose of anti-D gammaglobulin within 72 hours after delivery.

All nonsensitized pregnant women who are Rh-negative should routinely be given two 120-microgram IM or IV doses of gammaglobulin anti-D at 28 and 34 weeks of gestation.

After a miscarriage, spontaneous or induced abortion, or ectopic pregnancy during the first 12 weeks of gestation, nonsensitized Rh-negative women should be given a minimum of 120 micrograms of anti-D gammaglobulin IM or IV. If more than 12 weeks of gestation have passed, 300 micrograms should be administered.

After invasive procedures such as amniocentesis or cordocentesis in a nonsensitized Rh-negative pregnant woman, administer 300 micrograms of anti-D gammaglobulin IM or IV.

<b>Evidence</b>	The consequences of Rh isoimmunization for the fetus include ascites, edema, hydrops, and intrauterine death, while consequences for the newborn include anemia, edema, severe jaundice, kernicterus, pleural effusion, cardiac insufficiency, and death (442–456).
-----------------	---

## 23. Detecting and treating periodontal disease

<b>Background</b>	Poor oral health during pregnancy has been linked to greater perinatal and neonatal morbidity.
<b>Diagnosis</b>	Dental check-up (twice during pregnancy).
<b>Evidence</b>	Periodontal disease in pregnancy is associated with preeclampsia, and in newborns it is associated with prematurity, LBW, and IUGR (457–468).

## 24. Detecting, preventing, and managing domestic violence and abuse, and depression

<b>Background</b>	Direct violence against a pregnant woman and her fetus by her partner usually goes unrecognized by health professionals, or it is suspected but not dealt with, because of the multiple repercussions involved. This type of violence occurs in all social, economic, educational, and professional strata. In any form (intimidation, aggression, abuse, etc.) it requires careful evaluation and quick, efficient intervention in order to protect both mother and fetus from trauma, miscarriage, and death.
<b>Diagnosis</b>	Repeated injury, vaginal bleeding, a threat of abortion, low self-esteem, depression, personality alterations, neglect of personal hygiene.
<b>Evidence</b>	Violence diminishes motivation and morale, produces physical and psychological injury, depression, and post-traumatic stress, and provokes preterm delivery, as well as fetal and/or maternal death. One out of 7 women (14%) may develop depression during pregnancy or after delivery, the incidence being greater in women under 30 (468–479).

## 25. Detecting and preventing exposure to tobacco

<b>Background</b>	Consumption of or exposure to tobacco during gestation is known to increase the probability of miscarriage, perinatal mortality, and smaller weight gain in newborns. Inhalation of tobacco smoke with the gamut of toxic substances that it contains has an enormous effect on the conditions under which the fetus develops during the gestation.
	Nicotine induces acute hypoxemia (lack of oxygen) by reducing blood flow in the placenta, but the effect is transitory in the fetus's blood. Carbon monoxide, on the other hand, produces chronic hypoxia in the fetus, since it increases the level of carboxyhemoglobin (COHb) in the blood of both mother and fetus, thus reducing oxygen transport and its release in the fetus's tissues and organs, producing hypoxia and affecting the fetus's development.
<b>Prevention</b>	There is no consensus on the minimum quantity of tobacco consumption that can create problems in the newborn. Thus, all tobacco consumption should be avoided throughout pregnancy.
<b>Evidence</b>	Smoking by the mother, as well as exposure to tobacco smoke, is strongly associated with low birthweight,

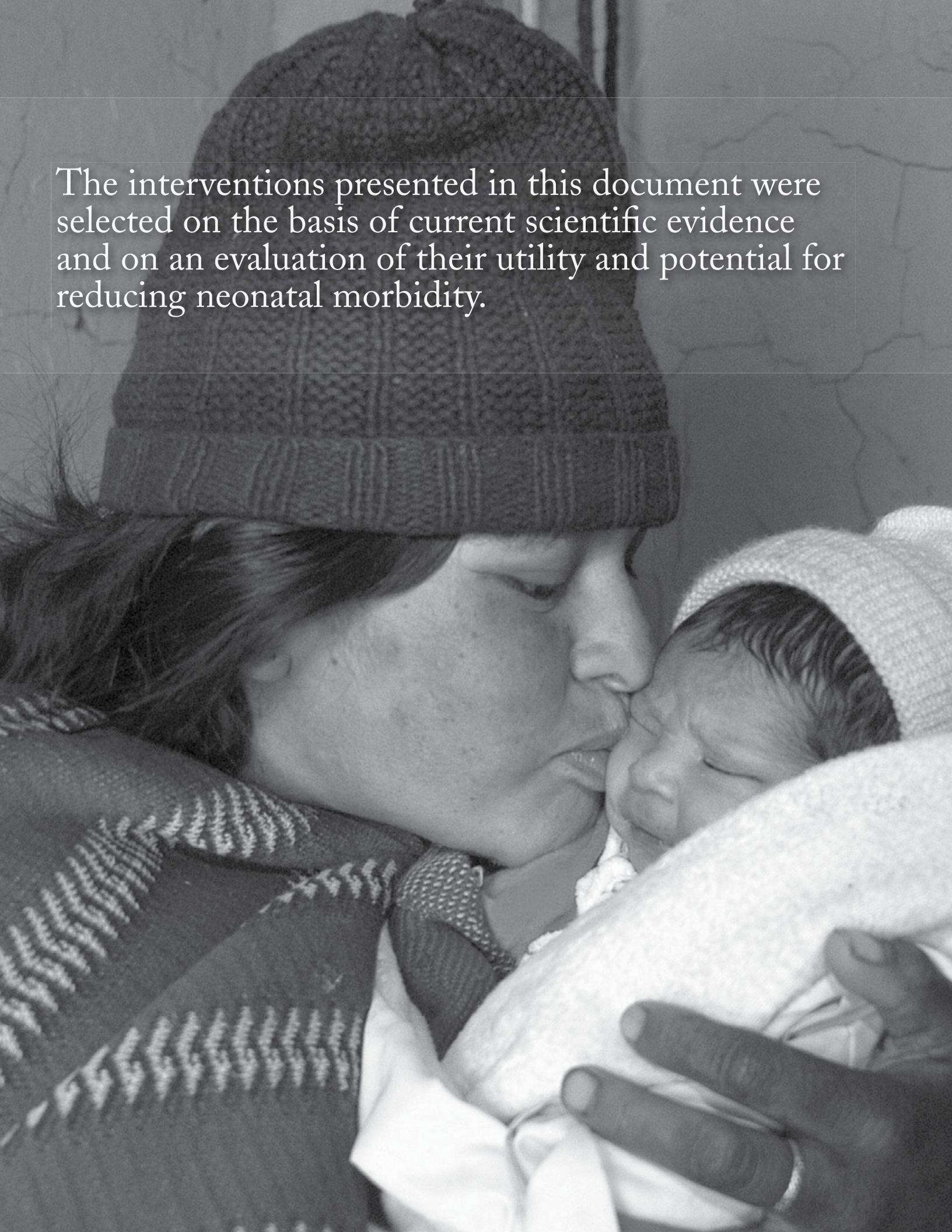
intrauterine growth restriction and sudden infant death syndrome. It has also been shown to lead to higher probabilities of miscarriage, premature birth, and more numerous complications during pregnancy and childbirth (480–493).

## **26. Detecting and preventing exposure to alcohol**

<b>Background</b>	Alcohol consumption during pregnancy is a serious public health problem, and a cause of birth defects and retarded development – a pattern known as fetal alcohol syndrome (FAS). FAS reflects congenital damage from alcohol and causes mental retardation, facial anomalies, and deficient growth. Affected newborns go on to develop cognitive deficiencies, learning and memory disorders, problem-solving difficulties, and attention deficit, as well as mental health problems and difficulties with social interaction. However, some children, while lacking the characteristic facial defects and not suffering from deficient growth, do suffer from an alcohol-related neurological development disorder.
<b>Prevention</b>	There is no consensus on the minimum quantity of alcohol consumption that can cause problems in the newborn. Thus, all alcohol consumption should be avoided throughout pregnancy.
<b>Evidence</b>	Alcohol consumption by a mother during pregnancy can cause severe birth defects, of which fetal alcohol syndrome (FAS) is the most devastating (494–505).

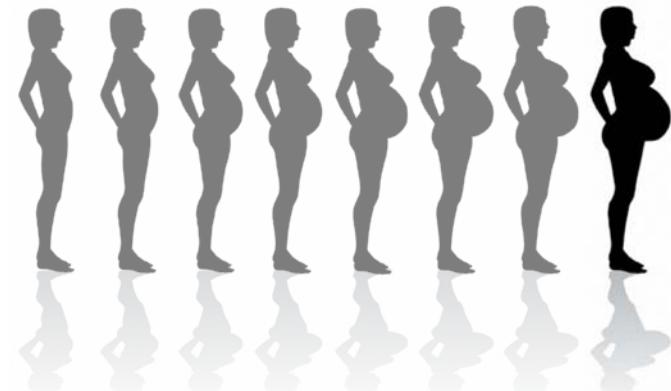
## **27. Detecting and preventing exposure to recreational or illicit drugs**

<b>Background</b>	Recreational drugs that, when introduced into the organism, alter physical and psychological balance and can produce dependency or stimulate the nervous system, modify one or more aspects of its functioning, or change an individual's normal behavior.
	Maternal drug use has various effects on the fetus. It can alter organogenesis and morphogenesis, fetal growth, lead to various malformations, cause developmental alterations, and produce a broad range of neurological pathologies. All drugs pass from the mother's bloodstream to the fetus through the placenta. Illicit substances that induce drug dependency and addiction in the mother also cause dependency in the fetus. At birth, the baby's dependency on the substance continues, but since the drug is no longer available, the infant's central nervous system becomes hyperstimulated, and withdrawal symptoms appear.
<b>Diagnosis</b>	Threatened miscarriage or premature birth, low self-esteem, depression, personality alterations, neglect of personal hygiene, or references to consuming toxic substances.
<b>Evidence</b>	Heroin and other opiates, including methadone, can induce a severe withdrawal syndrome in the baby, some symptoms lasting from four to six months. Convulsions can also occur in the babies of mothers who take methadone. The prenatal use of amphetamines is associated with low birthweight, premature birth, and intracranial hemorrhage. The use of cocaine is associated with a greater risk of sudden infant death syndrome (SIDS). Marijuana consumption is associated with lower birthweight and smaller size at birth (IUGR). Long-term effects reported in children exposed to prenatal drugs include increased risk of cognitive deficits, lower IQ, lower scores in short-term memory and verbal reasoning, as well as a need for further-reaching psychological intervention during childhood and adolescence (506–521).



The interventions presented in this document were selected on the basis of current scientific evidence and on an evaluation of their utility and potential for reducing neonatal morbidity.

# III. Interventions during delivery



## 1. Clean and safe delivery practices (with skilled personnel)

<b>Background</b>	Access to skilled care by personnel at a health care facility during pregnancy, childbirth and puerperium is critical to ensuring the health of mothers and newborns.
<b>Evidence</b>	Deliveries assisted by skilled personnel in high-quality health facilities have fewer complications and less perinatal and neonatal morbidity and mortality (522-525).

## 2. Accompanying the mother during labor and delivery

<b>Background</b>	Birth is not a disease. Delivery is a normal physiological process with emotional aspects that are fundamental and must be respected. The presence of a family member or other trusted person (doula) during pregnancy, childbirth, and postpartum is a woman's right. It provides her physical and emotional support, as well as helping to keep her properly informed.
<b>Evidence</b>	The continuous presence of a support person during labor and delivery reduces the duration of labor, the need for caesarean section or instruments to assist delivery, the need for pain medication, and the incidence of newborns with Apgar scores < 7 at 5 minutes (526-532).

## 3. Erythromycin in PROM before delivery in the first 37 weeks of gestation

<b>Background</b>	One factor associated with premature rupture of membranes (PROM) before delivery in the first 37 weeks of gestation, or close to term, is infection in the woman and/or fetus. Treating the mother prophylactically with antibiotics can reduce this risk. Bacterial resistance to penicillin and the fact that amoxicillin/clavulanate creates a risk of enterocolitis have led to the use of erythromycin, and with good results.
<b>Diagnosis</b>	PROM before labor, and clear suspicions of chorioamnionitis.
<b>Treatment</b>	Erythromycin, 250 mg orally, every 6 hours for 10 days.
<b>Evidence</b>	Statistically significant reduction of: chorioamnionitis; neonatal morbidity, including infection (pneumonia); use of surfactant; use of oxygen; and abnormality in cerebral ultrasound tests conducted before discharge (533-539).

#### 4. Monitoring labor with partograms

**Background** A partogram is a graphic visual representation of the measurements and events related to the course of the labor. The relevant measurements included in a partogram can include statistics such as cervical dilation over time, fetal heart rate, and the mother's vital signs. Clinical monitoring of the course of labor can prevent, detect, and facilitate managing the appearance of complications that can do damage that is sometimes irreversible or fatal to the mother and neonate.

**Evidence** The use of partograms reduces unnecessary interventions and perinatal complications (540–548).

#### 5. Caesarean section in breech presentation with LBW infants

**Background** The preferred route for the birth of neonates with LBW or who are premature, in cases of breech presentation, has been a matter of controversy. However, it has been demonstrated that, in practice, caesarean section diminishes perinatal complications.

**Treatment** Elective caesarean section.

**Evidence** Reduced neonatal morbidity and mortality, less asphyxiation and birth trauma than with vaginal deliveries of newborns with LBW (549–554).

#### 6. Antibiotic prophylaxis in caesarean section

**Background** Caesarean section is currently one of the most common operations in hospitals. The four most frequent indications are previous caesarean section, dystocia or arrested labor, breech presentation, and fetal stress. The most frequent type of complication related to caesarean section is infection. Without prophylactic antibiotics, endometritis occurs in almost 40% of caesarean sections where risk factors are present.

**Treatment** Cefalotin, single dose, 2 grams IV, during the caesarean section.

**Evidence** The use of prophylactic antibiotics in elective or non-elective caesarean section effectively reduces the incidence of endometritis and severe sequelae such as abscesses, septic shock, and septic thrombophlebitis (555–561).

#### 7. Modified active management of the third stage of labor

**Background** Birth extends from the emergence of the baby until the placenta is expelled. It is a period of great risk and remains the leading cause of maternal death throughout the world because of hemorrhage.

**Treatment** Oxytocine, 5-10 IU, IM, or

Methylergonovine, 0.5 mg, IM, or

Misoprostol, 0.4-0.8 mg, orally or sublingually,

+ Umbilical cord traction and uterine contraction + abdominal uterine massage

**Evidence** All of these have been shown to reduce bleeding and the number of transfusions for the mother by 60%. All increase the risk of placental retention, as well as anemia in the newborn, though oxytocics double the frequency of placental retention. Additional advantages of misoprostol are a longer half-life, stability at room temperature, and the possibility of oral administration. Oxytocine plus misoprostol can reduce postpartum hemorrhage after caesarean section (562–580).

#### 8. Late clamping of the umbilical cord

**Background** The practice of clamping the cord early, before placental transfusion ceases, or within 10-15 seconds after birth is frequent in health care facilities. It is not supported by scientific evidence and is harmful to the baby.

**Evidence**

Late clamping of the umbilical cord – when it stops pulsing (between 2 and 3 minutes) – is physiologically sound [TN: original simply says “es fisiológico”] and increases neonatal hematocrit by up to 50% over levels found with early clamping. It increases the newborn’s iron reserves, and reduces the prevalence of anemia in the first 4 to 6 months of life. It also improves cerebral oxygenation in premature newborns during the first 24 hours of life. A recent report states that late clamping of the cord facilitates the first transfusion of hematopoietic stem cells, precursors of endothelial cells, mesenchymal precursor cells, and pluripotent stem cells. (581-592).

WHO and UNICEF recommend at least two home visits to mothers and newborns: the first within 24 hours after birth (day 1) and a second on day 3, and if possible, a third after the first week of life (day 7).



# IV. Maternal interventions after delivery

## 1. Early skin-to-skin contact

**Background** Early contact between mother and baby fosters bonding. The greater the baby's social integration with its mother is, the stronger its bond with her will be. The first hours of life are the critical period for successful bonding. The quality of early relations is crucial for development.

**Intervention** Place the newborn in skin-to-skin contact with its mother within the first half hour after delivery.

**Evidence** This intervention fosters the beginning and continuance of exclusive breastfeeding, early bonding between mother and newborn, stability of temperature, cardiorespiratory variables, and crying patterns. No negative effects, either short- or long-term, have been reported (593-602).

## 2. Mother-newborn rooming-in

**Background** The formation of affective ties is a vital human need. Such ties provide the basis for the individual's psychological and social development. From birth, a human baby depends on an adult for survival, and the bond formed with the adult becomes the means by which the young person achieves a unique and specific sense of itself, thus acquiring the sense of security needed to become progressively more autonomous and independent.

**Intervention** The newborn remains beside the mother 24 hours a day.

**Evidence** Mother-newborn rooming-in increases the mother's confidence and self-esteem, and fosters the development of her skills. It facilitates bonding and the beginning and continuance of breastfeeding, as well as the parents' identity as parents, and their knowledge of their child. Finally, it gives the parents the peace of mind and enthusiasm that they need when they leave the care facility (603-608).

## 3. Detecting and managing puerperal endometritis

**Background** This is the most common infectious process during the puerperium. The endometrium is infected by germs that enter through the wound remaining in the area of the endometrium where the placenta was inserted, and from there the infection spreads to the entire endometrial mucous membrane. Endometritis can be caused by a single microorganism, but is most commonly caused by several. Only in one third of cases can the causal organism be identified.

**Diagnosis** Fever of over 38° C in the mother on the third or fourth day of puerperium, with tachycardia and chills. The lochia become thick, purulent-looking or chocolatey, and fetid.

**Risk factors** Primiparas and multiparas, premature rupture of membranes more than 6 hours before delivery, caesarean section, prolonged labor, induced delivery, preeclampsia, preterm delivery, anemia, lack of prenatal check-up, sexually transmitted infections, and low socioeconomic stratum.

**Treatment** The combination of gentamicin + clindamycin would seem an appropriate treatment, but susceptibility to antibiotics in the particular country will play a role in the decision.

**Evidence** The detection and timely management of puerperal endometritis is associated with less morbidity and maternal mortality (609–613).

## 4. Detecting, preventing, and managing postpartum depression

**Background** Postpartum depression is one of the most frequent postpartum diseases, affecting 1 out of 8 women giving birth. If not treated properly, it can persist for months and even years. Postpartum depression is present when the mother develops depression after the birth of a child. Sometimes, the phenomenon can be explained by the fact that the child was not wanted, or is not normal. In most cases, however, there is no explanation for the depression.

**Diagnosis** At any time within the first three months after delivery, a woman may present with:

- Difficulty going to sleep, or sleeping too much
- Lack of interest in usual activities or pastimes
- Feelings of guilt
- Loss of energy
- Difficulty concentrating or making decisions
- Changes in appetite or eating
- Feeling tired all the time
- Thoughts of suicide or death
- Physical violence and death of the child(ren)
- Personal or family history of depression
- An undesired or unplanned pregnancy
- Marital or financial problems
- Significant life changes, such as the death of a relative or a change of jobs
- Medical problems, or complications with the pregnancy
- Miscarriage in a previous pregnancy

**Treatment** The common antidepressants for postpartum use are the selective serotonin inhibitors such as sertraline, paroxetine, citalopram, and fluoxetine. If the mother is breastfeeding, sertraline chlorhydrate in a dose of 50 mg/day orally is the drug of choice, since it passes through to the milk in lower quantities.

Some women with mild to moderate depression can be treated with support groups. Severe depression requires psychotherapy.

**Evidence** In the mother: feelings of guilt, frustration, lack of interest, neglect of baby. In the newborn: greater morbidity and mortality, more prolonged colic and crying (614–628).

# V. Neonatal interventions after delivery

Immediate neonatal adaptation is the set of cardiac, hemodynamic, respiratory, and other changes on which an adequate transition from intrauterine life to life in the outside environment depends. The majority of newborns make this transition without complications, but others require some immediate intervention, such as neonatal resuscitation. Those that survive healthy depend on other basic interventions to prevent complications like neonatal ophthalmia, omphalitis, skin infection, or hemorrhagic disease of the newborn.

Beginning and continuing exclusive breastfeeding is one of the principal cost-effective interventions to reduce morbidity and mortality, both neonatal and in childhood. Premature or low birthweight newborns can be cared for with the kangaroo mother technique, which fosters bonding and breastfeeding and reduces postnatal complications.

Other interventions – prevention of hypothermia or hypoglycemia, and hand washing to prevent infection – can dramatically reduce neonatal mortality simply and at low cost. Although some newborns will need more sophisticated and expensive interventions – surfactant, mechanical ventilation, antibiotics for sepsis, for example – adequate prenatal intervention can considerably reduce morbidity.

## 1. Neonatal resuscitation with ambient air

**Background** Asphyxiation at birth is one of the leading causes of neonatal mortality and long-term neurological sequelae. In Latin America and the Caribbean, more than 50,000 children die every year from asphyxiation, accounting for 29% of all deaths before 28 days of age. Immediate and effective neonatal resuscitation can prevent the vast majority of these deaths. However, we often lack the proper equipment and oxygen supply where it is needed. Positive-pressure bag and mask ventilation can be used at facilities that lack staff trained in sophisticated resuscitation techniques (such as ET intubation, cardiac massage, drugs). It has been suggested that resuscitation can be as effective with ambient air as with 100% oxygen. If available, pulse oximetry should be used, with the sensor attached to the right-side upper limb (the wrist or palm of the hand), in order to evaluate the need for additional oxygen. In the case of neonates born at term, it is best to begin resuscitation with oxygen in ambient air (21%), and proceed to 100% oxygen if bradycardia is present after 90 seconds. The administration of supplemental oxygen should be regulated by mixing oxygen and air.

**Evidence** Ambient air is as good as 100% oxygen for the resuscitation of asphyxiated newborns, reducing both mortality and neurological sequelae (629–636).

## 2. Immediate care for the healthy newborn

<b>Background</b>	At the world level, 90%-95% of children are born in good shape and do not require any resuscitation, but only immediate care: preventing heat loss, attending properly to the umbilical cord (with a sterile/clean cut and ligature), beginning breastfeeding in the first hour, and avoiding any separation from the mother.
<b>Diagnosis</b>	Breathing or crying Good muscle tone
<b>Treatment</b>	<i>At birth:</i> <ul style="list-style-type: none"> <li>■ Maintain room temperature in the delivery room between 28° C and 30° C.</li> <li>■ Dry and cover.</li> <li>■ Aspiration of the mouth or nose should not be routine (but only used when necessary to remove obstructions or meconium).</li> <li>■ When it stops pulsing, the umbilical cord is closed with umbilical tape or a clamp 2-3 cm from the base, and cut with sterile scissors.</li> <li>■ Apply 1 mg of vitamin K1 intramuscularly.</li> <li>■ Administer eye infection prophylaxis in the form of silver nitrate (1%), tetracycline (1%), or erythromycin.</li> <li>■ Put an identification bracelet on the newborn.</li> <li>■ Keep the newborn in skin-to-skin contact with its mother from birth, and begin breastfeeding within an hour of birth.</li> <li>■ Erythromycin (0.5%)</li> </ul>
<b>Evidence</b>	Routine immediate care for healthy newborns prevents hypothermia, hypoglycemia, anemia, hemorrhagic disease of the newborn, eye infection, switching or loss of a newborn, and a delayed start to exclusive breastfeeding (637-651).

## 3. Care of the umbilical cord

<b>Background</b>	Omphalitis is one of the leading causes of sepsis in the neonatal period, often secondary to traditional treatment of the umbilical cord, to the way of cutting it and instruments used for the purpose, and to the application of dressings with creams, powders, lotions, and even some herbs. Omphalitis contributes to neonatal morbidity and mortality in developing countries. Various means are used to reduce omphalitis infection and mortality: simply keeping the umbilical cord dry, and using antiseptics, antibacterial agents, or alcohol.
<b>Treatment</b>	Use an antiseptic solution initially (triple stain or alcohol), and then keep clean and dry.
<b>Evidence</b>	Antiseptic solutions such as triple stain (Tween 80, gentian violet, and proflavine sulfate) or alcohol have been effective, although chlorhexidine or povidone-iodine can also be used. A case-control study in Nepal showed that the use of chlorhexidine on the umbilical stump reduced neonatal mortality by 24%, and up to 34% if applied during the first 24 hours, as compared with simple clean-and-dry management of the cord. The traditional treatment of the umbilical cord with breast milk seems not to have adverse effects, and is associated with the cord falling off more quickly. The use of alcohol or chlorhexidine delays the event by several days. During omphalitis epidemics, triple stain has been a more effective preventive agent, but can delay the cord's falling off (652-662).

## 4. Skin care at birth

<b>Background</b>	The surface of the newborn's skin, the vernix, and the amniotic fluid are full of substances that help protect against bacterial invasion.
-------------------	--

<b>Treatment</b>	The following are recommended for the newborn's bodily hygiene:
	<ul style="list-style-type: none"> <li>■ Avoid immediate bathing of preterm or low birthweight babies.</li> <li>■ Avoid removing the vernix from the skin.</li> <li>■ Full-term and stable newborns can be bathed with a sponge, but not by immersion, and care should be taken not to wet the umbilical cord.</li> <li>■ The scalp should be cleaned carefully, as should excess blood.</li> <li>■ If covered with blood or amniotic fluid, the baby can be cleaned with cotton impregnated with sterile water and a non-antiseptic neutral soap.</li> <li>■ If a fetid odor is present, or amniotitis is suspected, use soap to clean.</li> <li>■ Careful hygiene of ears, nostrils, and places where blood has accumulated should be carried out.</li> </ul>
<b>Evidence</b>	The surface of the skin, the vernix, and the amniotic fluid protect the newborn against bacterial invasion at birth (663-672).

## 5. Breastfeeding in the first hour

<b>Background</b>	Early skin-to-skin contact means placing the naked healthy neonate on its stomach on the mother's naked breast [immediately after birth, or shortly thereafter (within 24 hours)]. Although, from an evolutionary perspective, skin-to-skin contact is the norm, current practice in many places is to separate the newborn from the mother shortly after birth. Early skin-to-skin contact has been shown to be associated with exclusive and continuing breastfeeding, as well as with positive physiological and behavioral characteristics in the mother and newborn.
<b>Intervention</b>	Place the newborn in skin-to-skin contact with its mother, and initiate breastfeeding within the first hour after delivery, whether the birth be vaginal or caesarean.
<b>Evidence</b>	Initiating breastfeeding for the full-term newborn in the first hour of life reduces neonatal mortality by 22%. Delaying it more than 1 hour after birth increases the probability of neonatal death by as much as a factor of 2.5; delaying it by a day increases neonatal mortality by a factor of 4.5; and a delay of 3 days raises it by a factor of 11.

Early skin-to-skin contact (at any time from 1 to 24 hours after birth) has been shown to have statistically significant positive effects on the continuation of breastfeeding 1 to 4 months after birth, and to be associated with more optimal duration of breastfeeding. Statistically significant benefits of early skin-to-skin contact have also been demonstrated in newborns' maintenance of their temperature in the neutral range, in their crying patterns, in their blood glucose, and in overall scores for love and affectionate contact behavior on the part of the mother during breastfeeding in the first days of life. Premature newborns who are almost full-term have shown improved cardiorespiratory stability when exposed to early skin-to-skin contact (673-685).

## 6. Kangaroo mother program (skin-to-skin contact)

<b>Background</b>	The kangaroo mother program arose in a framework of primary care strategy, as an option to be used in the comprehensive care of premature and low birthweight infants, and is valid for both developed and developing countries. It has evolved into a form of care that is emphatically human-oriented and low-cost in a range of care scenarios – from situations with no resources whatsoever to facilities with funding and advanced technology.
	<p>The components of the program are as follows. Position: skin-to-skin contact 24 hours a day 7 days a week on the chest of the mother (or another family member). Nutrition: exclusive breastfeeding. Follow-up: early discharge with strict ambulatory monitoring up to 40 weeks of corrected age. Intervention Skin-to-skin contact and breastfeeding start as soon as possible in the neonatology unit, and the newborn is to be kept in skin-to-skin contact at home with its mother 24 hours a day.</p>

<b>Evidence</b>	A systematic review began with 6,127 studies, and selected 15, including 9 random controlled trials (RCTs) and 6 observational studies in low- and medium-income countries, all involving children in hospitals < 2000 g (~ 32–34 weeks of GA). The meta-analysis of the RCTs, as well as a separate meta-analysis of the observational studies of the kangaroo mother program starting in the first week of life, compared the results of the program with the results of standard care, and showed (for the first time) a statistically significant reduction of neonatal mortality (51%, CI 29%–82%) and severe morbidity – mostly infection (66%, CI 35%–83%).  The following benefits have been observed. The physical growth of the children in the kangaroo mother program exceeds that of premature or low birthweight infants cared for in incubators. The duration of exclusive breastfeeding is longer for children in the program; more breast milk is produced; and hospital stays are reduced, which reduces costs and diminishes the risk of hospital infections. Children subject to environmental risks are also protected (they have fewer rehospitalizations due to hypothermia, bronchoaspiration, and infection). The program improves mother-newborn and family-newborn bonds, and unites families more effectively (686–698).
-----------------	--

## 7. Stabilizing and transporting the neonate

<b>Background</b>	The purpose of transferring critically sick neonates is to reduce mortality and morbidity, and to minimize sequelae to the extent possible, using technically appropriate, optimal management. The ideal is to anticipate the need and transport the infant “in utero.” When this is not possible, the neonate should be transferred by a specialized team equipped to deal with the infant’s vulnerability, its disease or abnormality, and the physical process involved in the transfer. The recognition of potentially destabilizing factors minimizes these risks.
<b>Criteria</b>	The following criteria should be assessed for any transfer: <ul style="list-style-type: none"> <li>■ The neonate has a condition that cannot be treated in the facility where it was born.</li> <li>■ The referral center provides treatment and stabilization insofar as it is able. The center receiving the referral has the space, capacity, and trained personnel available to treat the patient, and has accepted the referral.</li> <li>■ The referring facility sends the relevant medical records, (ideally along with cord blood and placenta as well).</li> <li>■ The transfer is handled by skilled personnel, and uses appropriate equipment and means of transportation.</li> </ul>
<b>Evidence</b>	High-risk pregnancies and births in health units without adequate neonatal intensive care have a higher incidence of destabilization, perinatal mortality, and neurological sequelae. They should therefore have skilled personnel, and appropriate equipment and means of transportation (699–723).

## 8. Treating neonatal sepsis with a single daily dose of gentamicin

<b>Background</b>	Gentamicin is widely used to treat presumed or confirmed neonatal septicemia. Animal studies and systematic reviews of studies of older children and adults suggest that a daily dose is better than multiple doses, providing adequate levels in the blood without nephrotoxic or auditory effects.
<b>Treatment</b>	Gentamicin, 4 mg/kg, one dose daily, IV or IM, for 7–10 days.
<b>Evidence</b>	Pharmacokinetic evidence indicates that a single daily dose is better than a regimen of multiple doses to treat septicemia in newborns with over 32 weeks of gestation (724–731).

## 9. Treating respiratory distress syndrome (RDS) with natural surfactant

<b>Background</b>	Respiratory distress syndrome (RDS) is caused by a deficiency or dysfunction of the pulmonary tensioactive substance called surfactant. A broad range of products made from tensioactive agents – both
-------------------	--

synthetic and animal-derived preparations – has been formulated and studied in clinical trials. These substitute surfactants have been tested as a way of preventing RDS in premature newborns at high risk, as well as for treating those already experiencing breathing problems.

<b>Diagnosis</b>	Gestational age < 37 weeks. Progressive respiratory insufficiency. Hypoxemia with reduction of SaO <sub>2</sub> and elevated CO <sub>2</sub> . Chest x-ray: reduction of pulmonary volume with diffuse reticulogranular infiltrate described as looking like rough glass, and aerial bronchograms.
<b>Treatment</b>	<i>Artificial surfactant</i> : 4 mL/kg per dose, IT <i>Synthetic surfactant</i> : 5 mL/kg per dose, IT
<b>Evidence</b>	In newborns already experiencing breathing problems, multiple doses of natural surfactant produce better oxygenation, and better results in terms of ventilation parameters, while reducing the risk of pneumothorax, pulmonary interstitial emphysema, bronchopulmonary dysplasia, and necrotizing enterocolitis. In all cases, it is associated with lower mortality (732–741).

## 10. Neonatal screening to detect congenital hypothyroidism, galactosemia, phenylketonuria, and cystic fibrosis

<b>Background</b>	Neonatal screening tests are used to identify newborns who are carriers of an endocrine pathology, infection, or metabolic errors, before the disease manifests itself, and to prevent, if possible, any physical or mental disability or death. The principal screening tests detect an endless number of pathologies, but the most common in our environments are hypothyroidism, galactosemia, phenylketonuria, and cystic fibrosis.  Hypothyroidism is due to a deficiency of circulating thyroid hormones, and if it occurs during embryonic development or in the first months or years of life, it can have serious impact on intellectual, somatic, motor, bone, and functional development.  Galactosemia is a rare recessive autosomal disease characterized by a reduced ability to convert galactose in the diet into glucose, as a result of a deficiency of one of the three enzymes necessary for the conversion. Symptoms usually appear in the first days or weeks of life, and can be fatal since they lead to acute hepatic toxicity, hemorrhagic diathesis, and a predisposition to sepsis by <i>Escherichia coli</i> .  The primary defect in phenylketonuria is absence or deficiency of the liver enzyme phenylalanine hydroxylase. This prevents the conversion of phenylalanine into tyrosine, which in turn precludes the normal subsequent conversions. Children with this type of disorder show limited signs of it until they develop mental retardation – which may not be discernable until the second year of life, by which time it is irreversible.  A diagnosis of cystic fibrosis is usually suggested by phenotypic clinical manifestations (chronic sinopulmonary disease, pancreatic insufficiency, malnutrition, liver disease, or salt-loss syndrome) and confirmed by findings of a high concentration of sodium and chlorine in pharmacologically stimulated perspiration. The analysis of mutations can be useful in: identifying carriers, obtaining diagnostic confirmation in patients with atypical clinical presentations, and making prenatal diagnoses in high-risk pregnancies.
<b>Diagnosis</b>	Diagnosis is made by taking a blood sample from the cord, or drops of fresh capillary blood, usually from the heel, 4–7 days after birth.

**Confirmation**

<b>Pathology</b>	<b>Normal value</b>
Hypothyroidism	12-24 hours TSH < 37 mIU/mL of serum
	24 hours - 14 days TSH < 30 m IU/mL
	>14 days TSH < 20 m IU/mL
Galactosemia	Total galactose: < 10 mg/dL of blood
Phenylketonuria	Phenylalanine 26-91 mM
Cystic fibrosis	Immunoreactive trypsinogen <90 n...g/mL of blood

**Treatment**

*Hypothyroidism:* levothyroxin, 10-15 µg/kg/day until TSH is normalized, then adjust dose.

*Galactosemia:* Strict elimination of galactose and lactose from the diet; breastfeeding is contraindicated, and soy-based formula can be used.

*Phenylketonuria:* Restrict phenylalanine in the diet, as well as proteins in general.

*Cystic fibrosis:* There is no specific treatment. Treatment only relieves symptoms and delays the progress of the disease.

**Evidence**

Very early diagnosis and treatment of confirmed cases diminishes or eliminates the risk of suffering from the disease, and reduces alterations of intellectual, somatic, motor, bone, and functional development (742-760).

**11. Detecting and preventing tuberculosis and hepatitis B****Background**

The hepatitis B virus continues to be a serious public health problem. In pediatric patients, vertical transmission through the placenta is the most frequent route of infection. The incidence of perinatal transmission is 70%-80% when the mother is HBsAg-positive and HBeAg-positive. For tuberculosis, the BCG vaccine has proven to be relatively effective (> 80%) for the serious forms of the disease such as tubercular meningitis and miliary tuberculosis, especially in the pediatric population.

**Treatment**

Neonates exposed to a HBsAg-positive mother should immediately (ideally in the first 12 hours) receive hyperimmune immunoglobulin for hepatitis B (HBIG), and should begin the hepatitis B vaccination series at the latest within the first month of life, with the recombinant vaccine administered at a different site from that used for the immunoglobulin injection.

All unexposed healthy newborns should receive an obligatory first dose of hepatitis B vaccine intramuscularly immediately upon birth.

The BCG (bacillus Calmette-Guérin) vaccine was developed in 1921 by Calmette and Guérin of the Pasteur Institute. It is composed of an attenuated live strain of mycobacterium bovis that over many years in different laboratories has been developed from multiple cultivated substrains. Neonates weighing over 2,000 g at birth should receive an obligatory dose of BCG intradermally before being discharged from the hospital.

**Evidence**

Protection against perinatal transmission of hepatitis B and tuberculosis (761-766).

**12. Detecting and treating retinopathy of prematurity (ROP) in the newborn****Background**

Newborns with very low birthweights (< 1,500 g) or gestational age ≤ 32 weeks, and premature babies with a history of prolonged oxygen supplementation or of mechanical ventilation, are at greater risk of

developing retinopathy of prematurity. ROP is the abnormal development of the blood vessels in the retina. It commonly begins during the first days of life, and can progress rapidly, causing blindness in a matter of weeks.

<b>Diagnosis</b>	Pupillary dilation and indirect ophthalmoscopy begin at week 31, and the neonate's condition should be monitored until the retina matures at 40 weeks.
<b>Treatment</b>	Once the premature baby is diagnosed, photocoagulation should be used. Laser is considered the best way of treating ROP.
<b>Evidence</b>	Preventing exposure to high oxygen levels by controlled use of oxygen or pulse oximetry can reduce the number of newborns with serious and irreversible stages of the disease (767–780).

### 13. Early home visits for newborn care

<b>Background</b>	Three fourths of all neonatal deaths occur in the first week of life, and 25% to 45% in first 24 hours. This is also the period of most maternal deaths. Nevertheless, 47% of mothers and newborns do not receive adequate care during birth, and 72% of births outside institutional settings are not followed up by a postpartum check-up. It is recommended that health care services conduct the first postnatal visit after the first week of life, when the period of highest vulnerability has passed. The importance of providing care in the first hours and days of life has not been sufficiently stressed. This is the period in which the care continuum is most frequently interrupted.
<b>Intervention</b>	WHO and UNICEF recommend at least two home visits to mothers and newborns: the first within 24 hours after birth (day 1) and a second on day 3. If possible, a third visit should be made after the first week of life (day 7).
<b>Evidence</b>	Several studies have demonstrated that early home visits for the care of the newborn effectively reduce neonatal mortality in high-risk populations. They have also been shown to yield improvements in key neonatal care practices, such as commencing breastfeeding, exclusive breastfeeding, skin-to-skin contact, delayed bathing, and better hygiene, as well as hand washing with clean water and soap, and care of the umbilical cord (781–787).

### 14. Supine sleeping position and sudden infant death syndrome (SIDS)

<b>Background</b>	SIDS is the sudden inexplicable death of a child under one year of age.
<b>Risk factors</b>	Male sex, prematurity, low birthweight.
<b>Intervention</b>	Neonates and infants should always sleep on their backs.
<b>Evidence</b>	There is sufficient evidence that putting a baby to rest on its back, avoiding exposure to tobacco smoke, and breastfeeding considerably reduce the incidence of sudden infant death syndrome (788–795).

### 15. Exclusive breastfeeding

<b>Background</b>	Breast milk should be the first thing that the newborn tries, which means that nothing (such as water, other liquids, or ritual foods) should be given before breast milk. Breast milk completely meets an infant's nutritional needs and fluid requirements during the first six months of life. Water or other liquids are not needed to maintain good hydration, even in warm climates.
<b>Intervention</b>	Exclusive breastfeeding, 8 or more times in 24 hours, day and night.
<b>Evaluation</b>	If a baby urinates at least 6 times in 24 hours, its ingestion of breast milk is adequate. Otherwise, the number of feedings should be increased, or the breastfeeding technique being used should be evaluated.
<b>Evidence</b>	<i>For the infant:</i> Exclusive breastfeeding protects against disease and improves the immune system, protecting in the long term against obesity, diabetes, cancer, and high cholesterol.

*For the mother:* Exclusive breastfeeding reduces the risk of postpartum hemorrhage, fosters uterine involution, and delays the return of menstruation, which helps the mother prevent new pregnancies, and protects her against anemia by conserving iron. It also reduces the risk of breast or ovarian cancer before menopause.

*Psychological and developmental benefits:* Exclusive breastfeeding fosters the mother-infant bond, as well as optimal growth and development, including cerebral growth.

*Economic benefits:* Exclusive breastfeeding saves families the cost of buying breast milk substitutes, and reduces health care costs.

*Environmental benefits:* Exclusive breastfeeding conserves natural resources and reduces pollution.

*Impact on mortality:* 16% of neonatal deaths can be prevented if all newborns are exclusively breastfed from their first day of life, and 22% of neonates will be saved if breastfeeding begins in the first hour of life (796-818).

# VI. Summary of interventions and their management

<b>Interventions</b>	<b>Management</b>
<b>I. Prepregnancy interventions</b>	
1. Using body mass index (BMI) to monitor nutritional status prior to pregnancy	<p><b>When:</b> At each check-up.  <b>How:</b> Weight (kg)/height (m)<sup>2</sup>.  <b>Why:</b> To detect deviations in nutrition and correct them.  <b>How long:</b> All the time.</p>
2. Folic acid supplementation	<p><b>When:</b> 2-3 months before pregnancy.  <b>How:</b> 400 µg/day (0.4 mg/day) orally.  <b>Why:</b> To prevent folate deficiency and, during pregnancy, anomalies of the neural tube and other birth defects.  <b>How long:</b> Indefinitely</p>
3. Detecting, preventing and treating. anemia with iron	<p><b>When:</b> Pallor, or Hb &lt; 12 g/dL: 60 mg of elemental iron/day, taken orally. 120 mg of elemental iron/day, taken orally. Severe pallor, or Hb &lt; 7 g/dL: 120 mg elemental iron/day. Administered orally.  <b>How:</b> With food or at bedtime.  <b>Why:</b> To address anemia.  <b>How long to treat:</b> Until anemia improves.</p>
4. Micronutrients and reproductive health in women	<p><b>When:</b> In the course of health surveillance, 3 months before conception and as long during the postpartum period as breastfeeding continues.  <b>How:</b> Multivitamin supplements and balanced diet.  <b>Why:</b> To prevent acute and chronic deficiencies, overeating, or congenital problems upon becoming pregnant.  <b>How long to treat:</b> Indefinitely</p>
5. Detecting and treating sexually transmitted infections (HIV, Herpes, Gonococcus, Papilloma, etc.)	<p><b>When:</b> In the course of health surveillance.  <b>How:</b> Medical evaluation and laboratory work.  <b>Why:</b> To prevent health problems and perinatal complications.  <b>How long to treat:</b> Depends on the disease.</p>

Interventions	Management
<b>I. Prepregnancy interventions (continuation)</b>	
6. Deparasitization in areas of high prevalence	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Albendazole, 400 mg orally, single dose; Mebendazole, 100 mg twice a day orally for 3 days; or Mebendazole 500 mg orally, single dose.</p> <p><b>Why:</b> To prevent malnutrition and anemia due to parasitemia.</p> <p><b>How long to treat:</b> Once a year.</p>
7. Detecting and treating chronic diseases (cardiovascular, nutritional, endocrine)	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Clinical evaluation and laboratory work.</p> <p><b>Why:</b> To prevent malnutrition, obesity, diabetes and future perinatal complications.</p> <p><b>How long to treat:</b> Depends on disease and protocols.</p>
8. Detecting, preventing, and managing domestic violence	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Medical evaluation of warning signs or signs of violence.</p> <p><b>Why:</b> To prevent mortality, depression, physical injury, and low self-esteem.</p> <p><b>How long to treat:</b> Depends on the injuries and on national protocols for violence management.</p>
9. Detecting, preventing, and managing alcohol and tobacco consumption	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Medical evaluation of warning signs in consumption patterns.</p> <p><b>Why:</b> To prevent mortality, depression, antisocial personality, poor school performance, and future reproductive dysfunction.</p> <p><b>How long to treat:</b> On an ongoing basis [TN: via support groups].</p>
10. Detecting, preventing, and managing depression	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Medical evaluation of warning signs.</p> <p><b>Why:</b> To prevent mortality, poor school performance and future postpartum depression.</p> <p><b>How long to treat:</b> On an ongoing basis via support groups.</p>
11. Detecting and preventing cervical cancer	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Cytology and colposcopy.</p> <p><b>Prevention target:</b> Human papillomavirus (HPV) in all adolescents 11-12 years old, and in women 13-26 who have not received the vaccine previously.</p> <p><b>Why:</b> To prevent cancer and mortality.</p> <p><b>How long to treat:</b> First dose when detected, second two months after the first, and a third 6 months after the first.</p>
12. Complete immunization series	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Complete the vaccination series according to the country's Ministry of Health guidelines.</p> <p><b>Why:</b> To prevent morbidity and future congenital infections.</p> <p><b>How long to treat:</b> Follow age guidelines for vaccination series.</p>
13. Detecting and treating periodontal disease	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Dental check-up (semiannual).</p> <p><b>Why:</b> To prevent oral morbidity and future occurrence of low birthweight, prematurity, sepsis, etc.</p> <p><b>How long to treat:</b> Depends on morbidity.</p>
14. Preventing pregnancy in adolescents	<p><b>When:</b> In the course of health surveillance.</p> <p><b>How:</b> Educational interventions targeting service providers and parents, plus programs on the use of contraceptives.</p> <p><b>Why:</b> To prevent perinatal mortality and other adverse situations, including future low birthweight or premature birth.</p> <p><b>How long to treat:</b> On an ongoing basis.</p>

Interventions	Management
<b>II. Interventions during pregnancy</b>	
1. Prenatal check-up	<p><b>When:</b> 5 check-ups in the course of the pregnancy.  <b>How:</b> Follow WHO/PAHO guidelines.  <b>Why:</b> To prevent and detect risk.  <b>How long:</b> During pregnancy.</p>
2. Preventing fetal death	<p><b>When:</b> Throughout pregnancy.  <b>How:</b> Adequate access to obstetric care, particularly during the prenatal period, labor, and delivery.  <b>Why:</b> To prevent and detect risk of fetal death.</p>
3. Preventing preterm birth	<p><b>When:</b> Throughout pregnancy.  <b>How:</b> Adequate access to obstetric care, particularly during the prenatal period, labor, and delivery. Greater use of evidence-based interventions.  <b>Why:</b> To prevent and detect risk of fetal and neonatal death.</p>
4. Managing premature rupture of membranes	<p><b>When:</b> During prenatal check-up and delivery.  <b>How:</b> <i>In the absence of preterm labor</i> where no pathology is present, administer erythromycin, 250 mg orally, every 6 hours for 7 days; if GA &gt; 26 or &lt; 36 weeks, administer prenatal corticoids. <i>In the presence of labor, with GA &lt; 37 weeks</i>, administer an initial dose of penicillin G, 5 million IU, IV, then 2.5 million IU, IV, every 4 hours.  <b>Why:</b> To reduce the risk of early neonatal infection, RDS, and neonatal mortality.</p>
5. Using the CLAP/WR perinatal clinical record (PCR) and perinatal card	<p><b>When:</b> During prenatal check-up, delivery, puerperium, and the neonatal stage.  <b>How:</b> Follow CLAP/WR guidelines.  <b>Why:</b> To ensure proper use of the guidelines.</p>
6. Weight gain during pregnancy	<p><b>When:</b> At each prenatal check-up.  <b>Why:</b> To detect deviations in the weight increase pattern.  <b>How:</b> Use tables from IOM, 2009.  <b>How long to treat:</b> At each check-up.</p>
7. Reducing unnecessary caesarean sections	<p><b>When:</b> During labor and delivery.  <b>How:</b> Avoid vaginal or caesarean births before the full 39 weeks of GA.  <b>Why:</b> To prevent prematurity and its complications</p>
8. Immunization with tetanus toxoid, taking previous vaccination status into account	<p><b>When:</b> At first contact and 4 weeks later.  <b>How:</b> 0.5 mL per dose, IM.  <b>Why:</b> To prevent neonatal tetanus.  <b>How long:</b> 2 doses, or until 5 doses have been received.</p>
9. Preventing influenza with vaccination	<p><b>When:</b> At first prenatal check-up.  <b>How:</b> Single dose of inactivated seasonal or pandemic influenza virus vaccine.  <b>Why:</b> To prevent influenza A (H1N1) 2009 and severe complications during pregnancy.</p>
10. Syphilis screening and case management	<p><b>When:</b> At first prenatal visit and before delivery.  <b>How:</b> Conduct VDRL or RPR tests twice during pregnancy.  <b>Why:</b> To detect syphilis and prevent congenital syphilis.  <b>How long to treat:</b> If test is positive, administer a single dose of benzathine penicillin, 2.4 M units.</p>

Interventions	Management
<b>II. Interventions during pregnancy (continuation)</b>	
11. Detecting and treating asymptomatic bacteriuria	<p><b>When:</b> At first prenatal visit when gestation is &lt; 12 weeks, and at weeks 28 and 32.</p> <p><b>How:</b> <i>Diagnosis:</i> reactive strips or urine culture. <i>Treatment:</i> Cefalexin, 500 mg, orally every 8 hours; amoxicillin, 500 mg, orally every 8 hours; or cefadroxil, nitrofurantoin. 7-10 day course.</p> <p><b>Why:</b> Pyelonephritis prevents preterm delivery and LBW.</p> <p><b>How long to treat:</b> 7-10 days.</p>
12. Treating and preventing anemia with iron	<p><b>When:</b> Throughout pregnancy, postpartum, and breastfeeding period.</p> <p><b>How:</b> Pallor, or Hb &lt; 12 g/dL: 60 mg/day elemental iron. Severe pallor, or Hb &lt; 7 g/dL: 120 mg elemental iron/day. Administered orally.</p> <p><b>Why:</b> To cure anemia.</p> <p><b>How long to treat:</b> Throughout pregnancy, and three months postpartum.</p>
13. Preventing preeclampsia and eclampsia with calcium	<p><b>When:</b> In all pregnant women with risk factors for preeclampsia (&lt; 3 portions of dairy products/day, or BMI &lt; 18.5 or &gt; 25 in the first trimester).</p> <p><b>How:</b> 1-2 g calcium/day orally.</p> <p><b>Why:</b> To prevent hypertension and preeclampsia in women at risk.</p> <p><b>How long to treat:</b> Throughout pregnancy.</p>
14. Preventing preeclampsia and eclampsia with aspirin	<p><b>When:</b> In all pregnant women with risk factors for preeclampsia (BMI &lt; 18.5 or &gt; 25.0 in the first trimester).</p> <p><b>How:</b> 75-100 mg/day orally.</p> <p><b>Why:</b> To prevent preeclampsia, premature delivery, low birthweight, and perinatal mortality.</p> <p><b>How long to treat:</b> Throughout pregnancy.</p>
15. Detecting and treating sexually transmitted infections	<p><b>When:</b> At prenatal check-ups.</p> <p><b>How:</b> Clinical evaluation and laboratory tests.</p> <p><b>Why:</b> To prevent infections, birth defects, and perinatal complications.</p> <p><b>How long to treat:</b> Depends on disease.</p>
16. Antiretrovirals and other measures to reduce the risk of vertical transmission of HIV	<p><b>When:</b> During pregnancy and childbirth.</p> <p><b>How:</b> Zidovudine: <i>During pregnancy:</i> 100 mg orally 5 times/day beginning between weeks 14 and 34 and continuing throughout pregnancy. <i>During elective caesarean section:</i> 2 mg/kg IV followed by continuous infusion of 1 mg/kg/hour up to delivery. Should be accompanied by: elective caesarean section, education for parents, and use of formula for feeding.</p> <p><b>Why:</b> To reduce vertical transmission.</p> <p><b>How long to treat:</b> Until delivery is complete.</p>
17. Detecting and treating diabetes	<p><b>When:</b> At prenatal check-up.</p> <p><b>How:</b> A family history of diabetes, BMI &gt; 25.0, fasting blood glucose &gt; 105 mg/dL.</p> <p><b>Why:</b> To prevent birth defects and perinatal complications.</p> <p><b>How long to treat:</b> Depends on disease and protocols.</p>
18. Deparasitization in areas of high prevalence	<p><b>When:</b> Second and third trimester.</p> <p><b>How:</b> Albendazole, 400 mg orally, single dose.</p> <p><b>Why:</b> To prevent anemia.</p> <p><b>How long to treat:</b> A single dose 2 times.</p>

Interventions	Management
<b>II. Interventions during pregnancy (continuation)</b>	
19. Detecting and treating group B streptococcus	<p><b>When:</b> During pregnancy.</p> <p><b>How:</b> Rectal and vaginal culture between weeks 35 and 37. For intrapartum prophylaxis: Penicillin G: 5 million units IV (initial dose), then 2.5 million units IV every 6 hours for 48 hours, or ampicillin, initial dose 2g IV, then 1g IV every 4 hours up to delivery. If woman is allergic to penicillin, administer cefazolin, initial dose 2g IV, then 1g IV every 8 hours up to delivery.</p> <p><b>Why:</b> Prevents early neonatal infection by group B streptococcus.</p>
20. Nifedipine during preterm labor	<p><b>When:</b> During preterm labor.</p> <p><b>How:</b> 10 mg orally. If uterine activity continues, administer 10 mg every 20 minutes orally, with a total of 3 doses.</p> <p><b>Why:</b> Prevents premature birth, RDS, intraventricular hemorrhage, and jaundice.</p> <p><b>How long to treat:</b> Every 20 minutes, 3 doses in all.</p>
21. Prenatal corticosteroids (betamethasone, dexamethasone) to foster lung maturation	<p><b>When:</b> From 26 weeks to &lt; 35 weeks of gestation.</p> <p><b>How:</b> BMT: 12 mg IM every 24 hours in 2 doses. DMT: 6 mg IM every 12 hours in 4 doses.</p> <p><b>Why:</b> Reduces neonatal mortality, RDS, intraventricular hemorrhage, NNE, sepsis</p> <p><b>How long to treat:</b> Only cycle.</p>
22. Detecting, preventing, and treating Rh isoimmunization	<p><b>When:</b> If there is a history, administer gammaglobulin at weeks 28 and 34 of gestation. In the absence of a history, administer within 72 hours after delivery if baby is Rh positive.</p> <p><b>How:</b> 120-300 milligrams, IM or IV.</p> <p><b>Why:</b> To reduce intrauterine death, fetal hydrops, jaundice, and/or severe anemia.</p>
23. Detecting and treating periodontal disease	<p><b>When:</b> 2 check-ups during pregnancy.</p> <p><b>How:</b> Dental check-up.</p> <p><b>Why:</b> To reduce risk of premature birth.</p> <p><b>How long to treat:</b> Per dentist's recommendation.</p>
24. Detecting, preventing, and managing domestic violence, abuse and depression	<p><b>When:</b> Always.</p> <p><b>How:</b> Seek support from some family member, and psychosocial support.</p> <p><b>Why:</b> To prevent death or disability, and emotional trauma.</p> <p><b>How long to treat:</b> Duration of support group on a case-by-case basis.</p>
25. Detecting and preventing exposure to tobacco	<p><b>When:</b> Throughout pregnancy.</p> <p><b>How:</b> Psychosocial support.</p> <p><b>Why:</b> Prevent IUGR, low birthweight, and sudden infant death syndrome.</p> <p><b>How long to treat:</b> On a case-by-case basis.</p>
26. Detecting and preventing exposure to alcohol	<p><b>When:</b> Throughout pregnancy.</p> <p><b>How:</b> Psychosocial support.</p> <p><b>Why:</b> To prevent fetal alcohol syndrome.</p> <p><b>How long to treat:</b> On a case-by-case basis.</p>
27. Detecting and preventing exposure to drugs	<p><b>When:</b> Throughout pregnancy.</p> <p><b>How:</b> Psychosocial support.</p> <p><b>Why:</b> To prevent low birthweight, withdrawal syndrome, birth defects, and sudden infant death syndrome.</p> <p><b>How long to treat:</b> On a case-by-case basis.</p>

Interventions	Management
<b>III. Interventions during delivery</b>	
1. Clean and safe delivery practices (with skilled personnel)	<p><b>When:</b> During delivery.  <b>How:</b> Hand washing, clean instruments, facilitating environment.  <b>Why:</b> To prevent infections and complications.  <b>How long to treat:</b> During delivery and postpartum.</p>
2. Accompanying the mother during labor and delivery	<p><b>When:</b> During labor and delivery.  <b>How:</b> Permitting and encouraging a family member or other person to be present.  <b>Why:</b> To prevent obstetric complications, minimize the duration of labor, and reduce the need for caesarean section or pain medication.</p>
3. Erythromycin in PROM before delivery in first 37 weeks of gestation	<p><b>When:</b> In the absence of labor.  <b>How:</b> Erythromycin, 500 mg orally every 8 hours.  <b>Why:</b> Prevents preterm birth, neonatal sepsis, and chorioamnionitis.  <b>How long to treat:</b> 7 days.</p>
4. Monitoring labor with partograms	<p><b>When:</b> During labor.  <b>How:</b> Partogram.  <b>Why:</b> To reduce unnecessary interventions and perinatal complications.  <b>How long to treat:</b> Throughout labor.</p>
5. Caesarean section in breech presentation with LBW infants	<p><b>When:</b> During labor.  <b>How:</b> Elective caesarean section.  <b>Why:</b> To reduce perinatal complications, asphyxiation, and birth trauma.</p>
6. Antibiotic prophylaxis in caesarean section	<p><b>When:</b> Elective or non-elective caesarean section.  <b>How:</b> Cefalotin, 2g IV, single dose, during caesarean section.  <b>Why:</b> To prevent endometritis and its severe sequelae.  <b>How long to treat:</b> A single dose.</p>
7. Modified active management of the third stage of labor	<p><b>When:</b> During the third stage of labor (birth).  <b>How:</b> Oxytocine, 5-10 IU, IM; methylergonovine 0.5 mg, IM; or misoprostol, 0.4-0.8 mg orally; + traction/contraction + abdominal uterine massage.  <b>Why:</b> To reduce postpartum bleeding.  <b>How long to treat:</b> One dose.</p>
8. Late clamping of the umbilical cord	<p><b>When:</b> During delivery.  <b>How:</b> Clamp at 2-3 minutes.  <b>Why:</b> To permit more deposit of iron in the newborn, preventing anemia during the first 4-6 months of life, and improving the newborn's cerebral oxygenation.</p>

<b>Interventions</b>	<b>Management</b>
<b>IV. Maternal interventions after delivery</b>	
1. Early skin-to-skin contact	<p><b>When:</b> Immediately after birth.</p> <p><b>How:</b> Place baby in skin-to-skin contact with its mother.</p> <p><b>Why:</b> To promote early bonding and the beginning of breastfeeding.</p> <p><b>How long to treat:</b> As long as possible.</p>
2. Mother-newborn rooming-in	<p><b>When:</b> 24 hours a day.</p> <p><b>How:</b> Place the baby in contact with its mother.</p> <p><b>Why:</b> To promote early bonding and the beginning of breastfeeding, as well as parents' confidence in caring for the baby.</p> <p><b>How long to treat:</b> As long as possible.</p>
3. Detecting and managing puerperal endometritis	<p><b>When:</b> Fever in the mother third or fourth day after birth.</p> <p><b>How:</b> Gentamicin + clindamycin.</p> <p><b>Why:</b> To reduce maternal mortality from sepsis.</p> <p><b>How long to treat:</b> 10 days.</p>
4. Detecting, preventing, and managing postpartum depression	<p><b>When:</b> At any time within 3 months after delivery.</p> <p><b>How:</b> Detect through warning signs; treat with support groups or, in severe cases, antidepressants.</p> <p><b>Why:</b> To reduce maternal and neonatal morbidity and mortality.</p> <p><b>How long to treat:</b> Until symptoms recede.</p>
<b>V. Neonatal interventions after delivery</b>	
1. Neonatal resuscitation with ambient air	<p><b>When:</b> Immediately after birth when required.</p> <p><b>How:</b> Follow neonatal resuscitation protocol.</p> <p><b>Why:</b> To avoid the toxicity of oxygen and ROP sequelae as well as bronchopulmonary dysplasia.</p> <p><b>How long to treat:</b> Up to 10 minutes.</p>
2. Immediate care for the healthy newborn	<p><b>When:</b> Upon birth.</p> <p><b>How:</b> Following immediate care protocol.</p> <p><b>Why:</b> To prevent hypothermia, hypoglycemia, sepsis, and other complications at birth.</p>
3. Care of the umbilical cord	<p><b>When:</b> Upon birth, and until it falls off.</p> <p><b>How:</b> Initially apply antiseptic solution, then keep clean and dry.</p> <p><b>Why:</b> To prevent omphalitis and sepsis.</p> <p><b>How long to treat:</b> Until it falls off.</p>
4. Skin care at birth	<p><b>When:</b> Avoid immediate bathing of the preterm or low birthweight newborn.</p> <p><b>How:</b> Keep clean by sponge bath; do not immerse.</p> <p><b>Why:</b> To prevent hypothermia and infection.</p> <p><b>How long to treat:</b> Always.</p>
5. Breastfeeding in the first hour	<p><b>When:</b> From the first hour after birth.</p> <p><b>How:</b> Place the newborn in skin-to-skin contact with its mother, and initiate breastfeeding.</p> <p><b>Why:</b> To foster exclusive breastfeeding and a longer period of breastfeeding.</p> <p><b>How long to treat:</b> Continue exclusive breastfeeding until 6 months old.</p>
6. Kangaroo mother program (skin-to-skin contact)	<p><b>When:</b> When the newborn is stable and can suck.</p> <p><b>How:</b> Keep the newborn in skin-to-skin contact with its mother at home 24 hours a day.</p> <p><b>Why:</b> To foster bonding, provide psychological benefits, extend breastfeeding, and reduce neonatal mortality.</p> <p><b>How long to treat:</b> Generally a month.</p>

Interventions	Management
<b>V. Neonatal interventions after delivery (continuation)</b>	
7. Stabilizing and transporting the neonate	<p><b>When:</b> When it is necessary to transfer a newborn from the health care facility to a facility of greater complexity.</p> <p><b>How:</b> By any means of transportation adequately equipped for transporting newborns, and with skilled personnel, following established criteria for transfer.</p> <p><b>Why:</b> To reduce complications and neonatal mortality.</p> <p><b>How long to treat:</b> Throughout the transport process.</p>
8. Treating neonatal sepsis with a single daily dose of gentamicin	<p><b>When:</b> When neonatal sepsis is suspected.</p> <p><b>How:</b> Gentamicin, 4 mg/kg, one dose daily, IV or IM, for 7-10 days.</p> <p><b>Why:</b> To reduce complications and neonatal mortality.</p> <p><b>How long to treat:</b> 7-10 days.</p>
9. Treating respiratory distress syndrome (RDS) with natural surfactant	<p><b>When:</b> When hyaline membrane disease is suspected in a preterm newborn.</p> <p><b>How:</b> <i>Artificial surfactant:</i> 4 mL/kg per dose, IT.      <i>Synthetic surfactant:</i> 5 mL/kg per dose, IT.</p> <p><b>Why:</b> To reduce complications and neonatal mortality.</p> <p><b>How long to treat:</b> Repeat the dose according to how the condition evolves.</p>
10. Neonatal screening to detect congenital hypothyroidism, galactosemia, phenylketonuria, and cystic fibrosis.	<p><b>When:</b> Between days 4 and 7 of extrauterine life.</p> <p><b>How:</b> Blood drawn from the newborn (heel).</p> <p><b>Why:</b> Early diagnosis to prevent sequelae in intellectual and somatic development.</p> <p><b>How long to treat:</b> Immediately upon diagnosis.</p>
11. Preventing tuberculosis and hepatitis B	<p><b>When:</b> Before discharge.</p> <p><b>How:</b> BCG and hepatitis B vaccine. [TN: that doesn't answer the question of "detecting," which the running text does touch on]</p> <p><b>Why:</b> To prevent sequelae and neonatal mortality.</p> <p><b>How long to treat:</b> BCG: once only; hepatitis B: first dose.</p>
12. Detecting and treating retinopathy of prematurity (ROP) in the newborn	<p><b>When:</b> At 4-6 weeks of age.</p> <p><b>How:</b> Ophthalmological evaluation and laser photocoagulation treatment.</p> <p><b>Why:</b> To prevent blindness.</p> <p><b>How long to treat:</b> Depends on degree of ROP.</p>
13. Early home visits for newborn care	<p><b>When:</b> On days 1, 3, and 7 after birth.</p> <p><b>How:</b> Evaluate basic neonatal care and check key practices.</p> <p><b>Why:</b> To detect neonatal morbidity and support exclusive breastfeeding.</p>
14. Supine sleeping position sudden infant death syndrome (SIDS)	<p><b>When:</b> Always up to 1 year of age.</p> <p><b>How:</b> By ensuring that the baby sleeps on its back.</p> <p><b>Why:</b> To reduce the possibility of sudden death.</p>
15. Exclusive breastfeeding	<p><b>When:</b> From first hour after birth.</p> <p><b>How:</b> Eight times or more per 24 hours, day and night.</p> <p><b>Why:</b> To foster bonding, reduce neonatal infection, and provide psychological, economic, and environmental benefits.</p> <p><b>How long to treat:</b> Continue exclusive breastfeeding until 6 months old.</p>
15. Lactancia materna exclusiva	<p><b>Cuándo:</b> Desde la primera hora después de nacer.</p> <p><b>Cómo:</b> Ocho veces o más en 24 horas, de día y de noche.</p> <p><b>Por qué:</b> Favorece apego, disminuye infecciones neonatales, así como beneficios psicológicos, económicos y al medio ambiente.</p> <p><b>Cuánto tiempo tratar:</b> En forma exclusiva hasta los 6 meses de edad.</p>

Early skin-to-skin contact (at any time from 1 to 24 hours after birth) has been shown to have statistically significant positive effects on the continuation of breastfeeding 1 to 4 months after birth, and to be associated with more optimal duration of breastfeeding.



# VII. Bibliographic references

## Introduction

- Dramstadt GL, Bhutta ZA, Cousens S, et al. Evidence-based, cost-effective interventions: how many newborn babies can we save? *The Lancet* 2005;March:19-30.
- Acolet D. Quality of neonatal care and outcome. *Arch Dis Fetal Neonatal Ed* 2008;93:F69-F73.
- Burgers JS, Grol RP, Zaat JO, et al. Characteristics of effective clinical guidelines for general practice. *Br J Gen Pract* 2003;53:15-19.
- Grol R, Dalhuijsen J, Thomas S, et al. Attributes of clinical guidelines that influence use of guidelines in general practice: observational study. *BMJ* 1998;317:202-13.
- Rogowski JA, Horbar JD, Staiger DO, et al. Indirect vs direct hospital quality indicators for very low-birth-weight infants. *JAMA*. 2004 Jan 14;291(2):202-9.
- Mercier CE, Barry SE, Paul K, et al. Improving newborn preventive services at the birth hospitalization: a collaborative, hospital-based quality-improvement project. *Pediatrics*. 2007 Sep;120(3):481-8.
- Horbard JD. The Vermont Oxford Network: evidence-based quality improvement for neonatology. *Pediatrics* 1999;103(1 Suppl):350-9.
- Sackett DL, Rosenberg WM, Gray JA, et al. Evidence-based medicine: what it is and what it isn't? *Br Med J*. 1996;312:71-72.

## I. Prepregnancy interventions

- Using body mass index (BMI) to monitor nutritional status prior to pregnancy**
- Fescina RH, De Mucio B, Diaz Rossello JL, et al. Guías para el continuo de atención de la mujer y el recién nacido focalizadas en APS. CLAP/SMR, Pub. Cient. No. 1562, 2008.
- Summers L, Price RA. Preconception care. An opportunity to maximize health in pregnancy. *J Nurse Midwifery*. 1993 Jul-Aug;38(4):188-98.
- Perry LE. Preconception care: a health promotion opportunity. *Nurse Pract*. 1996 Nov;21(11):24-6, 32, 34 passim.
- Morrison EH. Periconception care. *Prim Care*. 2000 Mar;27(1):1-12.

- Riskin-Mashiah S. Preconception counseling--for all. *Harefuah*. 2004 Jul;143(7):530-6, 547.
- Heyes T, Long S, Mathers N. Preconception care: practice and beliefs of primary care workers. *Fam Pract*. 2004 Feb;21(1):22-7.
- IOM (Institute of Medicine), 2009. Weight Gain During Pregnancy: Reexamining the Guidelines. Washington, DC: The National Academies Press. IOM, [www.iom.edu](http://www.iom.edu).
- Canaval HO, Vargas J. Compendio de guías latinoamericanas para el manejo de la anemia ferropénica. AWLA/ALF, 2006.
- Stoltzfus RJ, Dreyfuss ML. Guidelines for the use of iron supplements to prevent and treat iron deficiency anemia. INACG/WHO/UNICEF, 1998.
- Chaparro CM, Lutter CK. Anemia among adolescent and young adult women in Latin America and the Caribbean: A cause for concern. Washington DC, PAHO, 2008.
- WHO, Recommendations on prevention of postpartum hemorrhage. Geneva, WHO, 2007.
- Siega-Riz AM, Hobel CJ. Predictors of poor maternal weight gain from baseline anthropometric, psychosocial, and demographic information in a Hispanic population. *J Am Diet Assoc*. 1997 Nov;97(11):1264-8.
- Schieve LA, Cogswell ME, Scanlon KS, et al. Prepregnancy body mass index and pregnancy weight gain: associations with preterm delivery. The NMIHS Collaborative Study Group. *Obstet Gynecol*. 2000 Aug;96(2):194-200.
- Borkowski W, Mieleniczuk H. Preterm delivery in relation to combined pregnancy weight gain and prepregnancy body mass. *Przegl Epidemiol*. 2007;61(3):577-8.
- Biggio JR, Chapman V, Neely Ch et al. Fetal Anomalies in Obese women. The contribution of Diabetes. *Obstet Gynecol* Feb 2010;115(2):290-6.
- Folic acid (FA) and multivitamin (MV) supplementation**
- Van Allen MI, Fraser FC, Dallaire L, et al. Recommendations on the use of folic acid supplementation to prevent the recurrence of neural tube defects. Clinical Teratology Committee, Canadian College of Medical Geneticists. *CMAJ*. 1993 Nov 1;149(9):1239-43.

25. Czeizel AE. Reduction of urinary tract and cardiovascular defects by periconceptional multivitamin supplementation. *Am J Med Genetics* 1996; 62:179-183.
26. American Academy of Pediatrics. Committee on Genetics. Folic acid for the prevention of neural tube defects. *Pediatrics*. 1999 Aug;104(2 Pt 1):325-7.
27. Czeizel AE. Primary prevention of neural-tube defects and some other major congenital abnormalities: recommendations for the appropriate use of folic acid during pregnancy. *Paediatr Drugs*. 2000 Nov-Dec;2(6):437-49.
28. Wilson RD, Davies G, Désilets V, et al. The use of folic acid for the prevention of neural tube defects and other congenital anomalies. *J Obstet Gynaecol Can*. 2003 Nov;25(11):959-73.
29. Lumley J, Watson L, Watson M, Bower C. Periconceptional supplementation with folate and/or multivitamins for preventing neural tube defects. In: *The Cochrane Library*, Issue 1, 2004.
30. Lumley J, Watson L, Watson M, Bower C.. Periconceptional supplementation with folate and/or multivitamins for preventing neural tube defects. *Cochrane Database Syst Rev*. 2011 Apr 13;(4):CD001056. Review.
31. Martínez de Villarreal LE, Arredondo P, et al. Weekly administration of folic acid and epidemiology of neural tube defects. *Matern Child Health J*. 2006 Sep;10(5):397-401.
32. Wilson RD, Johnson JA, Wyatt P, et al. Pre-conceptional vitamin/folic acid supplementation 2007: the use of folic acid in combination with a multivitamin supplement for the prevention of neural tube defects and other congenital anomalies. *J Obstet Gynaecol Can*. 2007 Dec;29(12):1003-26.
33. Goh YI, Bnollano E, Einarsen TR, Koren G. Motherisk Update 2007. Prenatal multivitamin supplementation and rates of pediatric cancers: a meta-analysis. *Clin Pharm Ther* 2007;81:685-9.
34. Catov JM, Bodnar LM, Ness RB, et al Association of Periconceptional Multivitamin Use and Risk of Preterm or small-for-Gestational-Age Births. *Am J Epidemiol* Vol.166, No. 3, 2007.
35. Wilson RD, Johnson JA, Wyatt P et al. Pre-conceptional vitamin/folic acid supplementation 2007: the use of folic acid in combination with a multivitamin supplement for the prevention of neural tube defects and other congenital anomalies. *J Obstet Gynaecol Can* 2007 Dec; 29 (12):1003-26.
36. Ryan-Harshman M, Aldoori W. Folic acid and prevention of neural tube defects. *Can Fam Physician*. 2008 Jan;54(1):36-8.
37. The Supplementation with Multiple Micronutrients Intervention Trial (SUMMIT) Study Group. *Lancet* 19- 25 Jan 2008: vol 371:(9608):215-227.
38. Czeizel AE. Periconceptional folic acid and multivitamin supplementation for the prevention of neural tube defects and other congenital abnormalities. *Birth Defects Res A Clin Mol Teratol*. 2009 Apr;85(4):260-8.
39. Czeizel AE. Periconceptional Folic Acid and Multivitamin Supplementation for the Prevention of Neural Tube Defects and Other Congenital Abnormalities. *Birth Defects Research (Part A)* 2009;85:260-268.
40. Matok I, Gorodischer R, Koren G, et al "Exposure To Folic Acid Antagonists During The First Trimester of Pregnancy and the Risk of Major Malformations" *Br J Clin Pharmacol* 2009; 68:(6), 956-62.
41. Bukowski R, Malonez FD, Porters FT, Nyberg DA et al. Preconceptional Folate Supplementation and the Risk of Spontaneous Preterm Birth: A Cohort Study. *PLoS Medicine* May 2009/ Vol 6 | Issue 5. www.plosmedicine.org.
- 3. Preventing and treating anemia with iron**
42. Looker AC, Dallman PR, Carroll MD, et al. Prevalence of iron deficiency in the United States. *JAMA*. 1997 Mar 26;277(12):973-6.
43. Beard JL. Iron requirements in adolescent females. *J Nutr*. 2000 Feb;130(2S Suppl):440S-442S.
44. Lynch SR. The potential impact of iron supplementation during adolescence on iron status in pregnancy. *J Nutr*. 2000 Feb;130(2S Suppl):448S-451S.
45. Murray-Kolb LE, Beard JL. Iron treatment normalizes cognitive functioning in young women. *Am J Clin Nutr*. 2007 Mar;85(3):778-87.
46. Ferreira MU, da Silva-Nunes M, Bertolino CN, Malafronte RS, Muniz PT, Cardoso MA. Anemia and iron deficiency in school children, adolescents, and adults: a community-based study in rural Amazonia. *Am J Public Health*. 2007 Feb;97(2):237-9.
47. Zimmermann MB, Hurrell RF. Nutritional iron deficiency. *Lancet*. 2007 Aug 11;370(9586):511-20.
48. Brotanek JM, Gosz J, Weitzman M, Flores G. Iron deficiency in early childhood in the United States: risk factors and racial/ ethnic disparities. *Pediatrics*. 2007;120 (3):568-575.
49. Ginder GD. Microcytic and hypochromic anemias. In: Goldman L, Ausiello D, eds. *Cecil Medicine*. 23rd ed. Philadelphia, Pa: Saunders Elsevier; 2007: chap 163.
50. Killip S, Bennett JM, Chambers MD. Iron deficiency anemia. *Am Fam Physician*. 2007;75(5):671-678.
51. Alleyne M, Horne MK, Miller JL. Individualized treatment for iron-deficiency anemia in adults. *Am J Med*. 2008;(121):943-948.
- 4. Micronutrients and reproductive health in women**
52. Kontic-Vucinic O, Sulovic N, Radunovic N. Micronutrients in women's reproductive health: I. Vitamins. *Int J Fertil Womens Med*. 2006 May-Jun;51(3):106-15.
53. Kontic-Vucinic O, Sulovic N, Radunovic N. Micronutrients in women's reproductive health: II. Minerals and trace elements. *Int J Fertil Womens Med*. 2006 May-Jun;51(3):116-24.
54. Wilson RD, Johnson JA, Wyatt P et al. Pre-conceptional vitamin/folic acid supplementation 2007: the use of folic acid in combination with a multivitamin supplement for the prevention of neural tube defects and other congenital anomalies. *J Obstet Gynaecol Can*. 2007 Dec;29(12):1003-26.

55. Evans L, Weisman CS. Folic acid supplementation in younger and older nonpregnant women of reproductive age: findings from the Central Pennsylvania Women's Health Study (CePAWHS). *Womens Health Issues*. 2010 Jan-Feb;20(1):50-7.
- 5. Detecting and treating sexually transmitted infections (STIs) before pregnancy**
56. Paavonen J, Eggert-Kruse W. Chlamydia trachomatis: impact on human reproduction. *Hum Reprod Update*. 1999 Sep-Oct;5(5):433-47.
57. Casalini C, Signorini L, Beltrame A, et al. Vertical transmission of human immunodeficiency virus (HIV) and other sexually transmitted infections (STI). *Minerva Ginecol*. 2001 Jun;53(3):177-92.
58. Wilkinson D, Rutherford G. Population-based interventions for reducing sexually transmitted infections, including HIV infection. *Cochrane Database Syst Rev*. 2001;(2):CD001220.
59. Gutierrez JP, Conde-González CJ, Walker DM, Bertozzi SM. Herpes simplex virus type 2 among Mexican high school adolescents: prevalence and association with community characteristics. *Arch Med Res*. 2007 Oct;38(7):774-82.
60. Nagot N, Ouedraogo A, Defer MC, et al. Association between bacterial vaginosis and Herpes simplex virus type-2 infection: implications for HIV acquisition studies. *Sex Transm Infect*. 2007 Aug;83(5):365-8.
61. Kaul R, Nagelkerke NJ, Kimani J, et al. Prevalent herpes simplex virus type 2 infection is associated with altered vaginal flora and an increased susceptibility to multiple sexually transmitted infections. *J Infect Dis*. 2007 Dec 1;196(11):1692-7.
62. Sanchez J, Lama JR, Kusunoki L, et al. HIV-1, sexually transmitted infections, and sexual behavior trends among men who have sex with men in Lima, Peru. *J Acquir Immune Defic Syndr*. 2007 Apr 15;44(5):578-85.
63. Ochsendorf FR. Sexually transmitted infections: impact on male fertility. *Andrologia*. 2008 Apr;40(2):72-5.
64. Cunningham KA, Beagley KW. Male Genital Tract Chlamydial Infection: Implications for Pathology and Infertility. *Biol Reprod*. 2008 May 14.
65. Porras C, Safaeian M, González P, et al. Epidemiology of genital chlamydia trachomatis infection among young women in Costa Rica. *Sex Transm Dis*. 2008 May;35(5):461-8.
66. Campero L, Walker D, Rouvier M, Atienzo E. First Steps Toward Successful Communication About Sexual Health Between Adolescents and Parents in Mexico. *Qual Health Res*. 2010 May 6.
- 6. Deparasitization in areas of high prevalence**
67. Gyorkos TW, Larocque R, Casapia M, Gotuzzo E. Lack of risk of adverse birth outcomes after deworming in pregnant women. *Pediatr Infect Dis J*. 2006 Sep;25(9):791-4.
68. Brooker S, Hotez PJ, Bundy DA. Hookworm-related anaemia among pregnant women: a systematic review. *PLoS Negl Trop Dis*. 2008 Sep 17;2(9):e291.
- 7. Detecting and treating chronic diseases**
69. de Onis M, Blossner M. Prevalence and trends of overweight among preschool children in developing countries. *Am J Clin Nutr*. 2000;72:1032-9.
70. Hoffman RP. Practical management of type 1 diabetes mellitus in adolescent patients: challenges and goals. *Treat Endocrinol*. 2004;3(1):27-39.
71. Peña M, Bacallao J. La obesidad en la pobreza: un problema emergente en las Américas. *Revista Futuros* 2005;10, Vol III.
72. Boney CM, Verma A, Tucker R, Vohr BR. Metabolic syndrome in childhood: association with birth weight, maternal obesity, and gestational diabetes mellitus. *Pediatrics*. 2005 Mar;115(3):e290-6.
73. Flynn MA, McNeil DA, Maloff B, et al. Reducing obesity and related chronic disease risk in children and youth: a synthesis of evidence with 'best practice' recommendations. *Obes Rev*. 2006 Feb;7 Suppl 1:7-66.
74. Roman H, Robillard PY, Hulsey TC, et al. Obstetrical and neonatal outcomes in obese women. *West Indian Med J*. 2007 Oct;56(5):421-6.
75. Buzinaro EF, Berchieri CB, Haddad AL, et al. Overweight in adolescent offspring of women with hyperglycemia during pregnancy. *Arg Bras Endocrinol Metabol*. 2008 Feb;52(1):85-92.
76. Cali AM, Caprio S. Prediabetes and type 2 diabetes in youth: an emerging epidemic disease? *Curr Opin Endocrinol Diabetes Obes*. 2008 Apr;15(2):123-7.
77. Weiss R, Gillis D. Patho-physiology and dynamics of altered glucose metabolism in obese children and adolescents. *Int J Pediatr Obes*. 2008;3 Suppl 1:15-20.
78. Lawrence S, Hazlett R, Hightower P. Understanding and acting on the growing childhood and adolescent weight crisis: a role for social work. *Health Soc Work*. 2010 May;35(2):147-53.
79. Baker JL, Farpour-Lambert NJ, et al. Evaluation of the overweight/obese child - practical tips for the primary health care provider: recommendations from the childhood obesity task force of the European association for the study of obesity. *Obes Facts*. 2010;3(2):131-7.
- 8. Detecting, preventing, and managing domestic violence**
80. WHO Global Consultation on Violence and Health. Violence: a public health priority. Geneva, World Health Organization, 1996 (Document WHO/EHA/SPI.POA.2).
81. WHO multi-country study on women's health and domestic violence. Geneva, World Health Organization, 1999 (Document WHO/FCH/GWH/02.01).
82. WHO. Putting women first: ethical and safety recommendations for research on domestic violence against women. Geneva, Worl Health Organization, 2001 (Document WHO/FCH/GWH/01.01).
83. Krug EG, Dahlberg LL, Mercy JA, et al. World report on violence and health. Geneva, World Health Organization, 2002.
84. Plichta SB. Intimate partner violence and physical health consequences: policy and practice implications. *J Interpers Violence*. 2004 Nov;19(11):1296-323.

85. Hussain R, Khan A. Women's perceptions and experiences of sexual violence in marital relationships and its effect on reproductive health. *Health Care Women Int.* 2008 May;29(5):468-83.
86. Sarkar NN. The impact of intimate partner violence on women's reproductive health and pregnancy outcome. *J Obstet Gynaecol.* 2008 Apr;28(3):266-71.
87. Robert J. Jagers; Antonio A. Morgan-Lopez; Brian R. Flay. The Impact of Age and Type of Intervention on Youth Violent Behaviors. *J Prim Prev.* 2009;30(6):642-658.
88. da Fonseca Phd RM, Egry Phd EY, Guedes RN, et al. Violence against women: a study of the reports to police in the city of Itapevi, São Paulo, Brazil. *Midwifery.* 2010 May 20.
89. Miranda MP, de Paula CS, Bordin IA. Life-long domestic violence against women: prevalence and immediate impact on health, work, and family. *Rev Panam Salud Publica.* 2010 Apr;27(4):300-8.
- 9. Detecting, preventing and managing alcohol and tobacco consumption**
90. Rice JP, Neuman RJ, Saccone NL, et al. Age and birth cohort effects on rates of alcohol dependence. *Alcohol Clin Exp Res.* 2003 Jan;27(1):93-9.
91. Grucza RA, Bierut LJ. Co-occurring risk factors for alcohol dependence and habitual smoking: update on findings from the Collaborative Study on the Genetics of Alcoholism. *Alcohol Res Health.* 2006;29(3):172-8.
92. Edenberg HJ, Foroud T. The genetics of alcoholism: identifying specific genes through family studies. *Addict Biol.* 2006 Sep;11(3-4):386-96.
93. Waldron M, Heath AC, Bucholz KK, Madden PA, Martin NG. Alcohol dependence and reproductive onset: findings in two Australian twin cohorts. *Alcohol Clin Exp Res.* 2008 Nov;32(11):1865-74.
94. Clinical Practice Guideline Treating Tobacco Use and Dependence 2008 Update Panel, Liaisons, and Staff. A clinical practice guideline for treating tobacco use and dependence: 2008 update. A U.S. Public Health Service report. *Am J Prev Med.* 2008 Aug;35(2):158-76.
95. Tripodi SJ, Bender K, Litschge C, Vaughn MG. Interventions for reducing adolescent alcohol abuse: a meta-analytic review. *Arch Pediatr Adolesc Med.* 2010 Jan;164(1):85-91.
- 10. Detecting, preventing, and managing depression**
96. Rushton JL, Forcier M, Schectman RM. Epidemiology of depressive symptoms in the National Longitudinal Study of Adolescent Health. *J Am Acad Child Adolesc Psychiatry.* 2002 Feb;41(2):199-205.
97. Merry S, McDowell H, Hetrick S, Bir J, Muller N. Psychological and/or educational interventions for the prevention of depression in children and adolescents. *Cochrane Database Syst Rev.* 2004;(1):CD003380.
98. Lillebeth Larun, Lena V. Nordheim, Eilin Ekeland, Kåre Birger Hagen, Frode Heian. Exercise in prevention and treatment of anxiety and depression among children and young people. *Cochrane Database of Systematic Reviews* 2006, Issue 3. Art. No.: CD004691.
99. Park S, Romer D. Associations between smoking and depression in adolescence: an integrative review. *Taehan Kanho Hakhoe Chi.* 2007 Mar;37(2):227-41.
100. Anthony FJorm, Amy J Morgan, Sarah E Hetrick. Relaxation for depression. *Cochrane Database of Systematic Reviews* 2008, Issue 4. Art. No.: CD007142.
101. Munafò MR, Hitsman B, Rende R, Metcalfe C, Niaura R. Effects of progression to cigarette smoking on depressed mood in adolescents: evidence from the National Longitudinal Study of Adolescent Health. *Addiction.* 2008 Jan;103(1):162-71.
102. Chaiton MO, Cohen JE, O'Loughlin J, Rehm J. A systematic review of longitudinal studies on the association between depression and smoking in adolescents. *BMC Public Health.* 2009 Sep 22;9:356.
103. Lancaster CA, Gold KJ, Flynn HA et al. Risk factors for depressive symptoms during pregnancy: a systematic review. *AJOG Vol 202(1), Jan 2010;* 5-14.
- 11. Detecting and treating cervical cancer**
104. Shepherd JJ, Greet Peersman, Ibrahim Napuli. Interventions for encouraging sexual lifestyles and behaviours intended to prevent cervical cancer. *Cochrane Database of Systematic Reviews* 1999, Issue 4. Art. No.: CD001035.
105. Monsonégo J. Cervical cancer prevention: the impact of HPV vaccination. *Cynecol Obstet Fertil.* 2006 Mar;34(3):189-201.
106. Monsonego J. Prevention of cervical cancer (II): prophylactic HPV vaccination, current knowledge, practical procedures and new issues. *Presse Med.* 2007 Apr;36(4 Pt 2):640-66.
107. Adams M, Jasani B, Fiander A. Human papilloma virus (HPV) prophylactic vaccination: challenges for public health and implications for screening. *Vaccine.* 2007 Apr 20;25(16):3007-13.
108. Miller MM, Wilson JM, Waldrop J. Current acceptance of the HPV vaccine. *Nurse Pract.* 2008 Apr;33(4):18-22; quiz 22-3.
109. Farrell RM, Rome ES. Adolescents' access and consent to the human papillomavirus vaccine: a critical aspect for immunization success. *Pediatrics.* 2007 Aug;120(2):434-7.
110. Luciani S, Vardy L, Paci E, et al. Cancer prevention and population-based screening. *Tumori.* 2009 Sep-Oct;95(5):597-609.
111. Franco EL, Coutlée F, Ferenczy A. Integrating human papillomavirus vaccination in cervical cancer control programmes. *Public Health Genomics.* 2009;12(5-6):352-61.
112. Adamopoulou M, Kalkani E, Charvalos E, et al. Comparison of cytology, colposcopy, HPV typing and biomarker analysis in cervical neoplasia. *Anticancer Res.* 2009 Aug;29(8):3401-9.

113. Ronco G, Giorgi-Rossi P, Carozzi F, et al. Efficacy of human papillomavirus testing for the detection of invasive cervical cancers and cervical intraepithelial neoplasia: a randomised controlled trial. *Lancet Oncol.* 2010 Jan 18.
114. Warman J. Cervical cancer screening in young women: saving lives with prevention and detection. *Oncol Nurs Forum.* 2010 Jan;37(1):33-8.
- 12. Complete immunization series**
115. Mast EE, Margolis HS, Fiore AE, et al. A comprehensive immunization strategy to eliminate transmission of hepatitis B virus infection in the United States: recommendations of the Advisory Committee on Immunization Practices (ACIP) part 1: immunization of infants, children, and adolescents. *MMWR Recomm Rep.* 2005 Dec 23;54(RR-16):1-31.
116. Pichichero ME, Rennels MB, Edwards KM, et al. Combined tetanus, diphtheria, and 5-component pertussis vaccine for use in adolescents and adults. *JAMA.* 2005 Jun 22;293(24):3003-11.
117. Broder KR, Cortese MM, Iskander JK, et al. Preventing tetanus, diphtheria, and pertussis among adolescents: use of tetanus toxoid, reduced diphtheria toxoid and acellular pertussis vaccines recommendations of the Advisory Committee on Immunization Practices (ACIP). *MMWR Recomm Rep.* 2006 Mar 24;55(RR-3):1-34.
118. America Academy of Pediatrics Committee on Infectious Diseases. Prevention of pertussis among adolescents: recommendations for use of tetanus toxoid, reduced diphtheria toxoid, and acellular pertussis (Tdap) vaccine. *Pediatrics.* 2006 Mar;117(3):965-78. Epub 2005 Dec 28.
119. Murphy TV, Slade BA, Broder KR, et al. Prevention of pertussis, tetanus, and diphtheria among pregnant and postpartum women and their infants recommendations of the Advisory Committee on Immunization Practices (ACIP). *MMWR Recomm Rep.* 2008 May 30;57(RR-4):1-51.
120. Centers for Disease Control and Prevention (CDC). Update on influenza A (H1N1) 2009 monovalent vaccines. *MMWR Morb Mortal Wkly Rep.* 2009 Oct 9;58(39):1100-1.
121. Keeton VF, Chen AK. Immunization updates and challenges. *Curr Opin Pediatr.* 2010 Jan 30.
122. Centers for Disease Control and Prevention (CDC). The recommended immunization schedules for persons aged 0 through 18 years, CDC/AAP/AAFP 2010.
- 13. Detecting and treating periodontal disease**
123. Hollister MC, Weintraub JA. The association of oral status with systemic health, quality of life, and economic productivity. *J Dent Educ.* 1993 Dec;57(12):901-12.
124. Ostberg AL, Jarkman K, Lindblad U, Halling A. Adolescents' perceptions of oral health and influencing factors: a qualitative study. *Acta Odontol Scand.* 2002 Jun;60(3):167-73.
125. Percy MS. Oral health of adolescents--it's more than dental caries. *MCN Am J Matern Child Nurs.* 2008 Jan-Feb;33(1):26-31.
126. Peres KG, Peres MA, Araujo CL, et al. Social and dental status along the life course and oral health impacts in adolescents: a population-based birth cohort. *Health Qual Life Outcomes.* 2009 Nov 22;7:95.
127. de Paula Júnior DF, Santos NC, da Silva ET, et al. Psychosocial impact of dental esthetics on quality of life in adolescents. *Angle Orthod.* 2009 Nov;79(6):1188-93.
128. Saliba-Garbin CA, Ispér-Garbin AJ, Moreira-Arcieri R, et al. Adolescents' perception of oral health. *Rev Salud Pública (Bogotá).* 2009 Mar-Apr;11(2):268-77.
129. Brukiene V, Aleksejneiene J. An overview of oral health promotion in adolescents. *Int J Paediatr Dent.* 2009 May;19(3):163-71.
130. Walsh T, Worthington HV, Glenny AM, et al. Fluoride toothpastes of different concentrations for preventing dental caries in children and adolescents. *Cochrane Database Syst Rev.* 2010 Jan 20;(1):CD007868.
131. Christensen LB, Twetman S, Sundby A. Oral health in children and adolescents with different socio-cultural and socio-economic backgrounds. *Acta Odontol Scand.* 2010 Jan;68(1):34-42.
- 14. Preventing pregnancy in adolescents**
132. Sant'Anna MJ, Carvalho KA, Melhado A, et al. Teenage pregnancy: impact of the integral attention given to the pregnant teenager and adolescent mother as a protective factor for repeat pregnancy. *Scientific World Journal.* 2007 Feb 9;7:187-94.
133. Milne D, Glasier A. Preventing repeat pregnancy in adolescents. *Curr Opin Obstet Gynecol.* 2008 Oct;20(5):442-6.
134. Chedraui P. Pregnancy among young adolescents: trends, risk factors and maternal-perinatal outcome. *J Perinat Med.* 2008;36(3):256-9.
135. Oringanje C, Meremikwu MM, Eko H, Esu E, et al. Interventions for preventing unintended pregnancies among adolescents. *Cochrane Database Syst Rev.* 2009 Oct 7;(4):CD005215.
136. Barnet B, Liu J, DeVoe M, Duggan AK, et al. Motivational intervention to reduce rapid subsequent births to adolescent mothers: a community-based randomized trial. *Ann Fam Med.* 2009 Sep-Oct;7(5):436-45.
137. Noll JG, Shenk CE, Putnam KT. Childhood sexual abuse and adolescent pregnancy: a meta-analytic update. *J Pediatr Psychol.* 2009 May;34(4):366-78.
138. No authors listed. Interventions for preventing unintended pregnancies among adolescents. *Obstet Gynecol.* 2010 Jan;115(1):171-2.
- II. Interventions during pregnancy**
- 1. Prenatal check-ups**
139. Villar J et al. The WHO antenatal care randomized controlled trial: rationale and study design. *Paediatric and Perinatal Epidemiology.* 1998, 12 (Suppl.): 27-58.

140. Carroli G, Rooney C, Villar J. How effective is antenatal care in preventing maternal mortality and serious morbidity? *Paediatric and Perinatal Epidemiology*, 2001, 15 (Suppl.1):1-42.
141. Villar J et al. for the WHO Antenatal Care Trial Research Group. WHO antenatal care randomised trial for the evaluation of a new model of routine antenatal care. *The Lancet*, 2001, 357: 1551-1564.
142. Carroli G et al. for the WHO Antenatal Care Trial Research Group. WHO systematic review of randomised controlled trials of routine antenatal care. *The Lancet*, 2001, 357: 1565-1570.
143. Villar J et al. Patterns of routine antenatal care for low-risk pregnancy The Cochrane Library, Issue No. 2. Oxford, Update Software, 2002.
144. Patrones de control prenatal de rutina para embarazos de bajo riesgo. De La Biblioteca Cochrane Plus, número 3, 2005.
145. Ickovics JR, Kershaw TS, Westdahl C, et al. Group prenatal care and perinatal outcomes: a randomized controlled trial. *Obstet Gynecol*. 2007 Aug;110(2 Pt 1):330-9.
146. Raatikainen K, Heiskanen N, Heinonen S. Under-attending free antenatal care is associated with adverse pregnancy outcomes. *BMC Public Health*. 2007 Sep 27;7(147):268.
147. Ickovics JR, Kershaw TS, Westdahl C, et al. Group prenatal care and perinatal outcomes: a randomized controlled trial. *Obstet Gynecol*. 2007 Aug;110(2 Pt 1):330-9.
148. García Alonso López A, Rosales Ortiz S, Jiménez Solís G; Colegio Mexicano de Especialistas en Ginecología y Obstetricia. Clinical practice guidelines 2008. Mexican College of Obstetrics and Gynecology Specialists. Prenatal monitoring in adolescents: risk maternal-infant. *Ginecol Obstet Mex*. 2009 May;77(5):S129-54.
149. Mukhopadhyay S, Wendel J. Are prenatal care resources distributed efficiently across high-risk and low-risk mothers? *Int J Health Care Finance Econ*. 2008 Sep;8(3):163-79.
150. Lauderdale DS, Vanderweele TJ, Siddique J, Lantos JD. Prenatal Care Utilization in Excess of Recommended Levels: Trends From 1985 to 2004. *Med Care Res Rev*. 2009 Nov 13.
151. Neilson JP. WITHDRAWN: Ultrasound for fetal assessment in early pregnancy. *Cochrane Database Syst Rev*. 2010 Apr 14;4:CD000182.
156. Smith GCS, Fretts RC. Stillbirth. *Seminar Vol* 370, November 17, 2007.
157. McClure EM, Wright LL, Goldenberg RL, et al. The global network: a prospective study of stillbirths in developing countries. *Am J Obstet Gynecol*. 2007 Sep;197(3):247.e1-5.
158. Goldenberg RL, McClure EM, Bann CM. The relationship of intrapartum and antepartum stillbirth rates to measures of obstetric care in developed and developing countries. *Acta Obstet Gynecol Scand*. 2007;86(11):1303-9.
159. Facchinetti F, Reddy U, Stray-Pedersen B, et al. International issues in stillbirth. *J Matern Fetal Neonatal Med*. 2008 Jun;21(6):425-8.
160. Flenady V, Frøen JF, Pinar H, et al. An evaluation of classification systems for stillbirth. *BMC Pregnancy Childbirth*. 2009 Jun 19;9:24.
161. McClure EM, Saleem S, Pasha O, Goldenberg RL. Stillbirth in developing countries: a review of causes, risk factors and prevention strategies. *J Matern Fetal Neonatal Med*. 2009 Mar;22(3):183-90.
162. Goldenberg RL, McClure EM, Belizán JM. Commentary: reducing the world's stillbirths. *BMC Pregnancy Childbirth*. 2009 May 7;9 Suppl 1:S1.
163. Yakoob MY, Menezes EV, Soomro T, et al. Reducing stillbirths: behavioural and nutritional interventions before and during pregnancy. *BMC Pregnancy Childbirth*. 2009 May 7;9 Suppl 1:S3.
164. Menezes EV, Yakoob MY, Soomro T, et al. Reducing stillbirths: prevention and management of medical disorders and infections during pregnancy. *BMC Pregnancy Childbirth*. 2009 May 7;9 Suppl 1:S4.
165. Haws RA, Yakoob MY, Soomro T, et al. Reducing stillbirths: screening and monitoring during pregnancy and labour. *BMC Pregnancy Childbirth*. 2009 May 7;9 Suppl 1:S5.
166. Darmstadt GL, Yakoob MY, Haws RA, et al. Reducing stillbirths: interventions during labour. *BMC Pregnancy Childbirth*. 2009 May 7;9 Suppl 1:S6.
167. Bhutta ZA, Darmstadt GL, Haws RA, et al. Delivering interventions to reduce the global burden of stillbirths: improving service supply and community demand. *BMC Pregnancy Childbirth*. 2009 May 7;9 Suppl 1:S7.
168. Cham M, Sundby J, Vangen S. Fetal outcome in severe maternal morbidity: too many stillbirths. *Acta Obstet Gynecol Scand*. 2009;88(3):343-9.
169. No authors listed. NewsCaps: Stillbirths in developing countries decrease when nurses or midwives assist births. *Am J Nurs*. 2010 May;110(5):18.
170. Monari F, Facchinetti F. Management of subsequent pregnancy after antepartum stillbirth. A review. *J Matern Fetal Neonatal Med*. 2010 May 26.

## 2. Preventing fetal death

152. World Health Organization. Definitions and indicators in family planning maternal & child health and reproductive health. Geneva: WHO Press, 2001.
153. Lawn J, Shibuya K, Stein C. No cry at birth: global estimates of intrapartum stillbirths and intrapartum-related neonatal deaths. *Bull World Health Organ* 2005;83:409-17.
154. World Health Organization. Neonatal and perinatal mortality: country, regional and global estimates. Geneva: WHO Press. 2006.
155. McClure EM, Nalubamba-Phiri M, Goldenberg RL. Stillbirth in developing countries. *Int J Gynaecol Obstet*. 2006 Aug;94(2):82-90.

## 3. Preventing preterm birth

171. McCormick MC. The contribution of low birth weight to infant mortality and childhood morbidity. *N Eng J Med* 1985;312:82-90.

172. Goldemberg RL, Rouse DJ. The prevention of premature birth. *N Eng J Med* 1998;339:313-20.
173. Slattery MM, Morrison JJ. Preterm delivery. *Lancet* 2002;360:1489-97.
174. Maloni JA, Damato EG. Reducing the risk for preterm birth: evidence and implications for neonatal nurses. *Adv Neonatal Care*. 2004 Jun;4(3):166-74.
175. Goldemberg RL, Culhane JF. Prepregnancy health status and the risk of preterm delivery. *Arch Pediatr Adolesc Med* 2005;159: 89-90.
176. Ananth CV, Vintzileos AM. Epidemiology of preterm birth and its clinical subtypes. *J Matern Fetal Neonatal Med* 2006;19:773-82.
177. Romero R, Espinoza J, Kusanovic J, et al. The preterm parturition syndrome. *BJOG* 2006;113:17-42.
178. Behrman RE, Butler AS. Preterm Birth: causes, consequences and prevention. IOM, National Academies Press, 2007. Washington DC. <http://www.nap.edu>.
179. Catov JM, Roberta B. Ness RB, Lisa M. Bodnar LM et al. Association of Periconceptional Multivitamin Use and Risk of Preterm or small-for-Gestational-Age Births. *Am J Epidemiol* 2007;166:296-303.
180. Goldenberg RL, Culhane JF, Iams JD, Romero R. Epidemiology and causes of preterm birth. *Lancet*. 2008 Jan 5;371(9606):75-84.
181. McElrath TF, Hecht JL, Dammann O, et al. Pregnancy disorders that lead to delivery before the 28th week of gestation: an epidemiologic approach to classification. *Am J Epidemiol*. 2008 Nov 1;168(9):980-9.
182. Damus K. Prevention of preterm birth: a renewed national priority. *Curr Opin Obstet Gynecol*. 2008 Dec;20(6):590-6.
183. Torloni MR, Betrán AP, Daher S, et al. Maternal BMI and preterm birth: a systematic review of the literature with meta-analysis. *J Matern Fetal Neonatal Med*. 2009 Nov;22(11): 957-70.
184. Shah PS, Zao J; Knowledge Synthesis Group of Determinants of preterm/LBW births. Induced termination of pregnancy and low birthweight and preterm birth: a systematic review and meta-analyses. *BJOG*. 2009 Oct;116(11):1425-42.
185. Offenbacher S, Beck JD, Jared HL, et al. Effects of periodontal therapy on rate of preterm delivery: a randomized controlled trial. *Obstet Gynecol*. 2009 Sep;114(3):551-9.
186. Zhang Q, Ananth CV, Li Z, Smulian JC. Maternal anaemia and preterm birth: a prospective cohort study. *Int J Epidemiol*. 2009 Oct;38(5):1380-9. Epub 2009 Jul 3.
187. Kramer MS, Kahn SR, Platt RW, Genest J, et al. Antioxidant vitamins, long-chain fatty acids, and spontaneous preterm birth. *Epidemiology*. 2009 Sep;20(5):707-13.
188. Novák T, Radnai M, Gorzó I, et al. Prevention of preterm delivery with periodontal treatment. *Fetal Diagn Ther*. 2009;25(2):230-3. Epub 2009 May 28.
189. Guerra GV, Cecatti JG, Souza JP, et al. Factors and outcomes associated with the induction of labour in Latin America. *BJOG*. 2009 Dec;116(13):1762-72.
190. Owen J, Hankins G, Iams JD, et al. Multicenter randomized trial of cerclage for preterm birth prevention in high-risk women with shortened midtrimester cervical length. *Am J Obstet Gynecol*. 2009 Oct;201(4):375.e1-8.
191. Polyzos NP, Polyzos IP, Mauri D, et al. Effect of periodontal disease treatment during pregnancy on preterm birth incidence: a metaanalysis of randomized trials. *Am J Obstet Gynecol* Vol 200(3);225-32 March 2009.
192. Ashton DM, Lawrence HC, Adams NL. Surgeon General's Conference on the Prevention of Preterm Birth. *Obstet Gynecol* Vol.113, No. 4, APRIL 2009.
193. Beck S, Wojdyla D, Say L, Betran AP, Merialdi M, et al. The worldwide incidence of preterm birth, a systematic review of maternal mortality and morbidity. *Bull World Health Organ* 2010;88:31-38.
194. Muglia L, Katz M. The enigma of spontaneous preterm birth. *NEJM* 11 Feb 2010;236(6):529/535.
195. Dodd JM, Flenady V, Cincotta R, Crowther CA. Prenatal administration of progesterone for preventing preterm birth in women considered to be at risk of preterm birth. *Cochrane Review* 2010.
196. Macones GA. Elective delivery before 39 weeks: reason for Caution. *Am J Obstet Gynecol MARCH* 2010.
197. Barros FC, Bhutta ZA, Batra M, et al. Global report on preterm birth and stillbirth (3 of 7): evidence for effectiveness of interventions. *BMC Pregnancy and Childbirth* 2010, 10(Suppl 1):S3doi.
198. The Ohio Perinatal Quality Collaborative Writing committee. A statewide initiative to reduce inappropriate scheduled births at 36 0/37-38 6/7 weeks' gestation. *Am J Obstet Gynecol* March 2010;202:243.e1-8.

#### 4. Managing premature rupture of membranes

199. IMPAC. Manejo de las complicaciones del embarazo y el parto: Guia para obstetricas y medicos. OMS/OPS, FNUAP,UNICEF, Banco Mundial 2000.
200. Kenyon S, Boulvain M, Neilson J. Antibiotics for preterm premature rupture of membranes. *Cochrane Database Syst Rev*. 2001;(4):CD001058.
201. Martínez Guillen F. Amenaza y trabajo de Parto Pretérmino. En, Martínez Guillen F. "Medicina Perinatal basada en Evidencia", Publicación auspiciada por OPS/OMS, Litografía Nicaragüense (Litonic), 2005, p 18-32.
202. Royal College of Obstetricians and Gynaecologists (RCOG). Prevention of early onset neonatal group B streptococcal disease. London (UK); NGC, [www.guideline.gov](http://www.guideline.gov), USA. Date Modified: 4/2/2007.
203. American College of Obstetricians and Gynecologists (ACOG). Premature rupture of membranes. Washington (DC), ACOG practice bulletin; no. 1) National Guideline Clearinghouse, [www.guideline.gov](http://www.guideline.gov) USA. Date Modified: 4/2/2008.

204. Kenyon S, Boulvain M, Neilson J. Antibiotics for preterm rupture of membranes. *Cochrane Library* 2010.
205. Hass DM. Antibiotic treatment for preterm rupture of the membranes. *BMJ Clinical Evidence*, 2010.
206. Yudin MH, van Schalkwyk J, Van Eyk N, et al. Antibiotic therapy in preterm premature rupture of the membranes. *J Obstet Gynaecol Can*. 2009 Sep;31(9):863-7, 868-74.
207. Abou El Senoun G, Dowswell T, Mousa HA. Planned home versus hospital care for preterm prelabour rupture of the membranes (PPROM) prior to 37 weeks' gestation. *Cochrane Database Syst Rev*. 2010 Apr 14;4:CD008053.
208. Cousens S, Blencowe H, Gravett M, Lawn JE. Antibiotics for pre-term pre-labour rupture of membranes: prevention of neonatal deaths due to complications of pre-term birth and infection. *Int J Epidemiol*. 2010 Apr;39 Suppl 1:i134-43.
- 5. Using the CLAP/WR perinatal clinical record (PCR) and perinatal card**
209. Schwarcz R, Gonzalo Diaz A, Fescina R, y col. Historia Clínica Perinatal Simplificada. Propuesta de un modelo para la atención primaria de baja complejidad. *Bol Oficina Sanit Panam* 1983;95(2):163-172.
210. Centro Latinoamericano de Perinatología y Desarrollo Humano. Sistema Informático Perinatal (SIP). CLAP/OPS-OMS, Publ. Cient. No. 1524, agosto 2005.
- 6. Weight gain during pregnancy**
211. Martínez Guillen F. Nutrición en la embarazada y su impacto en la niñez. Medicina Perinatal Basada en Evidencia, Litografía Nicaragüense, Managua 2005.
212. Dietz PM, Callaghan WM, Cogswell ME, et al. Combined effects of prepregnancy body mass index and weight gain during pregnancy on the risk of preterm delivery. *Epidemiology*. 2006 Mar;17(2):170-7.
213. Hedderson MM, Weiss NS, Sacks DA, Pettitt DJ. Pregnancy Weight Gain and Risk of Neonatal Complications. *Obstet Gynecol* 2006;108:1153-61.
214. DeVader SR, Neeley HL, Myles TD, Leet TL. Evaluation of gestational weight gain guidelines for women with normal prepregnancy body mass index. *Obstet Gynecol*. 2007 Oct;110(4):745-51.
215. Kiel DW, Dodson EA, Artal R, et al. Gestational weight gain and pregnancy outcomes in obese women: how much is enough? *Obstet Gynecol*. 2007 Oct;110(4):752-8.
216. Rode L, Hegaard HK, Kjaergaard H, et al. Association between maternal weight gain and birth weight. *Obstet Gynecol*. 2007 Jun;109(6):1309-15.
217. Viswanathan M, Siega-Riz AM, Moos MK, et al. Outcomes of maternal weight gain. *Evid Rep Technol Assess (Full Rep)*. 2008 May;(168):1-223.
218. Olson CM. Achieving a healthy weight gain during pregnancy. *Annu Rev Nutr*. 2008;28:411-23.
219. Amorim AR, Linné Y, Kac G, Lourenço PM. Assessment of weight changes during and after pregnancy: practical approaches. *Matern Child Nutr*. 2008 Jan;4(1):1-13.
220. Nohr EA, Vaeth M, Baker JL, et al. Combined associations of pregnancy body mass index and gestational weight gain with the outcome of pregnancy. *Am J Clin Nutr*. 2008 Jun;87(6):1750-9.
221. Chasan-Taber L, Schmidt MD, Pekow P, et al. Predictors of excessive and inadequate gestation weight gain in Hispanic women. *Obesity (Silver Spring)*. 2008 Jul;16(7):1657-66.
222. Wax JR. Risks and management of obesity in pregnancy: current controversies. *Curr Opin Obstet Gynecol*. 2009 Apr;21(2):117-23.
223. Institute of Medicine of the National Academies. Weight gain during pregnancy: Reexamining the Guidelines. Report Brief, May 2009.
224. Stothard KJ, Tennant PW, Bell R, Ranking J. Maternal Overweight and Obesity and Risk of Congenital Anomalies. A systematic Review and Meta-analysis. *JAMA* 2009; 301(6) 636-650.
225. Baker VJL, Sørensen T I, Olsen J, and Rasmussen KM. Pregnancy outcomes related to gestational weight gain in women defined by their body mass index, parity, height, and smoking status. *Am J Clin Nutr* 2009;90:1288-94.
226. Bodnar LM, Wisner KL, Moses-Kolko E et al. Prepregnancy body mass index, gestational weight gain, and the likelihood of major depressive disorder during pregnancy. *J Clin Psychiatry*. 2009 Sep;70(9):1290-6.
227. Caughey RW, Michels KB. Birth weight and childhood leukemia: A meta analysis and review of the current evidence. Department of Epidemiology, Harvard School of Public Health. *Int J Cancer* 2009, 124:2658-2670.
228. Barker DJP, Osmond C, Kajante E, Eriksson JG. Growth and chronic disease: findings in the Helsinki Birth Cohort. *Ann Hum Biol Sep-Oct 2009;36(5)* 445.
229. Prevention and management of obesity (mature adolescents and adults). *National Guideline Clearinghouse* 2009. 11/23/2009 www.guideline.gov.
230. Horta B, Gigante DP, Osmond C, et al. Intergenerational effect of weight gain in childhood on offspring birthweight. *Internat J Epidemiol* 2009;38:724-732.
231. Siega-Riz A, Knaack J, Viswanathan M, et al. A systematic review of outcomes of maternal weight gain according to the Institute of Medicine recommendations: birthweight, fetal growth, and postpartum weight retention. *Am J Obstet Gynecol Oct 2009;201:339.e1-14*.
232. Artal R, Lockwood CJ, Brown HL. Weight Gain Recommendations in Pregnancy and the Obesity Epidemic. *Obst Gynecol VOL. 115, No. 1, JAN 2010*.
233. Davies GA, Maxwell C, McLeod L, et al. Obesity in pregnancy. *J Obstet Gynaecol Can*. 2010 Feb;32(2):165-73.
234. Shaikh H, Robinson S, Teoh TG. Management of maternal obesity prior to and during pregnancy. *Seminars in Fetal and Neonatal Medicine Vol 15, Issue 2, April 2010, Pag 77-82*.

235. Fitzsimons KJ, Modder J. Setting maternity care standards for women with obesity in pregnancy. *Seminars in Fetal and Neonatal Medicine* Vol 15, Issue 2, April 2010, Pag 100-107.
236. McGuire W, Dyson L, Renfrew M. Maternal obesity: consequences for children, challenges for clinicians and carers. *Seminars in Fetal and Neonatal Medicine* Vol 15, Issue 2, April 2010, Pag 108-112.
- 7. Reducing the use of unnecessary caesarean section**
237. Barros FC, Victora CG, Barros A JD et al. The challenge of reducing neonatal mortality in middle-income countries: findings from three Brazilian birth cohorts in 1982, 1993, and 2004. *Lancet* Vol 365 Issue 9462, 5 March 2005, pag 847-854.
238. Villar J, Valladares E, Wojdyla D et al. Caesarean delivery rates and pregnancy outcomes: the 2005 WHO global survey on maternal and perinatal health in Latin America. *Lancet* Vol 367, Issue 9525, 3-9 June 2006, Pages 1819-1829.
239. Chaillet N, Dumont A. Evidence-Based Strategies for Reducing Cesarean Section Rates: A Meta-Analysis. *BIRTH* 34:1, 53-64 March 2007.
240. Hansen AK, Wisborg K, et al. Elective caesarean section and respiratory morbidity in the term and near-term neonate. *Acta Obstetricia et Gynecologica*. 2007; 86: 389\_394.
241. Damus K. Prevention of Preterm birth: a renewed national priority. *Current Opinion Obstet Gynecol* 2008;20: 590.
242. Malloy MH. Impact of Cesarean Section on Intermediate and Late Preterm Births: United States, 2000–2003. *BIRTH* 36:1, 26-33 March 2009.
243. Pulver LS, Guest-Warnick G, Stoddard GJ, et al. Weight for Gestational Age Affects the Mortality of Late Preterm Infants. *Pediatrics*. Volume 123, Number 6, June 2009 e1072-77.
244. Darmstadt GL, Yakoob MY, Haws RA, Menezes EV, Soomro T, Bhutta ZA. Reducing stillbirths: interventions during labour. *BMC Pregnancy Childbirth*. 2009; 9(Suppl 1): S6.
245. Karlstrom A, Radestad M L, Eriksson C, et al. Cesarean Section without Medical Reason, 1997 to 2006: A Swedish Register Study. *BIRTH* 37:1, 11-19 March 2010.
246. Tzur T, Weinraub AY, Sheiner E, Wiznitzer A. Timing of elective repeat caesarean section: maternal and neonatal morbidity and mortality. *J Matern Fetal Neonatal Med* 2010 Early Online, 1-7.
247. Wilimink FA, Hukkelhoven CH, Lunshof S et al. Neonatal outcome following elective cesarean section beyond 37 weeks of gestation: a 7-year retrospective analysis of a national registry. *Am J Obst Gynecol*. March 2010; Vol 202, No 3, 250.
248. NIH Consensus Development Conference on Vaginal Birth After Cesarean: New Insights. *National Institute of Health (NIH)*, March 8–10, 2010 Bethesda, Maryland USA.
249. Cesarean Rates Reach Record High, US. Health Day. *Medline Plus March* 2010.

## 8. Immunization with tetanus toxoid, taking previous vaccination status into account

250. Vandelaer J, Birmingham M, Gasse F, et al. Tetanus in developing countries: an update on the Maternal and Neonatal Tetanus Elimination Initiative. *Vaccine*. 2003 Jul 28;21(24):3442-5.
251. Pichichero ME, Rennels MB, Edwards KM et al. Combined tetanus, diphtheria, and 5-component pertussis vaccine for use in adolescents and adults. *JAMA*. 2005 Jun 22;293(24):3003-11.
252. Organización Panamericana de la Salud. Eliminación del tétanos neonatal: Guía práctica. OPS/OMS 2005. Publicación científica No. 602.
253. Pichichero ME, Blatter MM, Kennedy WA, et al. Acellular pertussis vaccine booster combined with diphtheria and tetanus toxoids for adolescents. *Pediatrics*. 2006 Apr;117(4):1084-93.
254. Roper MH, Vandelaer JH, Gasse FL. Maternal and neonatal tetanus. *Lancet*. 2007 Dec 8;370(9603):1947-59.
255. Murphy TV, Slade BA, Broder KR, Kretsinger K, et al. Prevention of Pertussis, Tetanus, and Diphtheria Among Pregnant and Postpartum Women and their Infants Recommendations of the Advisory Committee on Immunization Practices (ACIP). *MMWR Recomm Rep*. 2008 May 30;57(RR-4):1-47.
256. Saffar M, Khalilian A, Ajami A, et al. Seroimmunity to diphtheria and tetanus among mother-infant pairs; the role of maternal immunity on infant immune response to diphtheria-tetanus vaccination. *Swiss Med Wkly*. 2008 May 3;138(17-18):256-60.
257. Gall SA. Vaccines for pertussis and influenza: recommendations for use in pregnancy. *Clin Obstet Gynecol*. 2008 Sep;51(3):486-97.
258. Za'tsev EM, Krasnoproschina LI, Astakhova TI, Zakharova NS. Monitoring of antibodies against diphtheria, tetanus and pertussis in pregnant women. *Zh Mikrobiol Epidemiol Immunobiol*. 2010 Jan-Feb;(1):32-5.
- 9. Preventing influenza by vaccination**
259. McNeil SA, Dodds L, Allen VM, et al. Influenza vaccine programs and pregnancy: new Canadian evidence for immunization. *J Obstet Gynaecol Can*. 2007 Aug;29(8):674-6.
260. Zaman K, Roy E, Arifeen SE, et al. Effectiveness of maternal influenza immunization in mothers and infants. *N Engl J Med*. 2008 Oct 9;359(15):1555-64.
261. Mak TK, Mangtani P, Leese J, et al. Influenza vaccination in pregnancy: current evidence and selected national policies. *Lancet Infect Dis*. 2008 Jan;8(1):44-52.
262. MacDonald NE, Riley LE, Steinhoff MC. Influenza immunization in pregnancy. *Obstet Gynecol*. 2009 Aug;114(2 Pt 1):365-8.
263. [No authors listed]. Organizations encourage seasonal flu vaccine, H1N1 vaccine for pregnant women. *Am Fam Physician*. 2009 Nov 1;80(9):913.

264. Tamma PD, Ault KA, del Rio C, et al. Safety of influenza vaccination during pregnancy. *Am J Obstet Gynecol.* 2009 Dec;201(6):547-52.
265. CDC. Interim Guidance - Pregnant women and swine influenza: considerations for clinicians. Centers for Disease Control and Prevention. Available at [http://www.cdc.gov/swineflu/clinician\\_pregnant.htm](http://www.cdc.gov/swineflu/clinician_pregnant.htm). Accessed April 29, 2009.
266. CDC. Interim Guidance: Considerations Regarding 2009 H1N1 Influenza in Intrapartum and Postpartum Hospital Settings. November 10, 2009.
267. Gruslin A, Steben M, Halperin S. Immunization in Pregnancy. Society of Obstetricians and Gynaecologists of Canada. SOGC CLINICAL PRACTICE GUIDELINE. *J Obstet Gynaecol Can* 2009;31(11):1085-1092.
268. FDA y CDC. Use of Influenza A (H1N1) 2009 Monovalent Influenza Vaccine in Pregnant Women. October 27, 2009.
269. Creanga AA, Johnson TF, Graifcer SB, et al. Severity of 2009 pandemic influenza A (H1N1) virus infection in pregnant women. *Obstet Gynecol.* 2010 Apr;115(4):717-26.
270. Satpathy HK, Lindsay M, Kawaiss JF. Novel H1N1 virus infection and pregnancy. *Postgrad Med.* 2009 Nov;121(6):106-12.
271. Steinhoff MC, Omer SB, Roy E, et al. Influenza immunization in pregnancy--antibody responses in mothers and infants. *N Engl J Med.* 2010 Apr 29;362(17):1644-6.
272. Siston AM, Rasmussen SA, Honein MA, et al. Pandemic 2009 influenza A(H1N1) virus illness among pregnant women in the United States. *JAMA.* 2010 Apr 21;303(15):1517-25.
273. Charlier C, Launay O, Coignard-Biebler H, et al. Pregnancy: a high risk factor in influenza infection. *Med Sci (Paris).* 2010 Jan;26(1):100-4.
274. CDC. 2009 pandemic influenza A (H1N1) in pregnant women requiring intensive care - New York City, 2009. *MMWR Morb Mortal Wkly Rep.* 2010 Mar 26;59(11):321-6.
- 10. Syphilis screening and case management**
275. Carey JC. Congenital syphilis in the 21st century. *Curr Womens Health Rep.* 2003 Aug;3(4):299-302.
276. Bhutta Z, Dramstadt G, Hasan B, Haws R. Community-based interventions for improving perinatal and neonatal outcomes in developing countries: a review of the evidence. *Pediatrics* 2005;115:519-64.
277. Majeroni BA, Ukkadam S. Screening and treatment for sexually transmitted infections in pregnancy. *Am Fam Physician.* 2007 Jul 15;76(2):265-70.
278. Carles G, Lochet S, Youssef M et al. Syphilis and pregnancy. *J Gynecol Obstet Biol Reprod (Paris).* 2007 Nov 12.
279. Leslie DE, Azzato F, Karapanagiotidis T, et al. Development of a real-time PCR assay to detect *Treponema pallidum* in clinical specimens and assessment of the assay's performance by comparison with serological testing. *J Clin Microbiol.* 2007 Jan;45(1):93-6.
280. O'Connor M, Kleinman S, Goff M. Syphilis in pregnancy. *J Midwifery Womens Health.* 2008 May-Jun;53(3):e17-21.
281. Chakraborty R, Luck S. Syphilis is on the increase: the implications for child health. *Arch Dis Child.* 2008 Feb;93(2):105-9.
282. Organización Mundial de la Salud. Eliminación mundial de la sífilis congénita: fundamentos y estrategia para la acción. OMS 2008.
283. U.S. Preventive Services Task Force. Screening for syphilis infection in pregnancy: U.S. Preventive Services Task Force reaffirmation recommendation statement. *Ann Intern Med.* 2009 May 19;150(10):705-9.
284. No authors listed. Summaries for patients. Screening pregnant women for syphilis infection: U.S. Preventive Services Task Force recommendation. *Ann Intern Med.* 2009 May 19;150(10):I-40.
285. Gayet-Ageron A, Ninet B, Toutous-Trellu L, et al. Assessment of a real-time PCR test to diagnose syphilis from diverse biological samples. *Sex Transm Infect.* 2009 Aug;85(4):264-9.
286. Lomotey CJ, Lewis J, Gebrian B, et al. Maternal and congenital syphilis in rural Haiti. *Rev Panam Salud Publica.* 2009 Sep;26(3):197-202.
287. Díaz-Olavarrieta C, Wilson KS, García SG, et al. The co-occurrence of intimate partner violence and syphilis among pregnant women in Bolivia. *J Womens Health (Larchmt).* 2009 Dec;18(12):2077-86.
288. Macêdo VC, Bezerra AF, Frias PG, Andrade CL. Evaluation of measures to prevent vertical transmission of HIV and syphilis in public maternity hospitals in four municipalities in Northeast Brazil. *Cad Saude Publica.* 2009 Aug;25(8):1679-92.
289. Villazón-Vargas N, Conde-Glez CJ, Juárez-Figueroa L, Uribe-Salas F. Evaluation of a rapid diagnostic test to assess the prevalence of maternal syphilis in Bolivia. *Rev Med Chil.* 2009 Apr;137(4):515-21.
290. CDC. Congenital syphilis - United States, 2003-2008. *MMWR Morb Mortal Wkly Rep.* 2010 Apr 16;59(14):413-7.
291. Goldenberg RL, McClure EM, Saleem S, Reddy UM. Infection-related stillbirths. *Lancet.* 2010 Apr 24;375(9724):1482-90.
292. Stamm LV. Global challenge of antibiotic-resistant *Treponema pallidum*. *Antimicrob Agents Chemother.* 2010 Feb;54(2):583-9.
- 11. Detecting and treating symptomatic and asymptomatic bacteriuria**
293. Gilstrap LC 3rd, Ramin SM. Urinary tract infections during pregnancy. *Obstet Gynecol Clin North Am.* 2001 Sep;28(3):581-91.
294. Krcmery S, Hromec J, Demesova D. Treatment of lower urinary tract infection in pregnancy. *Int J Antimicrob Agents.* 2001 Apr;17(4):279-82.
295. Smail F, Vazquez JC. Antibiotics for asymptomatic bacteriuria in pregnancy. *Cochrane Database Syst Rev.* 2007 Apr 18;(2):CD000490.

296. Quiroga-Feuchter G, Robles-Torres RE, Ruelas-Morán A, Gómez-Alcalá AV. Asymptomatic bacteriuria among pregnant women. An underestimated threat. *Rev Med Inst Mex Seguro Soc.* 2007 Mar-Apr;45(2):169-72.
297. Ochoa-Brust GJ, Fernández AR, Villanueva-Ruiz GJ, et al. Daily intake of 100 mg ascorbic acid as urinary tract infection prophylactic agent during pregnancy. *Acta Obstet Gynecol Scand.* 2007;86(7):783-7.
298. Hazhir S. Asymptomatic bacteriuria in pregnant women. *Urol J.* 2007 Winter;4(1):24-7.
299. Macejko AM, Schaeffer AJ. Asymptomatic bacteriuria and symptomatic urinary tract infections during pregnancy. *Urol Clin North Am.* 2007 Feb;34(1):35-42.
300. Macejko AM, Schaeffer AJ. Asymptomatic bacteriuria and symptomatic urinary tract infections during pregnancy. *Urol Clin North Am.* 2007 Feb;34(1):35-42.
301. Swadpanich U, Lumbiganon P, Prasertcharoensook W, Laopaiboon M. Antenatal lower genital tract infection screening and treatment programs for preventing preterm delivery. *Cochrane Database of Systematic Reviews*, Issue 2, 2008.
302. Schnarr J, Smaill F. Asymptomatic bacteriuria and symptomatic urinary tract infections in pregnancy. *Eur J Clin Invest.* 2008 Oct;38 Suppl 2:50-7.
303. Sheiner E, Mazor-Drey E, Levy A. Asymptomatic bacteriuria during pregnancy. *J Matern Fetal Neonatal Med.* 2009 May;22(5):423-7.
304. Grabe M, Bishop MC, Bjerklund-Johansen TE et al Guidelines on Urological Infections. *European Association of Urology (EAU);2009. National Guidelines Clearinghouse.* www.guideline.gov. Date . Modified: 6/22/2009.
305. Mignini L, Carroli G, Abalos E, et al. Accuracy of diagnostic tests to detect asymptomatic bacteriuria during pregnancy. *Obstet Gynecol.* 2009 Feb;113(2 Pt 1):346-52.
306. Smaill FM, Vazquez JC. Antibiotics for asymptomatic bacteriuria in pregnancy. *Cochrane Library* 2010, Issue 1.
307. Bahadi A, El Kabbaj D, Elfazazi H, et al. Urinary tract infection in pregnancy. *Saudi J Kidney Dis Transpl.* 2010 Mar;21(2):342-4.
308. Haider G, Zehra N, Munir AA, Haider A. Risk factors of urinary tract infection in pregnancy. *J Pak Med Assoc.* 2010 Mar;60(3):213-6.
- 12. Preventing and treating anemia with iron**
309. Leblanc CP, Rioux FM. Iron deficiency anemia following prenatal nutrition interventions. *Can J Diet Pract Res.* 2007 Winter;68(4):222-5.
310. Heilmann L. Maternal hemoglobin and pregnancy outcome. *Clin Appl Thromb Hemost.* 2006 Apr;12(2):241.
311. El Guindi W, Pronost J, Carles G, et al. Severe maternal anemia and pregnancy outcome. *J Gynecol Obstet Biol Reprod (Paris).* 2004 Oct;33(6 Pt 1):506-9.
312. Meda N, Dao Y, Touré B, Yameogo B, et al. Assessing severe maternal anemia and its consequences: the value of a simple examination of the coloration of palpebral conjunctiva. *Sante.* 1999 Jan-Feb;9(1):12-7.
313. Patra S, Pasrija S, Trivedi SS, Puri M. Maternal and perinatal outcome in patients with severe anemia in pregnancy. *Int J Gynaecol Obstet.* 2005 Nov;91(2):164-5.
314. Boy E. Severe anemia in pregnancy. IDRC/CRDI, 2000
315. Levy A, Fraser D, Katz M, Mazor M, Sheiner E. Maternal anemia during pregnancy is an independent risk factor for low birthweight and preterm delivery. *Eur J Obstet Gynecol Reprod Biol.* 2005 Oct 1;122(2):182-6.
316. Peña-Rosas JP, Viteri FE. Effects of routine oral iron supplementation with or without folic acid for women during pregnancy. *Cochrane Database Syst Rev.* 2006 Jul 19;3:CD004736.
317. Reveiz L, Gyte GM, Cuervo LG. Treatments for iron-deficiency anaemia in pregnancy. *Cochrane Database Syst Rev.* 2007 Apr 18;(2):CD003094.
318. Peña-Rosas JP, Viteri FE. Effects and safety of preventive oral iron or iron+folic acid supplementation for women during pregnancy. *Cochrane Database Syst Rev.* 2009 Oct 7;(4):CD004736.
- 13. Preventing preeclampsia and eclampsia with calcium (in populations with diets low in calcium)**
319. Atallah AN, Hofmeyr GJ, Duley L. Calcium supplementation during pregnancy for preventing hypertensive disorders and related problems. *Cochrane Database Syst Rev.* 2000;(3):CD001059.
320. Sukonpan K, Phupong V. Serum calcium and serum magnesium in normal and preeclamptic pregnancy. *Arch Gynecol Obstet.* 2005 Nov;273(1):12-6.
321. Duckitt K, Harrington D. Risk factors for pre-eclampsia at antenatal booking: systematic review of controlled studies. *BMJ* 2005;330:565 (12 March).
322. Suplementación con calcio durante el embarazo para la prevención de trastornos hipertensivos y problemas relacionados. De La Biblioteca Cochrane Plus, número 3, 2005.
323. Beinder E. Calcium-supplementation in pregnancy--is it a must? *Ther Umsch.* 2007 May;64(5):243-7.
324. Hofmeyr GJ, Atallah AN, Duley L. Calcium supplementation during pregnancy for preventing hypertensive disorders and related problems. *Cochrane Database Syst Rev.* 2006 Jul 19;3:CD001059.
325. Hofmeyr GJ, Atallah AN, Duley L. Calcium supplementation during pregnancy for preventing hypertensive disorders and related problems. *Cochrane Database Syst Rev.* 2006 Jul 19;3:CD001059.
326. Villar J, Abdel-Aleem H, Meraldi M, Mathai M, et al. World Health Organization randomized trial of calcium supplementation among low calcium intake pregnant women. *Am J Obstet Gynecol.* 2006 Mar;194(3):639-49.

327. Punthumapol C, Kittichotpanich B. Serum calcium, magnesium and uric acid in preeclampsia and normal pregnancy. *J Med Assoc Thai.* 2008 Jul;91(7):968-73.
328. Magee LA, Helewa M, Moutquin JM, Dadelszen P et al. Diagnosis, evaluation, and Management of the Hypertensive Disorders of pregnancy. *SOGC Clinical Practice Guideline No 206. JOGC (The Official voice of reproductive health care in Canada)* March 2008.
329. Meads CA, Cnosen JS, Meher S, Juarez-Garcia A, et al. Methods of prediction and prevention of pre-eclampsia: systematic reviews of accuracy and effectiveness literature with economic modeling. NIHR Health Technology Assessment programme, NHS, UK. *Health Technol Assess* 2008;12(6) :1-27.
330. Duley L. Calcium supplementation. Pre-eclampsia and hypertension. *Pregnancy and childbirth. BMJ Clin Evidence* 2008.
331. Ettinger AS, Lamadrid-Figueroa H, Téllez-Rojo MM, et al. Effect of calcium supplementation on blood lead levels in pregnancy: a randomized placebo-controlled trial. *Environ Health Perspect.* 2009 Jan;117(1):26-31.
332. Kumar A, Devi SG, Batra S, et al. Calcium supplementation for the prevention of pre-eclampsia. *Int J Gynaecol Obstet.* 2009 Jan;104(1):32-6.
333. Duley L. The global impact of pre-eclampsia and eclampsia. *Semin Perinatol.* 2009 Jun;33(3):130-7.
334. Briceño-Pérez C, Briceño-Sanabria L, Vigil-De Gracia P. Prediction and prevention of preeclampsia. *Hypertens Pregnancy.* 2009 May;28(2):138-55.
335. Jain S, Sharma P, Kulshreshtha S, The role of calcium, magnesium, and zinc in pre-eclampsia. *Biol Trace Elem Res.* 2010 Feb;133(2):162-70.
336. Hofmeyer GJ, Atallah AN, Duley L. Calcium Supplementation during pregnancy for prevent hypertensive disorders and related problems. Cochrane Database of Systematic Reviews 2006, Issue 3. *Cochrane Library* 2010.
341. Knight M, Duley L, Henderson-Smart DJ, King JF. WITHDRAWN: Antiplatelet agents for preventing and treating pre-eclampsia. *Cochrane Database Syst Rev.* 2007 Jul 18;(2):CD000492.
342. Bujold E, Morency AM, Roberge S, et al. Acetylsalicylic acid for the prevention of preeclampsia and intra-uterine growth restriction in women with abnormal uterine artery Doppler: a systematic review and meta-analysis. *J Obstet Gynaecol Can.* 2009 Sep;31(9):818-26.
343. No authors listed. Pre-eclampsia: aspirin beneficial. *Prescribe Int.* 2009 Dec;18(104):274.
344. Duley L, Henderson-Smart DJ, Meher S, King JF. Antiplatelet Agents or preventing pre-eclampsia and its complications. *Cochrane Library 2010.*
345. Duley L. Pre-eclampsia and hypertension. *Prgnancy and childbirth. Antiplatelet drugs. BMJ Clinical Evidence* 2010.
346. Deruelle P, Girard JM, Couty N, Subtil D. Prevention of preeclampsia. *Ann Fr Anesth Reanim.* 2010 Mar;29(3):e31-5.

## **15. Detecting and treating sexually transmitted infections**

347. Read JS, Newell MK. Efficacy and safety of cesarean delivery for prevention of mother-to-child transmission of HIV-1. *Cochrane Database Syst Rev.* 2005 Oct 19;(4):CD005479.
348. Lee C, Gong Y, Brok J, et al. Effect of hepatitis B immunisation in newborn infants of mothers positive for hepatitis B surface antigen: systematic review and meta-analysis. *BMJ.* 2006 Feb 11;332(7537):328-36.
349. European Collaborative Study. Mother-to-child transmission of HIV infection in the era of highly active antiretroviral therapy. *Clin Infect Dis.* 2005 Feb 1;40(3):458-65.
350. Carles G, Lochet S, Youssef M, et al. Syphilis and pregnancy. *J Gynecol Obstet Biol Reprod (Paris).* 2007 Nov 12.
351. Majeroni BA, Ulkadam S. Screening and treatment for sexually transmitted infections in pregnancy. *Am Fam Physician.* 2007 Jul 15;76(2):265-70.
352. Pitsouni E, Iavazzo C, Athanasiou S, Falagas ME. Single-dose azithromycin versus erythromycin or amoxicillin for Chlamydia trachomatis infection during pregnancy: a meta-analysis of randomised controlled trials. *Int J Antimicrob Agents.* 2007 Sep;30(3):213-21.
353. Stringer EM, Chi BH, Chintu N, et al. Monitoring effectiveness of programmes to prevent mother-to-child HIV transmission in lower-income countries. *Bull World Health Organ.* 2008 Jan;86(1):57-62.
354. Cowan FM, Humphrey JH, Ntouzini R, et al. Maternal Herpes simplex virus type 2 infection, syphilis and risk of intra-partum transmission of HIV-1: results of a case control study. *AIDS.* 2008 Jan 11;22(2):193-201.
355. Jalil EM, Pinto VM, Benzaken AS, et al. Prevalence of Chlamydia and Neisseria gonorrhoeae infections in pregnant women in six Brazilian cities. *Rev Bras Ginecol Obstet.* 2008 Dec;30(12):614-9.
356. Bacq Y. Hepatitis B and pregnancy. *Gastroenterol Clin Biol.* 2008 Jan;32(1 Pt 2):S12-9.

357. Yang J, Zeng XM, Men YL, Zhao LS. Elective caesarean section versus vaginal delivery for preventing mother to child transmission of hepatitis B virus--a systematic review. *Virology*. 2008 Aug 28;5:100.
358. Chen MY, Fairley CK, De Guingand D, et al. Screening pregnant women for chlamydia: what are the predictors of infection? *Sex Transm Infect*. 2009 Feb;85(1):31-5.
359. U.S. Preventive Services Task Force. Screening for hepatitis B virus infection in pregnancy: U.S. Preventive Services Task Force reaffirmation recommendation statement. *Ann Intern Med*. 2009 Jun 16;150(12):869-73, W154.
360. Xu WM, Cui YT, Wang L, et al. Lamivudine in late pregnancy to prevent perinatal transmission of hepatitis B virus infection: a multicentre, randomized, double-blind, placebo-controlled study. *J Viral Hepat*. 2009 Feb;16(2):94-103.
361. Kershaw TS, Magriples U, Westdahl C, et al. Pregnancy as a window of opportunity for HIV prevention: effects of an HIV intervention delivered within prenatal care. *Am J Public Health*. 2009 Nov;99(11):2079-86.
362. Kalwij S, Macintosh M, Baraitser P. Screening and treatment of Chlamydia trachomatis infections. *BMJ*. 2010 Apr 21;340:c1915.
363. Schwebke JR, Desmond RA. A randomized controlled trial of partner notification methods for prevention of trichomoniasis in women. *Sex Transm Dis*. 2010 Jun;37(6):392-6.
- 16. Antiretrovirals to reduce risk of vertical transmission of HIV**
364. Nogueira AS, Abreu T, Oliveira R, et al. Successful prevention of HIV transmission from mother to infant in Brazil using a multidisciplinary team approach. *Brazilian Journal of Infectious Diseases*, 2001;5(2):78-86.
365. Brocklehurst P. Interventions for reducing the risk of mother-to-child transmission of HIV infection. *Cochrane Database Syst Rev*. 2002;(1): CD000102.
366. Brocklehurst P, Volmink J. Antiretrovirals for reducing the risk of mother-to-child transmission of HIV infection. *Cochrane Database Syst Rev*. 2002;(2):CD003510.
367. Suksomboon N, Poolsup N, Ket-Aim S. Systematic review of the efficacy of antiretroviral therapies for reducing the risk of mother-to-child transmission of HIV infection. *J Clin Pharm Ther*. 2007 Jun;32(3):293-311.
368. Volmink J, Siegfried NL, van der Merwe L, Brocklehurst P. Antiretrovirals for reducing the risk of mother-to-child transmission of HIV infection. *Cochrane Database Syst Rev*. 2007 Jan 24;(1):CD003510.
369. Wilkin TJ, Shalev N, Tieu HV, Hammer SM. Advances in antiretroviral therapy. *Top HIV Med*. 2010 Apr-May;18(2):66-92.
371. Slocum J, Barcio L, Darany J, et al. Preconception to postpartum: management of pregnancy complicated by diabetes. *Diabetes Educ*. 2004 Sep-Oct;30(5):740, 742-4, 747-53.
372. Nazer Herrera J, García Huidobro M, Cifuentes Ovalle L. Congenital malformations among offspring of diabetic women. *Rev Med Chil*. 2005 May;133(5):547-54.
373. Stotland NE, Cheng YW, Hopkins LM, Caughey AB. Gestational weight gain and adverse neonatal outcome among term infants. *Obstet Gynecol*. 2006 Sep;108(3 Pt 1):635-43.
374. HAPO Study Cooperative Research Group. Hyperglycemia and adverse pregnancy outcomes. *N Engl J Med*. 2008 May 8;358(19):1991-2002.
375. Hedderson MM, Weiss NS, Sacks DA, et al. Pregnancy weight gain and risk of neonatal complications: macrosomia, hypoglycemia, and hyperbilirubinemia. *Obstet Gynecol*. 2006 Nov;108(5):1153-61.
376. Jain NJ, Denk CE, Kruse LK, Dandolu V. Maternal obesity: can pregnancy weight gain modify risk of selected adverse pregnancy outcomes? *Am J Perinatol*. 2007 May;24(5):291-8.
377. Smith SA, Hulsey T, Goodnight W. Effects of obesity on pregnancy. *J Obstet Gynecol Neonatal Nurs*. 2008 Mar-Apr;37(2):176-84.
378. Guideline Development Group. Management of diabetes from preconception to the postnatal period: summary of NICE guidance. *BMJ*. 2008 Mar 29;336(7646):714-7.
379. Reece EA, Homko CJ. Prepregnancy care and the prevention of fetal malformations in the pregnancy complicated by diabetes. *Clin Obstet Gynecol*. 2007 Dec;50(4):990-7.
380. Kmietowicz Z. Screen all pregnant women for risk factors for gestational diabetes, says NICE. *BMJ*. 2008 Mar 29;336(7646):689.
381. Tieu J, Crowther CA, Middleton P. Dietary advice in pregnancy for preventing gestational diabetes mellitus. *Cochrane Database Syst Rev*. 2008 Apr 16;(2):CD006674.
382. Balsells M, García-Patterson A, Gich I, Corcay R. Maternal and fetal outcome in women with type 2 versus type 1 diabetes mellitus: a systematic review and metaanalysis. *J Clin Endocrinol Metab*. 2009 Nov;94(11):4284-91.
383. Landon MB, Spong CY, Thom E, et al. A multicenter, randomized trial of treatment for mild gestational diabetes. *N Engl J Med*. 2009 Oct 1;361(14):1339-48.
384. Perichart-Perera O, Balas-Nakash M, Parra-Covarrubias A, et al. A medical nutrition therapy program improves perinatal outcomes in Mexican pregnant women with gestational diabetes and type 2 diabetes mellitus. *Diabetes Educ*. 2009 Nov-Dec;35(6):1004-13.
385. Alwan N, Tuffnell DJ, West J. Treatments for gestational diabetes. *Cochrane Database Syst Rev*. 2009 Jul 8;(3):CD003395.
386. Vasudevan C, Renfrew M, McGuire W. Fetal and perinatal consequences of maternal obesity. *Arch Dis Child Fetal Neonatal Ed*. 2010 Jun 7.
387. Agarwal MM, Dhatt GS, Shah SM. Gestational diabetes: simplifying the IADPSG diagnostic algorithm using fasting plasma glucose. *Diabetes Care*. 2010 Jun 2.
- 17. Detecting and treating diabetes**
370. Bewley C. Diabetes in pregnancy. *Nurs Stand*. 2002 Mar 6-12;16(25):47-52.

388. Betti M, Ceccatelli G, Belcari F, et al. Neonatal outcome in newborns from mothers with endocrinopathies. *Gynecol Endocrinol.* 2010 Jun 2.
389. Petry CJ. Gestational diabetes: risk factors and recent advances in its genetics and treatment. *Br J Nutr.* 2010 May;21:1-13.
- 18. Deparasitization in areas of high prevalence using albendazole**
390. de Silva NR, Kodituwakku KK, Edirisinghe SS, de Silva HJ. Routine use of mebendazole in pregnancy. *Ceylon Med J.* 1996 Sep;41(3):99-101.
391. de Silva NR, Sirisena JL, Gunasekera DP, et al. Effect of mebendazole therapy during pregnancy on birth outcome. *Lancet.* 1999 Apr 3;353(9159):1145-9.
392. Diav-Citrin O, Shechtman S, Arnon J, et al. Pregnancy outcome after gestational exposure to mebendazole: a prospective controlled cohort study. *Am J Obstet Gynecol.* 2003 Jan;188(1):282-5.
393. Acs N, Bánhyi F, Puhó E, Czeizel AE. Population-based case-control study of mebendazole in pregnant women for birth outcomes. *Congenit Anom (Kyoto).* 2005 Sep;45(3):85-8.
394. Larocque R, Casapia M, Gotuzzo E, et al. A double-blind randomized controlled trial of antenatal mebendazole to reduce low birthweight in a hookworm-endemic area of Peru. *Trop Med Int Health.* 2006 Oct;11(10):1485-95.
395. Christian P, Shahid F, Rizvi A, et al. Treatment response to standard of care for severe anemia in pregnant women and effect of multivitamins and enhanced anthelmintics. *Am J Clin Nutr.* 2009 Mar;89(3):853-61.
- 19. Detecting and treating group B streptococcus**
396. Schrag S, Gorwitz R, Fultz-Butts K, Schuchat A. Prevention of perinatal group B streptococcal disease. Revised guidelines from CDC. *MMWR Recomm Rep.* 2002 Aug 16;51(RR-11):1-22.
397. Puopolo KM, Madoff LC, Eichenwald EC. Early-onset group B streptococcal disease in the era of maternal screening. *Pediatrics.* 2005 May;115(5):1240-6.
398. Illuzzi JL, Bracken MB. Duration of intrapartum prophylaxis for neonatal group B streptococcal disease: a systematic review. *Obstet Gynecol.* 2006 Nov;108(5):1254-65.
399. Mechurová A, Unzeitig V, Vlk R. Guidelines for the diagnosis and treatment of group B streptococcal colonization during pregnancy and delivery. *Klin Mikrobiol Infekc Lek.* 2006 Apr;12(2):76-7.
400. Renner RM, Renner A, Schmid S, et al. Efficacy of a strategy to prevent neonatal early-onset group B streptococcal (GBS) sepsis. *J Perinat Med.* 2006;34(1):32-8.
401. Heath PT, Schuchat A. Perinatal group B streptococcal disease. *Best Pract Res Clin Obstet Gynaecol.* 2007 Jun;21(3):411-24.
402. Hamada S, Vearncombe M, McGeer A, Shah PS. Neonatal group B streptococcal disease: incidence, presentation, and mortality. *J Matern Fetal Neonatal Med.* 2008 Jan;21(1):53-7.
403. Edwards RK, Novak-Weekley SM, Koty PP, et al. Rapid group B streptococci screening using a real-time polymerase chain reaction assay. *Obstet Gynecol.* 2008 Jun;111(6):1335-41.
404. Nandyal RR. Update on group B streptococcal infections: perinatal and neonatal periods. *J Perinat Neonatal Nurs.* 2008 Jul-Sep;22(3):230-7.
405. Centelles-Serrano MJ, Pérez-Moreno MO, Llovet-Lombarte, et al. Effectiveness of systematic investigation for Group B Streptococcus in urine samples to identify colonized pregnant women. *Enferm Infect Microbiol Clin.* 2009 Aug-Sep;27(7):394.
406. Centers for Disease Control and Prevention (CDC). Trends in perinatal group B streptococcal disease - United States, 2000-2006. *MMWR Morb Mortal Wkly Rep.* 2009 Feb 13;58(5):109-12.
407. Goins WP, Talbot TR, Schaffner W, et al. Adherence to perinatal group B streptococcal prevention guidelines. *Obstet Gynecol.* 2010 Jun;115(6):1217-24.
408. Valkenburg-van den Berg AW, Houtman-Roelofsen RL, Oostvogel PM, et al. Timing of group B streptococcus screening in pregnancy: a systematic review. *Gynecol Obstet Invest.* 2010;69(3):174-83.
- 20. Nifedipine during preterm labor**
409. Laohapojanart N, Soorapan S, Wacharaprechanont T, Ratanajamit C. Safety and efficacy of oral nifedipine versus terbutaline injection in preterm labor. *J Med Assoc Thai.* 2007 Nov;90(11):2461-9.
410. Lyell DJ, Pullen K, Campbell L, Ching S, et al. Magnesium sulfate compared with nifedipine for acute tocolysis of preterm labor: a randomized controlled trial. *Obstet Gynecol.* 2007 Jul;110(1):61-7.
411. Papatsonis DN, Bos JM, van Geijn HP, et al. Nifedipine pharmacokinetics and plasma levels in the management of preterm labor. *Am J Ther.* 2007 Jul-Aug;14(4):346-50.
412. Mawaldi L, Duminy P, Tamim H. Terbutaline versus nifedipine for prolongation of pregnancy in patients with preterm labor. *Int J Gynaecol Obstet.* 2008 Jan;100(1):65-8.
413. Silberschmidt AL, Kühn-Velten WN, Juon AM, et al. Nifedipine concentration in maternal and umbilical cord blood after nifedipine gastrointestinal therapeutic system for tocolysis. *BJOG.* 2008 Mar;115(4):480-5.
414. Kam KY, Lamont RF. Developments in the pharmacotherapeutic management of spontaneous preterm labor. *Expert Opin Pharmacother.* 2008 May;9(7):1153-68.
415. Nassar AH, Abu-Musa AA, Awwad J, et al. Two dose regimens of nifedipine for management of preterm labor: a randomized controlled trial. *Am J Perinatol.* 2009 Sep;26(8):575-81.
416. King JF, Flenady VJ, Papatsonis DNM, Dekker GA, Carbonne B. Calcium channel blockers for inhibiting preterm labour. *Cochrane Library* 2010.
417. Hass D. Calcium channel blockers. Preterm Birth. *BMJ Clinical Evidence* 2010.

## 21. Prenatal corticosteroids (betamethasone, dexamethasone) to foster lung maturation

418. Liggins GC. A controlled trial of antepartum glucocorticoid treatment for the prevention of respiratory distress syndrome in premature infants. *Pediatrics* 1972;50:515-525.
419. Sinclair JC. Meta-analysis of randomized controlled trials of antenatal corticosteroids for the prevention of respiratory distress syndrome: discussion. *Am J Obstet Gynecol* 1995;173:335-344.
420. Crowley P. Prophylactic corticosteroids for preterm birth. *Cochrane Database Syst Rev*. 2000;(2):CD000065.
421. Abbasi S, Hirsch D, Davis J, Tolosa J, et al. Effect of single versus multiple courses of antenatal corticosteroids on maternal and neonatal outcome. *Am J Obstet Gynecol*. 2000 May;182(5):1243-9.
422. Bar-Lev MR, Maayan-Metzger A, Matok I, et al. Short-term outcomes in low birth weight infants following antenatal exposure to betamethasone versus dexamethasone. *Obstet Gynecol*. 2004 Sep;104(3):484-8.
423. Peaceman AM, Bajaj K, Kumar P, Grobman WA. The interval between a single course of antenatal steroids and delivery and its association with neonatal outcomes. *Am J Obstet Gynecol*. 2005 Sep;193(3 Pt 2):1165-9.
424. Smrcek JM, Schwartau N, Kohl M, Berg C, et al. Antenatal corticosteroid therapy in premature infants. *Arch Gynecol Obstet*. 2005 Jan;271(1):26-32.
425. Roberts D, Dalziel S. Antenatal corticosteroids for accelerating fetal lung maturation for women at risk of preterm birth. *Cochrane Database Syst Rev*. 2006 Jul 19;3:CD004454.
426. Lee BH, Stoll BJ, McDonald SA, et al. Adverse neonatal outcomes associated with antenatal dexamethasone versus antenatal betamethasone. *Pediatrics*. 2006 May;117(5):1503-10.
427. Crowther CA, Harding JE. Repeat doses of prenatal corticosteroids for women at risk of preterm birth for preventing neonatal respiratory disease. *Cochrane Database Syst Rev*. 2007 Jul 18;(3):CD003935.
428. Feldman DM, Carbone J, Belden L, et al. Betamethasone vs dexamethasone for the prevention of morbidity in very-low-birthweight neonates. *Am J Obstet Gynecol*. 2007 Sep;197(3):284.e1-4.
429. Miracle X, Di Renzo GC, Stark A, et al. Guideline for the use of antenatal corticosteroids for fetal maturation. *J Perinat Med*. 2008;36(3):191-6.
430. American College of Obstetricians and Gynecologists Committee on Obstetric Practice. ACOG Committee Opinion No. 402: Antenatal corticosteroid therapy for fetal maturation. *Obstet Gynecol*. 2008 Mar;111(3):805-7.
431. Murphy KE, Hannan ME, Willan AR, et al. Multiple courses of antenatal corticosteroids for preterm birth (MACS): a randomised controlled trial. *Lancet*. 2008 Dec 20;372(9656):2143-51.

432. Brownfoot FC, Crowther CA, Middleton P. Different corticosteroids and regimens for accelerating fetal lung maturation for women at risk of preterm birth. *Cochrane Database Syst Rev*. 2008 Oct 8;(4):CD006764.
433. Mazumder P, Dutta S, Kaur J, Narang A. Single versus multiple courses of antenatal betamethasone and neonatal outcome: a randomized controlled trial. *Indian Pediatr*. 2008 Aug;45(8):661-7.
434. Finken MJ, Keijzer-Veen MG, et al. Antenatal glucocorticoid treatment is not associated with long-term metabolic risks in individuals born before 32 weeks of gestation. *Arch Dis Child Fetal Neonatal Ed*. 2008 Nov;93(6):F442-7.
435. Hayes EJ, Paul DA, Stahl GE, et al. Effect of antenatal corticosteroids on survival for neonates born at 23 weeks of gestation. *Obstet Gynecol*. 2008 Apr;111(4):921-6.
436. Garite TJ, Kurtzman J, Maurel K, et al. Impact of a 'rescue course' of antenatal corticosteroids: a multicenter randomized placebo-controlled trial. *Am J Obstet Gynecol*. 2009 Mar;200(3):248.e1-9.
437. Sotiriadis A, Makrydimas G, Papatheodorou S, Ioannidis JP. Corticosteroids for preventing neonatal respiratory morbidity after elective caesarean section at term. *Cochrane Database Syst Rev*. 2009 Oct 7;(4):CD006614.
438. Halliday HL, Ehrenkranz RA, Doyle LW. Late (>7 days) postnatal corticosteroids for chronic lung disease in preterm infants. *Cochrane Database Syst Rev*. 2009 Jan 21;(1):CD001145.
439. Halliday HL, Ehrenkranz RA, Doyle LW. Early (< 8 days) postnatal corticosteroids for preventing chronic lung disease in preterm infants. *Cochrane Database Syst Rev*. 2010 Jan 20;(1):CD001146.
440. Mwansa-Kambafwile J, Cousens S, Hansen T, Joy E, Lawn JE. Antenatal steroids in preterm labour for the prevention of neonatal deaths due to complications of preterm birth. *International Journal of Epidemiology* 2010;39:i122-i133.
441. Roberts D, Dalziel S. Antenatal corticosteroids for accelerating fetal lung maturation for women at risk of preterm birth. *Cochrane Library* 2010. *BMJ Clinical Evidence* 2010

## 22. Detecting, preventing, and treating Rh isoimmunization

442. Bowman JM, Chown B, Lewis M, Pollock JM. Rh isoimmunization during pregnancy: antenatal prophylaxis. *Can Med Assoc J*. 1978 Mar 18;118(6):623-7.
443. Urbaniak SJ. The scientific basis of antenatal prophylaxis. *Br J Obstet Gynaecol*. 1998 Nov;105 Suppl 18:11-8.
444. Crowther CA, Keirse MJ. Anti-D administration in pregnancy for preventing rhesus alloimmunisation. *Cochrane Database Syst Rev*. 2000;(2):CD000020.
445. Crowther C, Middleton P. Anti-D administration after childbirth for preventing Rhesus alloimmunisation. *Cochrane Database Syst Rev*. 2000;(2):CD000021.
446. Fung Kee Fung K, Eason E, et al. Prevention of Rh alloimmunization. *J Obstet Gynaecol Can*. 2003 Sep;25(9):765-73.

447. Abdel-Fattah SA, Shefras J, et al. Reassuring fetal middle cerebral artery doppler velocimetry in alloimmunised pregnancies: neonatal outcomes without invasive procedures. *Fetal Diagn Ther.* 2005 Sep-Oct;20(5):341-5
448. Cortey A, Brossard Y. Prevention of fetomaternal rhesus-D allo-immunization. Practical aspects. *J Gynecol Obstet Biol Reprod (Paris).* 2006 Feb;35(1 Suppl):1S123-1S130
449. American College of Obstetricians and Gynecologists. ACOG Practice Bulletin No. 75: management of alloimmunization. *Obstet Gynecol.* 2006 Aug;108(2):457-64
450. Oepkes D, Seaward PG, Vandebussche FP, et al. Doppler ultrasonography versus amniocentesis to predict fetal anemia. *N Engl J Med.* 2006 Jul 13;355(2):156-64
451. Collège National des Gynécologues et Obstétriciens Français. Recommendations for clinical practice. Prevention in maternofetal Rh immunization (December 2005). *Gynecol Obstet Fertil.* 2006 Apr;34(4):360-5
452. Chaffe B, Ford J, Bills V. Routine antenatal anti-D prophylaxis and patient compliance with the two-dose regimen. *Transfus Med.* 2007 Oct;17(5):399-403
453. Koelewijn JM, de Haas M, Vrijkotte TG, et al. Risk factors for RhD immunisation despite antenatal and postnatal anti-D prophylaxis. *BJOG.* 2009 Sep;116(10):1307-14.
454. Pretlove SJ, Fox CE, Khan KS, Kilby MD. Noninvasive methods of detecting fetal anaemia: a systematic review and meta-analysis. *BJOG.* 2009 Nov;116(12):1558-67
455. Pilgrim H, Lloyd-Jones M, Rees A. Routine antenatal anti-D prophylaxis for RhD-negative women: a systematic review and economic evaluation. *Health Technol Assess.* 2009 Feb;13(10):iii, ix-xi, 1-103
456. Boulet S, Krause C, Tixier H, Bardou M, Sagot P. Relevance of new recommendations on routine antenatal prevention of rhesus immunization: an appraisal based on a retrospective analysis of all cases observed in two French administrative areas of 3 million inhabitants. *Eur J Obstet Gynecol Reprod Biol.* 2009 Sep;146(1):65-70.
461. Siqueira FM, Cota LO, Costa JE, et al. Intrauterine growth restriction, low birth weight, and preterm birth: adverse pregnancy outcomes and their association with maternal periodontitis. *J Periodontol.* 2007 Dec;78(12):2266-76.
462. Ferguson JE 2nd, Hansen WF, Novak KF, Novak MJ. Should we treat periodontal disease during gestation to improve pregnancy outcomes? *Clin Obstet Gynecol.* 2007 Jun;50(2):454-67.
463. Ruma M, Boggess K, Moss K, et al. Enfermedad periodontal materna, inflamación sistémica y riesgo de pre-eclampsia. *Am J Obstet Gynecol.* 2008 Apr;198(4):389.e1-5.
464. Boggess KA. Maternal oral health in pregnancy. *Obstet Gynecol.* 2008 Apr;111(4):976-86.
465. Polyzos NP, Polyzos IP, Mauri D, et al. Effect of periodontal disease treatment during pregnancy on preterm birth incidence: a metaanalysis of randomized trials. *Am J Obstet Gynecol.* 2009 Mar;200(3):225-32.
466. Novák T, Radnai M, Gorzó I, et al. Prevention of preterm delivery with periodontal treatment. *Fetal Diagn Ther.* 2009;25(2):230-3.
467. Radnai M, Pál A, Novák T, et al. Benefits of periodontal therapy when preterm birth threatens. *J Dent Res.* 2009 Mar;88(3):280-4.
468. Reis DM, Pitta DR, Ferreira HM, et al. Health education as a strategy for the promotion of oral health in the pregnancy period. *Cien Saude Colet.* 2010 Jan;15(1):269-76.

## 24. Detecting, preventing, and managing domestic violence and abuse, and depression

### 23. Preventing and treating periodontal disease

457. Offenbacher S, Lieff S, Boggess KA, et al. Maternal periodontitis and prematurity. Part I: Obstetric outcome of prematurity and growth restriction. *Ann Periodontol.* 2001 Dec;6(1):164-74
458. Jeffcoat MK, Geurs NC, Reddy MS, et al. Periodontal infection and preterm birth: results of a prospective study. *J Am Dent Assoc.* 2001 Jul;132(7):875-80
459. López NJ, Smith PC, Gutierrez J. Periodontal therapy may reduce the risk of preterm low birth weight in women with periodontal disease: a randomized controlled trial. *J Periodontol.* 2002 Aug;73(8):911-24
460. Michalowicz BS, Hodges JS, DiAngelis AJ, et al. Treatment of periodontal disease and the risk of preterm birth. *N Engl J Med.* 2006 Nov 2;355(18):1885-94.
469. Espinoza H, Camacho AV. Muerte materna ocasionada por violencia doméstica: componente crítico e inadvertido de la mortalidad materna. *Rev Panam Salud Pública* v.17 n.2 Washington feb. 2005.
470. Monteiro Cde F, Costa NS, Nascimento PS, de Aguiar YA. Intrafamiliar violence against pregnant adolescents. *Rev Bras Enferm.* 2007 Jul-Aug;60(4):373.
471. Shoffner DH. We don't like to think about it: intimate partner violence during pregnancy and postpartum. *J Perinat Neonatal Nurs.* 2008 Jan-Mar;22(1):39-48.
472. Rodriguez MA, Heilemann MV, Fielder E, et al. Intimate partner violence, depression, and PTSD among pregnant Latina women. *Ann Fam Med.* 2008 Jan-Feb;6(1):44-52.
473. Tiwari A, Chan KL, Fong D, et al. The impact of psychological abuse by an intimate partner on the mental health of pregnant women. *BJOG.* 2008 Feb;115(3):377-84.
474. Rodrigues T, Rocha L, Barros H. Physical abuse during pregnancy and preterm delivery. *Am J Obstet Gynecol.* 2008 Feb;198(2):171.e1-6.
475. Sanchez SE, Qiu C, Perales MT, et al. Intimate partner violence (IPV) and preeclampsia among Peruvian women. *Eur J Obstet Gynecol Reprod Biol.* 2008 Mar;137(1):50-5.
476. Sela HY, Shveiky D, Laufer N, et al. Pregnant women injured in terror-related multiple casualty incidents: injuries and outcomes. *J Trauma.* 2008 Mar;64(3):727-32.

477. Taft AJ, Small R, Hegarty KL, et al. MOSAIC (MOthers' Advocates In the Community): protocol and sample description of a cluster randomised trial of mentor mother support to reduce intimate partner violence among pregnant or recent mothers. *BMC Public Health*. 2009 May 27;9:159.
478. Ramsay J, Carter Y, Davidson L, et al. Advocacy interventions to reduce or eliminate violence and promote the physical and psychosocial well-being of women who experience intimate partner abuse. *Cochrane Database Syst Rev*. 2009 Jul 8;(3):CD005043.
479. Kiely M, El-Mohandes AA, El-Khorazaty MN, Gantz MG. An integrated intervention to reduce intimate partner violence in pregnancy: a randomized controlled trial. *Obstet Gynecol*. 2010 Feb;115(2 Pt 1):273-83.
490. Leonardi-Bee J, Smyth A, Britton J, Coleman T, et al. Environmental tobacco smoke and fetal health: systematic review and meta-analysis. *Arch Dis Child Fetal Neonatal Ed*. 2008 Sep;93(5):F351-61.
491. Ruger JP, Weinstein MC, Hammond SK, et al. Cost-effectiveness of motivational interviewing for smoking cessation and relapse prevention among low-income pregnant women: a randomized controlled trial. *Value Health*. 2008 Mar-Apr;11(2):191-8.
492. Tsui HC, Wu HD, Lin CJ, et al. Prenatal smoking exposure and neonatal DNA damage in relation to birth outcomes. *Pediatr Res*. 2008 Aug;64(2):131-4.
493. Lumley J, Chamberlain C, Dowswell T, et al. Interventions for promoting smoking cessation during pregnancy. *Cochrane Database Syst Rev*. 2009 Jul 8;(3):CD001055.

## 25. Detecting and preventing exposure to tobacco

480. Ferraz EM, Gray RH, Cunha TM, et al. Determinants of preterm delivery and intrauterine growth retardation in north-east Brazil. *Int J Epidemiol*. 1990 Mar;19(1):101-8.
481. Vega J, Sáez G, Smith M, Agurto M, Morris NM, et al. Risk factors for low birth weight and intrauterine growth retardation in Santiago, Chile. *Rev Med Chil*. 1993 Oct;121(10):1210-9.
482. Roquer JM, Figueras J, Botet F, Jiménez R. Influence on fetal growth of exposure to tobacco smoke during pregnancy. *Acta Paediatr*. 1995 Feb;84(2):118-21.
483. Horta BL, Victora CG, Menezes AM, et al. Low birthweight, preterm births and intrauterine growth retardation in relation to maternal smoking. *Paediatr Perinat Epidemiol*. 1997 Apr;11(2):140-51.
484. Windham GC, Hopkins B, Fenster L, Swan SH. Prenatal active or passive tobacco smoke exposure and the risk of preterm delivery or low birth weight. *Epidemiology*. 2000 Jul;11(4):427-33.
485. Dejmek J, Solanský I, Podrazilová K, Srám RJ. The exposure of nonsmoking and smoking mothers to environmental tobacco smoke during different gestational phases and fetal growth. *Environ Health Perspect*. 2002 Jun;110(6):601-6.
486. Chełchowska M, Laskowska-Klita T, Leibschang J. The effect of tobacco smoking during pregnancy on concentration of malondialdehyde in blood of mothers and in umbilical cord blood. *Ginekol Pol*. 2005 Dec;76(12):960-5.
487. Szymanowski K, Chmaj-Wierzchowska K, Florek E, Opala T. Influence of tobacco smoking to development of the fetus, newborn and child--a review. *Przegl Lek*. 2006;63(10):1135-7.
488. Ward C, Lewis S, Coleman T. Prevalence of maternal smoking and environmental tobacco smoke exposure during pregnancy and impact on birth weight: retrospective study using Millennium Cohort. *BMC Public Health*. 2007 May 16;7:81.
489. de Chazeron I, Llorca PM, Ughetto S, et al. Occult maternal exposure to environmental tobacco smoke exposure. *Tob Control*. 2007 Feb;16(1):64-5.

## 26. Detecting and preventing exposure to alcohol

494. Weber MK, Floyd RL, Riley EP, et al. National Task Force on Fetal Alcohol Syndrome and Fetal Alcohol Effect: defining the national agenda for fetal alcohol syndrome and other prenatal alcohol-related effects. *MMWR Recomm Rep*. 2002 Sep 20;51(RR-14):9-12.
495. Floyd RL, O'Connor MJ, Sokol RJ, et al. Recognition and prevention of fetal alcohol syndrome. *Obstet Gynecol*. 2005 Nov;106(5 Pt 1):1059-64.
496. Bertrand J, Floyd LL, Weber MK; Fetal Alcohol Syndrome Prevention Team, Division of Birth Defects and Developmental Disabilities, National Center on Birth Defects and Developmental Disabilities, Centers for Disease Control and Prevention (CDC). Guidelines for identifying and referring persons with fetal alcohol syndrome. *MMWR Recomm Rep*. 2005 Oct 28;54(RR-11):1-14.
497. Krulewitch CJ. Alcohol consumption during pregnancy. *Annu Rev Nurs Res*. 2005;23:101-34.
498. Ripabelli G, Cimmino L, Grasso GM. Alcohol consumption, pregnancy and fetal alcohol syndrome: implications in public health and preventive strategies. *Ann Ig*. 2006 Sep-Oct;18(5):391-406.
499. Floyd RL, O'Connor MJ, Bertrand J, Sokol R. Reducing adverse outcomes from prenatal alcohol exposure: a clinical plan of action. *Alcohol Clin Exp Res*. 2006 Aug;30(8):1271-5.
500. Floyd RL, Sobell M, Velasquez MM, et al. Preventing alcohol-exposed pregnancies: a randomized controlled trial. *Am J Prev Med*. 2007 Jan;32(1):1-10.
501. Floyd RL, Weber MK, Denny C, O'Connor MJ. Prevention of fetal alcohol spectrum disorders. *Dev Disabil Res Rev*. 2009;15(3):193-9.
502. Floyd RL, Weber MK, Denny C, O'Connor MJ. Prevention of fetal alcohol spectrum disorders. *Dev Disabil Res Rev*. 2009;15(3):193-9.
503. Coles CD, Kable JA, Taddeo E. Math performance and behavior problems in children affected by prenatal alcohol exposure: intervention and follow-up. *J Dev Behav Pediatr*. 2009 Feb;30(1):7-15.

504. Stade BC, Bailey C, Dzendoletas D, et al. Psychological and/or educational interventions for reducing alcohol consumption in pregnant women and women planning pregnancy. *Cochrane Database Syst Rev.* 2009 Apr 15;(2):CD004228.
505. Hutson JR, Magri R, Gareri JN, Koren G. The incidence of prenatal alcohol exposure in Montevideo Uruguay as determined by meconium analysis. *Ter Drug Monit.* 2010 Jun;32(3):311-7.
519. Bagner DM, Sheinkopf SJ, Miller-Loncar C, et al. The effect of parenting stress on child behavior problems in high-risk children with prenatal drug exposure. *Child Psychiatry Hum Dev.* 2009 Mar;40(1):73-84..
520. Marin SJ, Keith L, Merrell M, McMillin GA. Comparison of drugs of abuse detection in meconium by EMIT II and ELISA. *J Anal Toxicol.* 2009 Apr;33(3):148-54.
521. Koren G, Nulman I. Drugs in pregnancy: associations, causation, and misperceptions. *Arch Pediatr Adolesc Med.* 2010 May;164(5):494-5.

## 27. Detecting and preventing exposure to drugs

506. Chiriboga CA. Neurological correlates of fetal cocaine exposure. *Ann NY Acad Sci.* 1998.
507. Bauer CR, Shankaran S, Bada HS, et al. The Maternal Lifestyle Study: drug exposure during pregnancy and short-term maternal outcomes. *Am J Obstet Gynecol.* 2002 Mar;186(3):487-95.
508. Singer LT, Arendt R, Minnes S, et al. Cognitive and motor outcomes of cocaine-exposed infants. *JAMA.* 2002 Apr 17;287(15):1952-60.
509. Singer LT, Minnes S, Short E, et al. Cognitive outcomes of preschool children with prenatal cocaine exposure. *JAMA.* 2004 May 26;291(20):2448-56.
510. Bauer CR, Langer JC, Shankaran S, et al. Acute neonatal effects of cocaine exposure during pregnancy. *Arch Pediatr Adolesc Med.* 2005 Sep;159(9):824-34.
511. Montgomery D, Plate C, Alder SC, et al. Testing for fetal exposure to illicit drugs using umbilical cord tissue vs meconium. *J Perinatol.* 2006 Jan 1;26(1):11-4.
512. Linares TJ, Singer LT, Kirchner HL, et al. Mental health outcomes of cocaine-exposed children at 6 years of age. *J Pediatr Psychol.* 2006 Jan-Feb;31(1):85-97.
513. Bada HS, Das A, Bauer CR, et al. Impact of prenatal cocaine exposure on child behavior problems through school age. *Pediatrics.* 2007 Feb;119(2):e348-59.
514. Montgomery DP, Plate CA, Jones M, et al. Using umbilical cord tissue to detect fetal exposure to illicit drugs: a multicentered study in Utah and New Jersey. *J Perinatol.* 2008 Nov;28(11):750-3.
515. Koren G, Hutson J, Gareri J. Novel methods for the detection of drug and alcohol exposure during pregnancy: implications for maternal and child health. *Clin Pharmacol Ther.* 2008 Apr;83(4):631-4.
516. Singer LT, Nelson S, Short E, et al. Prenatal cocaine exposure: drug and environmental effects at 9 years. *J Pediatr.* 2008 Jul;153(1):105-11.
517. Bennett DS, Bendersky M, Lewis M. Children's cognitive ability from 4 to 9 years old as a function of prenatal cocaine exposure, environmental risk, and maternal verbal intelligence. *Dev Psychol.* 2008 Jul;44(4):919-28.
518. Draper ES, Rankin J, Tonks AM, et al. Recreational drug use: a major risk factor for gastroschisis? *Am J Epidemiol.* 2008 Feb 15;167(4):485-91.

## III. Interventions during delivery

### 1. Clean and safe delivery practices (with skilled personnel)

522. Camacho V, de la Gálvez A, Murdock M, et al. Atención calificada del parto en América Latina y el Caribe: consulta técnica regional. *USAID/OPS,* 2004.
523. Sibley L, Ann Sipe T. What can a meta-analysis tell us about traditional birth attendant training and pregnancy outcomes? *Midwifery.* 2004 Mar;20(1):51-60.
524. Say L, Raine R. A systematic review of inequalities in the use of maternal health care in developing countries: examining the scale of the problem and the importance of context. *Bull World Health Organ.* 2007 Oct;85(10):812-9.
525. Harvey SA, Blandón YC, McCaw-Binns A, et al. Are skilled birth attendants really skilled? A measurement method, some disturbing results and a potential way forward. *Bull World Health Organ.* 2007 Oct;85(10):783-90.

### 2. Accompanying the mother during labor and delivery

526. Klaus MH, Kennell JH. The doula: an essential ingredient of childbirth rediscovered. *Acta Paediatr.* 1997 Oct;86(10):1034-6.
527. Madi BC, Sandall J, Bennett R, MacLeod C. Effects of female relative support in labor: a randomized controlled trial. *Birth.* 1999 Mar;26(1):4-8.
528. Campbell DA, Lake MF, Falk M, Backstrand JR. A randomized control trial of continuous support in labor by a lay doula. *J Obstet Gynecol Neonatal Nurs.* 2006 Jul-Aug;35(4):456-64.
529. Campbell D, Scott KD, Klaus MH, Falk M. Female relatives or friends trained as labor doulas: outcomes at 6 to 8 weeks postpartum. *Birth.* 2007 Sep;34(3):220-7.
530. Scott KD, Klaus PH, Klaus MH. The obstetrical and postpartum benefits of continuous support during childbirth. *J Womens Health Gend Based Med.* 1999 Dec;8(10):1257-64.
531. Hodnett ED. WITHDRAWN: Caregiver support for women during childbirth. *Cochrane Database Syst Rev.* 2007 Jul 18;(1):CD000199.
532. Hodnett ED, Gates S, Hofmeyr GJ, Sakala C. Continuous support for women during childbirth. *Cochrane Database Syst Rev.* 2007 Jul 18;(3):CD003766.

### 3. Erythromycin in PROM before delivery in first 37 weeks of gestation

533. Kenyon SL, Taylor DJ, Tarnow-Mordi W; ORACLE Collaborative Group. Broad-spectrum antibiotics for preterm, prelabour rupture of fetal membranes: the ORACLE I randomised trial. ORACLE Collaborative Group. *Lancet*. 2001 Mar 31;357(9261):979-88.
534. Buitendijk SE. Erythromycin for premature rupture of membranes is beneficial for infant. *Ned Tijdschr Geneeskd*. 2001 Sep 22;145(38):1828-31.
535. Kenyon S, Taylor DJ, Tarnow-Mordi WO; ORACLE Collaborative Group. ORACLE--antibiotics for preterm prelabour rupture of the membranes: short-term and long-term outcomes. *Acta Paediatr Suppl*. 2002;91(437):12-5.
536. Kenyon S, Boulvain M, Neilson J. Antibiotics for preterm rupture of membranes. *Cochrane Database Syst Rev*. 2003;(2):CD001058.
537. Kenyon S, Boulvain M, Neilson J. Antibiotics for preterm rupture of the membranes: a systematic review. *Obstet Gynecol*. 2004 Nov;104(5 Pt 1):1051-7.
538. Ehsanipoor RM, Chung JH, Clock CA, et al. A retrospective review of ampicillin-sulbactam and amoxicillin + clavulanate vs cefazolin/cephalexin and erythromycin in the setting of preterm premature rupture of membranes: maternal and neonatal outcomes. *Am J Obstet Gynecol*. 2008 May;198(5):e54-6.
539. Yudin MH, van Schalkwyk J, Van Eyk N, et al. Antibiotic therapy in preterm premature rupture of the membranes. *J Obstet Gynaecol Can*. 2009 Sep;31(9):863-7, 868-74.

### 4. Monitoring labor with partogram

540. Villacaqui R, Vidal Mosquera MO, Penaranda A, Herrera I. Partograma con curvas de alerta en pacientes de la altura. *Ginecología y Obstetricia Abril* 1998 Vol. 44 Nº 1.
541. Sanches, IH. Comparação Entre Dois Modelos de Partogramas Aplicados à Assistência Clínica ao Parto de Primigestas. *Rev. Bras. Ginecol. Obstet.* 2000, vol. 22, no. 4:245-245.
542. Nápoles D, Bajuelo AE, Téllez MS, Couto D. El partograma y las desviaciones del trabajo de parto. *Medisan* 2004; 8(4): 64-72.
543. Nápoles D. Alternativas en las desviaciones del trabajo de parto. *Medisan* 2005; 9(2).
544. Fatusi AO, Makinde ON, Adeyemi AB, et al. Evaluation of health workers' training in use of the partogram. *Int J Gynaecol Obstet*. 2008 Jan;100(1):41-4.
545. Organización Panamericana de la Salud. Instructivo para el Diligenciamiento de la Historia Clínica del Parto "Partograma" del CLAP - OPS/OMS, febrero, 2008.
546. Lavender T, Hart A, Smyth RM, Effect of partogram use on outcomes for women in spontaneous labour at term. *Cochrane Database Syst Rev*. 2008 Oct 8;(4):CD005461.

547. Rocha IM, de Oliveira SM, Schneck CA, et al. The partogram as an instrument to analyze care during labor and delivery. *Rev Esc Enferm USP*. 2009 Dec;43(4):880-8.
548. Basu JK, Buchmann EJ, Basu D. Role of a second stage partogram in predicting the outcome of normal labour. *Aust N Z J Obstet Gynaecol*. 2009 Apr;49(2):158-61.

### 5. Caesarean section in breech presentation with LBW infants

549. Lee KS, Khoshnood B, Sriram S, Hsieh HL, Singh J, Mittendorf R. Relationship of cesarean delivery to lower birth weight-specific neonatal mortality in singleton breech infants in the United States. *Obstet Gynecol*. 1998 Nov;92(5):769-74.
550. Makris N, Xygakis A, Chionis A, Sakellaropoulos G, Michalas S. The management of breech presentation in the last three decades. *Clin Exp Obstet Gynecol*. 1999;26(3-4):178-80.
551. Muhuri PK, Macdorman MF, Menacker F. Method of delivery and neonatal mortality among very low birth weight infants in the United States. *Matern Child Health J*. 2006 Jan;10(1):47-53.
552. Robilio PA, Boe NM, Danielsen B, Gilbert WM. Vaginal vs. cesarean delivery for preterm breech presentation of singleton infants in California: a population-based study. *J Reprod Med*. 2007 Jun;52(6):473-9.
553. Lee HC, Gould J. Survival advantage associated with cesarean delivery in very low birth weight vertex neonates. *Obstet Gynecol*. 2007 May;109(5):1203.
554. Malloy MH, Doshi S. Cesarean section and the outcome of very preterm and very low-birthweight infants. *Clin Perinatol*. 2008 Jun;35(2):421-35, viii.

### 6. Antibiotic prophylaxis in caesarean section

555. Martins AC, Krauss-Silva L. Systematic reviews of antibiotic prophylaxis in cesareans. *Cad Saude Publica*. 2006 Dec;22(12):2513-26.
556. Rudge MV, Atallah AN, Peraçoli JC, et al. Randomized controlled trial on prevention of postcesarean infection using penicillin and cephalothin in Brazil. *Acta Obstet Gynecol Scand*. 2006;85(8):945-8.
557. Alekwe LO, Kuti O, Orji EO, Ogunniyi SO. Comparison of ceftriaxone versus triple drug regimen in the prevention of cesarean section infectious morbidities. *J Matern Fetal Neonatal Med*. 2008 Sep;21(9):638-42.
558. Costantine MM, Rahman M, Ghulmiyah L, Timing of perioperative antibiotics for cesarean delivery: a metaanalysis. *Am J Obstet Gynecol*. 2008 Sep;199(3):301.e1-6.
559. Tita AT, Hauth JC, Grimes A, et al. Decreasing incidence of postcesarean endometritis with extended-spectrum antibiotic prophylaxis. *Obstet Gynecol*. 2008 Jan;111(1):51-6.
560. Tita AT, Rouse DJ, Blackwell S, et al. Emerging concepts in antibiotic prophylaxis for cesarean delivery: a systematic review. *Obstet Gynecol*. 2009 Mar;113(3):675-82.
561. Hofmeyr GJ, Smaill FM. Antibiotic prophylaxis for cesarean section. *Cochrane Database Syst Rev*. 2010 Jan 20;(1):CD000933.

## 7. Modified active management of the third stage of labor

562. Vimala N, Mittal S, Kumar S, Dadhwal V, Mehta S. Sublingual misoprostol versus methylergometrine for active management of the third stage of labor. *Int J Gynaecol Obstet.* 2004 Oct;87(1):1-5.
563. Vimala N, Mittal S, Kumar S. Sublingual misoprostol versus oxytocin infusion to reduce blood loss at cesarean section. *Int J Gynaecol Obstet.* 2006 Feb;92(2):106-10.
564. Chandhiok N, Dhillon BS, Datey S, Mathur A, Saxena NC. Oral misoprostol for prevention of postpartum hemorrhage by paramedical workers in India. *Int J Gynaecol Obstet.* 2006 Feb;92(2):170-5.
565. Langenbach C. Misoprostol in preventing postpartum hemorrhage: a meta-analysis. *Int J Gynaecol Obstet.* 2006 Jan;92(1):10-8.
566. Baskett TF, Persad VL, Clough HJ, Young DC. Misoprostol versus oxytocin for the reduction of postpartum blood loss. *Int J Gynaecol Obstet.* 2007 Apr;97(1):2-5.
567. Ng PS, Lai CY, Sahota DS, Yuen PM. A double-blind randomized controlled trial of oral misoprostol and intramuscular syntometrine in the management of the third stage of labor. *Gynecol Obstet Invest.* 2007;63(1):55-60.
568. Alfrevic Z, Blum J, Walraven G, Weeks A, Winikoff B. Prevention of postpartum hemorrhage with misoprostol. *Int J Gynaecol Obstet.* 2007 Dec;99 Suppl 2:S198-201.
569. Mousa HA, Alfrevic Z. Treatment for primary postpartum haemorrhage. *Cochrane Database Syst Rev.* 2007 Jan 24;(1):CD003249.
570. Su LL, Chong YS, Samuel M. Oxytocin agonists for preventing postpartum haemorrhage. *Cochrane Database Syst Rev.* 2007 Jul 18;(3):CD005457.
571. Gürmezoglu AM, Forna F, Villar J, Hofmeyr GJ. Prostaglandins for preventing postpartum haemorrhage. *Cochrane Database Syst Rev.* 2007 Jul 18;(3):CD000494.
572. Hofmeyr GJ, Abdel-Aleem H, Abdel-Aleem MA. Uterine massage for preventing postpartum haemorrhage. *Cochrane Database Syst Rev.* 2008 Jul 16;(3):CD006431.
573. Sosa CG, Althabe F, Belizán JM, Buekens P. Risk factors for postpartum hemorrhage in vaginal deliveries in a Latin-American population. *Obstet Gynecol.* 2009 Jun;113(6):1313-9.
574. Althabe F, Alemán A, Tomasso G, et al. A pilot randomized controlled trial of controlled cord traction to reduce postpartum blood loss. *Int J Gynaecol Obstet.* 2009 Oct;107(1):4-7.
575. Audureau E, Deneux-Tharaux C, et al. Practices for prevention, diagnosis and management of postpartum haemorrhage: impact of a regional multifaceted intervention. *BJOG.* 2009 Sep;116(10):1325-33.
576. Leduc D, Senikas V, Lalonde AB, et al. Active management of the third stage of labour: prevention and treatment of postpartum hemorrhage. *J Obstet Gynaecol Can.* 2009 Oct;31(10):980-93.

577. Fekih M, Jnifene A, Fathallah K, et al. Benefit of misoprostol for prevention of postpartum hemorrhage in cesarean section: a randomized controlled trial. *J Gynecol Obstet Biol Reprod (Paris).* 2009 Nov;38(7):588-93.
578. Eftekhari N, Doroodian M, Lashkarizadeh R. The effect of sublingual misoprostol versus intravenous oxytocin in reducing bleeding after caesarean section. *J Obstet Gynaecol.* 2009 Oct;29(7):633-6.
579. Dapo OT, Akinola OI, Fawole AO, et al. Active management of third stage of labor: evidence versus practice. *Acta Obstet Gynecol Scand.* 2009;88(11):1252-60.
580. Blum J, Winikoff B, Raghavan S, et al. Treatment of postpartum haemorrhage with sublingual misoprostol versus oxytocin in women receiving prophylactic oxytocin: a double-blind, randomised, non-inferiority trial. *Lancet.* 2010 Jan 16;375(9710):217-23.

## 8. Delayed clamping of the umbilical cord

581. Ceriani Cernadas JM, Carroli G, Pellegrini L, et al. The effect of timing of cord clamping on neonatal venous hematocrit values and clinical outcome at term: a randomized, controlled trial. *Pediatrics.* 2006 Apr;117(4):e779-86.
582. Chaparro CM, Neufeld LM, Tena Alavez G, et al. Effect of timing of umbilical cord clamping on iron status in Mexican infants: a randomised controlled trial. *Lancet.* 2006 Jun 17;367(9527):1997-2004.
583. van Rheenen PF, Gruschke S, Brabin BJ. Delayed umbilical cord clamping for reducing anaemia in low birthweight infants: implications for developing countries. *Ann Trop Paediatr.* 2006 Sep;26(3):157-67.
584. Hutton EK, Hassan ES. Late vs early clamping of the umbilical cord in full-term neonates: systematic review and meta-analysis of controlled trials. *JAMA.* 2007 Mar 21;297(11):1241-52.
585. Baenziger O, Stolkin F, Keel M, et al. The influence of the timing of cord clamping on postnatal cerebral oxygenation in preterm neonates: a randomized, controlled trial. *Pediatrics.* 2007 Mar;119(3):455-9.
586. Rabe H, Reynolds G, Diaz-Rosello J. A systematic review and meta-analysis of a brief delay in clamping the umbilical cord of preterm infants. *Neonatology.* 2008;93(2):138-44.
587. McDonald SJ, Middleton P. Effect of timing of umbilical cord clamping of term infants on maternal and neonatal outcomes. *Cochrane Database Syst Rev.* 2008 Apr 16;(2):CD004074
588. Venâncio SI, Levy RB, Saldiva SR, et al. Effects of delayed cord clamping on hemoglobin and ferritin levels in infants at three months of age. *Cad Saude Publica.* 2008;24 Suppl 2:S323-31.
589. Jaleel R, Deeba F, Khan A. Timing of umbilical cord clamping and neonatal haematological status. *J Pak Med Assoc.* 2009 Jul;59(7):468-70.
590. Eichenbaum-Pikser G, Zasloff JS. Delayed clamping of the umbilical cord: a review with implications for practice. *J Midwifery Womens Health.* 2009 Jul-Aug;54(4):321-6.

591. Coggins M, Mercer J. Delayed cord clamping: advantages for infants. *Nurs Womens Health.* 2009 Apr;13(2):132-9.
592. Ononeze AB, Hutchon DJ. Attitude of obstetricians towards delayed cord clamping: a questionnaire-based study. *J Obstet Gynaecol.* 2009 Apr;29(3):223-4.
605. Anderson GC. Risk in mother-infant separation postbirth. *Image J Nurs Sch.* 1989 Winter;21(4):196-9.
606. Cottrell BH, Grubbs LM. Women's satisfaction with couplet care nursing compared to traditional postpartum care with rooming-in. *Res Nurs Health.* 1994 Dec;17(6):401-9.
607. Prodromidis M, Field T, Arendt R, et al. Mothers touching newborns: a comparison of rooming-in versus minimal contact. *Birth.* 1995 Dec;22(4):196-200
608. Ball HL, Ward-Platt MP, Heslop E, et al. Randomised trial of infant sleep location on the postnatal ward. *Arch Dis Child.* 2006 Dec;91(12):1005-10.

## IV. Maternal interventions after delivery

### 1. Early skin-to-skin contact

593. Curry MA. Maternal attachment behavior and the mother's self-concept: the effect of early skin-to-skin contact. *Nurs Res.* 1982 Mar-Apr;31(2):73-8.
594. Mikiel-Kostyra K, Boltruszko I, Mazur J, Zielenska M. Skin-to-skin contact after birth as a factor determining breastfeeding duration. *Med Wiek u Rozwoj.* 2001 Apr-Jun;5(2):179-89.
595. Mikiel-Kostyra K, Mazur J, Bołtruszko I. Effect of early skin-to-skin contact after delivery on duration of breastfeeding: a prospective cohort study. *Acta Paediatr.* 2002;91(12):1301-6.
596. Mizuno K, Mizuno N, Shinohara T, Noda M. Mother-infant skin-to-skin contact after delivery results in early recognition of own mother's milk odour. *Acta Paediatr.* 2004 Dec;93(12):1640-5.
597. Vaidya K, Sharma A, Dhungel S. Effect of early mother-baby close contact over the duration of exclusive breastfeeding. *Nepal Med Coll J.* 2005 Dec;7(2):138-40.
598. Miles R, Cowan F, Glover V, et al. A controlled trial of skin-to-skin contact in extremely preterm infants. *Early Hum Dev.* 2006 Jul;82(7):447-55.
599. Moore ER, Anderson GC. Randomized controlled trial of very early mother-infant skin-to-skin contact and breastfeeding status. *J Midwifery Womens Health.* 2007 Mar-Apr;52(2):116-25.
600. Moore ER, Anderson GC, Bergman N. Early skin-to-skin contact for mothers and their healthy newborn infants. *Cochrane Database Syst Rev.* 2007 Jul 18;(3):CD003519.
601. Bystrova K, Ivanova V, Edhborg M, et al. Early contact versus separation: effects on mother-infant interaction one year later. *Birth.* 2009 Jun;36(2):97-109.
602. Chiu SH, Anderson GC. Effect of early skin-to-skin contact on mother-preterm infant interaction through 18 months: randomized controlled trial. *Int J Nurs Stud.* 2009 Sep;46(9):1168-80.

### 2. Mother-newborn rooming-in

603. Winkelstein ML, Carson VJ. Adolescents and rooming-in. *Matern Child Nurs J.* 1987 Spring;16(1):75-88.
604. Norr KF, Roberts JE, Freese U. Early postpartum rooming-in and maternal attachment behaviors in a group of medically indigent primiparas. *J Nurse Midwifery.* 1989 Mar-Apr;34(2):85-91.

### 3. Detecting and managing puerperal endometritis

609. Soper DE. Postpartum endometritis. Pathophysiology and prevention. *J Reprod Med.* 1988 Jan;33(1 Suppl):97-100.
610. Casey BM, Cox SM. Chorioamnionitis and endometritis. *Infect Dis Clin North Am* 1997 Mar;11(1):203-22
611. Livingston JC, Llata E, Rinehart E, et al. Gentamicin and clindamycin therapy in postpartum endometritis: the efficacy of daily dosing versus dosing every 8 hours. *Am J Obstet Gynecol.* 2003 Jan;188(1):149-52.
612. French LM, Smaill FM. Antibiotic regimens for endometritis after delivery. *Cochrane Database Syst Rev.* 2004 Oct 18;(4):CD001067.
613. Haas DM, Morgan Al, Darei S, Contreras K. Vaginal preparation with antiseptic solution before cesarean section for preventing postoperative infections. *Cochrane Database Syst Rev.* 2010 Mar 17;3:CD007892.

### 4. Detecting, preventing, and managing postpartum depression

614. Ueda M, Yamashita H, Yoshida K. Impact of infant health problems on postnatal depression: pilot study to evaluate a health visiting system. *Psychiatry Clin Neurosci.* 2006 Apr;60(2):182-9.
615. Dennis CL, Hodnett E. Psychosocial and psychological interventions for treating postpartum depression. *Cochrane Database Syst Rev.* 2007 Oct 17;(4):CD006116.
616. van Doesum KT, Riksen-Walraven JM, et al. A randomized controlled trial of a home-visiting intervention aimed at preventing relationship problems in depressed mothers and their infants. *Child Dev.* 2008 May-Jun;79(3):547-61.
617. Dennis CL, Allen K. Interventions (other than pharmacological, psychosocial or psychological) for treating antenatal depression. *Cochrane Database Syst Rev.* 2008 Oct 8;(4):CD006795.
618. Cho HJ, Kwon JH, Lee JJ. Antenatal cognitive-behavioral therapy for prevention of postpartum depression: a pilot study. *Yonsei Med J.* 2008 Aug 30;49(4):553-62.
619. Leis JA, Mendelson T, Tandon SD, Perry DF. A systematic review of home-based interventions to prevent and treat postpartum depression. *Arch Womens Ment Health.* 2009 Feb;12(1):3-13.

620. Gjerdingen D, Crow S, McGovern P, et al. Stepped care treatment of postpartum depression: impact on treatment, health, and work outcomes. *J Am Board Fam Med.* 2009 Sep-Oct;22(5):473-82
621. Tronick E, Reck C. Infants of depressed mothers. *Harv Rev Psychiatry.* 2009;17(2):147-56.
622. Feldman R, Granat A, Pariente C, et al. Maternal depression and anxiety across the postpartum year and infant social engagement, fear regulation, and stress reactivity. *J Am Acad Child Adolesc Psychiatry.* 2009 Sep;48(9):919-27
623. Vik T, Grote V, Escribano J, et al. Infantile colic, prolonged crying and maternal postnatal depression. *Acta Paediatr.* 2009 Aug;98(8):1344-8.
624. Knudson-Martin C, Silverstein R. Suffering in silence: a qualitative meta-data-analysis of postpartum depression. *J Marital Fam Ther.* 2009 Apr;35(2):145-58.
625. Morrell CJ, Slade P, Warner R, et al. Clinical effectiveness of health visitor training in psychologically informed approaches for depression in postnatal women: pragmatic cluster randomised trial in primary care. *BMJ.* 2009 Jan 15;338:a3045.
626. Paulden M, Palmer S, Hewitt C, Gilbody S. Screening for postnatal depression in primary care: cost effectiveness analysis. *BMJ.* 2009 Dec 22;339:b5203.
627. Hewitt C, Gilbody S, Brealey S, et al. Methods to identify postnatal depression in primary care: an integrated evidence synthesis and value of information analysis. *Health Technol Assess.* 2009 Jul;13(36):1-145, 147-230.
628. American College of Obstetricians and Gynecologists. Committee on Obstetric Practice. Committee opinion no. 453: Screening for depression during and after pregnancy. *Obstet Gynecol.* 2010 Feb;115(2 Pt 1):394-5.
635. Ten VS, Matsiukevich D. Room air or 100% oxygen for resuscitation of infants with perinatal depression. *Curr Opin Pediatr.* 2009 Apr;21(2):188-93
636. Roy R. Question 1: is room air better than 100% oxygen for the resuscitation of the depressed full-term newborn? *Arch Dis Child.* 2010 Jan;95(1):68-70

## 2. Immediate care for the healthy newborn

637. Chen JY. Prophylaxis of ophthalmia neonatorum: comparison of silver nitrate, tetracycline, erythromycin and no prophylaxis. *Pediatr Infect Dis J.* 1992 Dec;11(12):1026-30
638. Martell M, Estol P, Martinez G y cols. Atención inmediata del recién nacido. *CLAP, OPS/OMS.* 1993, Publicación científica No. 1206.
639. PROSALUD. Guía para prevenir la hipotermia y regular la temperatura del recién nacido. La Paz, Bolivia 1994.
640. Isenberg SJ, Apt L, Wood M. A controlled trial of povidone-iodine as prophylaxis against ophthalmia neonatorum. *N Engl J Med.* 1995 Mar 2;332(9):562-6.
641. Puckett RM, Offringa M. Prophylactic vitamin K for vitamin K deficiency bleeding in neonates. *Cochrane Database Syst Rev.* 2000;(4):CD002776.
642. Figueras Aloy J, García Alix A, Alomar Ribes A, et al. Recommendations for the care of healthy newborn infants. *An Esp Pediatr.* 2001 Aug;55(2):141-5
643. Autret-Leca E, Jonville-Béra AP. Vitamin K in neonates: how to administer, when and to whom. *Paediatr Drugs.* 2001;3(1):1-8.
644. World Health Organization. Principles of Newborn Baby Care. In "Managing Newborn problems: A guide for doctors, nurses, and midwives. WHO/UNFPA/UNICEF/ The World Bank Group. 2003.
645. Narayanan I. Componentes del cuidado esencial del recién nacido. *BASICS II.* Honduras 2004.
646. Beck D, Ganges F, Goldman S, Long P. Care of the Newborn: Reference Manual. Save the Children, Washington DC, 2004.
647. Ali Z, Khadije D, Elahe A, Mohammad M, et al. Prophylaxis of ophthalmia neonatorum comparison of betadine, erythromycin and no prophylaxis. *J Trop Pediatr.* 2007 Dec;53(6):388-92.
648. Matinzadeh ZK, Beiragdar F, Kavemanesh Z, et al. Efficacy of topical ophthalmic prophylaxis in prevention of ophthalmia neonatorum. *Trop Doct.* 2007 Jan;37(1):47-9.
649. Matinzadeh ZK, Beiragdar F, Kavemanesh Z, et al. Efficacy of topical ophthalmic prophylaxis in prevention of ophthalmia neonatorum. *Trop Doct.* 2007 Jan;37(1):47-9.
650. McCall EM, Alderdice FA, Halliday HL, et al. Interventions to prevent hypothermia at birth in preterm and/or low birthweight infants. *Cochrane Database Syst Rev.* 2008 Jan 23;(1):CD004210.
651. Luna MS, Alonso CR, Mussons FB, et al. Recommendations for the care of the healthy normal newborn at delivery and during the first postnatal hours. *An Pediatr (Barc).* 2009 Oct;71(4):349-61.

## V. Neonatal interventions after delivery

### 1. Neonatal resuscitation with ambient air

629. Ramji S, Rasaily R, Mishra PK, et al. Resuscitation of asphyxiated newborns with room air or 100% oxygen at birth: a multicentric clinical trial. *Indian Pediatr.* 2003 Jun;40(6):510-7
630. Tan A, Schulze A, O'Donnell CP, Davis PG. Air versus oxygen for resuscitation of infants at birth. *Cochrane Database Syst Rev.* 2005 Apr 18;(2):CD002273
631. Saugstad OD, Ramji S, Vento M. Resuscitation of depressed newborn infants with ambient air or pure oxygen: a meta-analysis. *Biol Neonate.* 2005;87(1):27-34.
632. Rabi Y, Rabi D, Yee W. Room air resuscitation of the depressed newborn: a systematic review and meta-analysis. *Resuscitation.* 2007 Mar;72(3):353-63
633. Zhu JJ, Wu MY. Which is better to resuscitate asphyxiated newborn infants: room air or pure oxygen? *Zhonghua Er Ke Za Zhi.* 2007 Sep;45(9):644-9
634. Chabernaud JL. Air or oxygen for neonatal resuscitation at birth? *Arch Pediatr.* 2009 Aug;16(8):1194-201

### 3. Care of the umbilical cord

652. Panyavudhikrai S, Danchaivijitr S, Vantanasiri C, et al. Antiseptics for preventing omphalitis. *J Med Assoc Thai.* 2002 Feb;85(2):229-34.
653. Axelsson I. A Cochrane review on the umbilical cord care and prevention of infections. Antiseptic solutions are not necessary in developed countries but life-saving in developing countries. *Lakartidningen.* 2002 Mar 19;99(14):1563-6.
654. Janssen PA, Selwood BL, Dobson SR, et al. To dye or not to dye: a randomized, clinical trial of a triple dye/alcohol regime versus dry cord care. *Pediatrics.* 2003 Jan;111(1):15-20.
655. Pezzati M, Rossi S, Tronchin M, et al. Umbilical cord care in premature infants: the effect of two different cord-care regimens (salicylic sugar powder vs chlorhexidine) on cord separation time and other outcomes. *Pediatrics.* 2003 Oct;112(4):e275.
656. Zupan J, Garner P, Omari AA. Topical umbilical cord care at birth. *Cochrane Database Syst Rev.* 2004;(3):CD001057.
657. Shoaeib FM, All SA, El-Barawy MA. Alcohol or traditional methods versus natural drying for newborn's cord care. *J Egypt Public Health Assoc.* 2005;80(1-2):169-201.
658. Vural G, Kisa S. Umbilical cord care: a pilot study comparing topical human milk, povidone-iodine, and dry care. *J Obstet Gynecol Neonatal Nurs.* 2006 Jan-Feb;35(1):123-8.
659. Ahmadpour-Kacho M, Zahedpasha Y, Hajian K, et al. The effect of topical application of human milk, ethyl alcohol 96%, and silver sulfadiazine on umbilical cord separation time in newborn infants. *Arch Iran Med.* 2006 Jan;9(1):33-8.
660. Mullany LC, Darmstadt GL, Khatry SK, et al. Topical applications of chlorhexidine to the umbilical cord for prevention of omphalitis and neonatal mortality in southern Nepal: a community-based, cluster-randomised trial. *Lancet.* 2006 Mar 18;367(9514):910-8.
661. Mullany LC, El Arifeen S, Winch PJ, et al. Impact of 4.0% chlorhexidine cleansing of the umbilical cord on mortality and omphalitis among newborns of Sylhet, Bangladesh: design of a community-based cluster randomized trial. *BMC Pediatr.* 2009 Oct 21;9:67.
662. Kapellen TM, Gebauer CM, Brosteanu O, et al. Higher rate of cord-related adverse events in neonates with dry umbilical cord care compared to chlorhexidine powder. Results of a randomized controlled study to compare efficacy and safety of chlorhexidine powder versus dry care in umbilical cord care of the newborn. *Neonatology.* 2009;96(1):13-8.

### 4. Skin care at birth

663. Yosipovitch G, Maayan-Metzger A, Merlob P, Sirota L. Skin barrier properties in different body areas in neonates. *Pediatrics.* 2000 Jul;106(1 Pt 1):105-8.
664. Yoshio H, Tollin M, Gudmundsson GH, et al. Antimicrobial polypeptides of human vernix caseosa and amniotic fluid: implications for newborn innate defense. *Pediatr Res.* 2003 Feb;53(2):211-6.
665. Akinbi HT, Narendran V, Pass AK, et al. Host defense proteins in vernix caseosa and amniotic fluid. *Am J Obstet Gynecol.* 2004 Dec;191(6):2090-6.
666. Tollin M, Bergsson G, Kai-Larsen Y, et al. Vernix caseosa as a multi-component defence system based on polypeptides, lipids and their interactions. *Cell Mol Life Sci.* 2005 Oct;62(19-20):2390-9.
667. Walker VP, Akinbi HT, Meinzen-Derr J, et al. Host defense proteins on the surface of neonatal skin: implications for innate immunity. *J Pediatr.* 2008 Jun;152(6):777-81.
668. Adam R. Skin care of the diaper area. *Pediatr Dermatol.* 2008 Jul-Aug;25(4):427-33.
669. Kiechl-Kohlendorfer U, Berger C, Inzinger R. The effect of daily treatment with an olive oil/lanolin emollient on skin integrity in preterm infants: a randomized controlled trial. *Pediatr Dermatol.* 2008 Mar-Apr;25(2):174-8.
670. Visscher M, Odio M, Taylor T, et al. Skin care in the NICU patient: effects of wipes versus cloth and water on stratum corneum integrity. *Neonatology.* 2009;96(4):226-34.
671. Körner A, Dinten-Schmid B, Stoffel L, et al. Skin care and skin protection in preterm babies. *Pflege.* 2009 Aug;22(4):266-76.
672. Berger C, Inzinger R. Study of skin care in premature and newborn infants. *Kinderkrankenschwester.* 2009 Mar;28(3):116-25.

### 5. Breastfeeding in the first hour

673. Berens PD. Prenatal, intrapartum, and postpartum support of the lactating mother. *Pediatr Clin North Am.* 2001 Apr;48(2):365-75.
674. Rowe-Murray HJ, Fisher JR. Baby friendly hospital practices: cesarean section is a persistent barrier to early initiation of breastfeeding. *Birth.* 2002 Jun;29(2):124-31.
675. Cooke M, Cantrill R, and Creedy D. The first breastfeed: a content analysis of midwifery textbooks. *Breastfeed. Review* 2003;11:5-11.
676. Awi DD, Alikor EA. The influence of pre- and post-partum factors on the time of contact between mother and her new-born after vaginal delivery. *Nigeria Journal of Medicine* 2004;13:272-275.
677. Edmond KM, Zandoh C, Quigley MA, et al. Delayed breastfeeding initiation increases risk of neonatal mortality. *Pediatrics.* 2006 Mar;117(3):e380-6.
678. Chien LY, Tai CJ. Effect of delivery method and timing of breastfeeding initiation on breastfeeding outcomes in Taiwan. *Birth.* 2007 Jun;34(2):123-30.
679. Forster DA, McLachlan HL. Breastfeeding initiation and birth setting practices: a review of the literature. *J Midwifery Womens Health.* 2007 May-Jun;52(3):273-80.
680. DiGirolamo AM, Grummer-Strawn LM, Fein SB. Effect of maternity-care practices on breastfeeding. *Pediatrics.* 2008 Oct;122 Suppl 2:S43-9.
681. Keister D, Roberts KT, Werner SL. Strategies for breastfeeding success. *Am Fam Physician.* 2008 Jul 15;78(2):225-32.

682. Saloojee H. Contacto temprano piel a piel entre las madres y sus recién nacidos sanos: Comentario de la BSR. *La Biblioteca de Salud Reproductiva de la OMS*; Ginebra: Organización Mundial de la Salud, 2008.
683. Sweet L. Birth of a very low birth weight preterm infant and the intention to breastfeed 'naturally'. *Women Birth*. 2008 Mar;21(1):13-20.
684. Cramton R, Zain-Ul-Abideen M, Whalen B. Optimizing successful breastfeeding in the newborn. *Curr Opin Pediatr*. 2009 Jun;21(3):386-96.
685. Nommsen-Rivers LA, Mastergeorge AM, Hansen RL, et al. Doula care, early breastfeeding outcomes, and breastfeeding status at 6 weeks postpartum among low-income primiparae. *J Obstet Gynecol Neonatal Nurs*. 2009 Mar-Apr;38(2):157-73
- 6. Kangaroo mother program (skin-to-skin contact)**
686. Charpak N, Ruiz-Peláez JG, Charpak Y, Rey-Martinez. Kangaroo Mother Program: an alternative way of caring for low birth weight infants? One year mortality in a two cohort study. *Pediatrics*. 1994 Dec;94(6 Pt 1):804-10.
687. Charpak N, Ruiz-Peláez JG, Figueroa de Calume Z. Current knowledge of Kangaroo Mother Intervention. *Curr Opin Pediatr*. 1996 Apr;8(2):108-12.
688. Charpak N, Ruiz-Peláez JG, Figueroa de C Z, Charpak Y. Kangaroo mother versus traditional care for newborn infants </=2000 grams: a randomized, controlled trial. *Pediatrics*. 1997 Oct;100(4):682-8.
689. Tessier R, Cristo M, Velez S, Giron M, et al. Kangaroo mother care and the bonding hypothesis. *Pediatrics*. 1998 Aug;102(2):e17.
690. Conde-Agudelo A, Diaz-Rosello JL, Belizan JM. Kangaroo mother care to reduce morbidity and mortality in low birthweight infants. *Cochrane Database Syst Rev*. 2003;(2):CD002771.
691. Venancio SI, de Almeida H. Kangaroo-Mother Care: scientific evidence and impact on breastfeeding. *J Pediatr (Rio J)*. 2004 Nov;80(5 Suppl):S173-80.
692. Charpak N, Ruiz JG, Zupan J, Cattaneo A, et al. Kangaroo Mother Care: 25 years after. *Acta Paediatr*. 2005 May;94(5):514-22.
693. Moore ER, Anderson GC, Bergman N. Early skin-to-skin contact for mothers and their healthy newborn infants. *Cochrane Database Syst Rev*. 2007 Jul 18;(3):CD003519.
694. Erlandsson K, Dsilna A, Fagerberg I, Christensson K. Skin-to-skin care with the father after cesarean birth and its effect on newborn crying and prefeeding behavior. *Birth*. 2007 Jun;34(2):105-14.
695. Hake-Brooks SJ, Anderson GC. Kangaroo care and breastfeeding of mother-preterm infant dyads 0-18 months: a randomized, controlled trial. *Neonatal Netw*. 2008 May-Jun;27(3):151-9.
696. Kostandy RR, Ludington-Hoe SM, Cong X, et al. Kangaroo Care (skin contact) reduces crying response to pain in preterm neonates: pilot results. *Pain Manag Nurs*. 2008 Jun;9(2):55-65.
697. Tessier R, Charpak N, Giron M, et al. Kangaroo Mother Care, home environment and father involvement in the first year of life: a randomized controlled study. *Acta Paediatr*. 2009 Sep;98(9):1444-50.
698. Lawn JE, Mwansa-Kambafwile J, L HortaB, Barros FC, Cousens S. Kangaroo mother care to prevent neonatal deaths due to preterm birth complications. *International Journal of Epidemiology* 2010;39:i144-i154.
- 7. Stabilizing and transporting the neonate**
699. American Academy of Pediatrics. Safe transportation of premature and low birth weight infants. *Pediatrics* 1966;97:758-60.
700. Comité de Estándares de la Sociedad Española de Neonatología. Recomendaciones para el transporte perinatal. *An Esp Pediatr* 2001;55:146-53.
701. Bowen SL. Transport of the mechanically ventilated neonate. *Respir Care Clin N Am*. 2002 Mar;8(1):67-82.
702. Cambero O, Mazzi E, Rivera L. Transporte del neonato en estado crítico. En: Perinatología. 2<sup>a</sup> Ed. Mazzi E, Sandoval O Eds. Elite Impresiones, Bolivia 2002.
703. Annibale DJ, Cahill JB, Tuttle DS, et al. Preparation of the critically ill neonate for transport. *J S C Med Assoc*. 2002 Jun;98(3):129-36.
704. Lee SK, Zupancic JA, Sale J, et al. Cost-effectiveness and choice of infant transport systems. *Med Care*. 2002 Aug;40(8):705-16.
705. Neto MT. Regionalization, networks and neonatal transport. *J Matern Fetal Neonatal Med*. 2002 Feb;11(2):140.
706. American Academy of Pediatrics, American Heart Association. Reanimación neonatal. 2<sup>nd</sup> Ed. 2003.
707. Buckland L, Austin N, Jackson A, Inder T. Excessive exposure of sick neonates to sound during transport. *Arch Dis Child Fetal Neonatal Ed*. 2003 Nov;88(6):F513-6.
708. Hatherill M, Waggie Z, Reynolds L, Argent A. Transport of critically ill children in a resource-limited setting. *Intensive Care Med*. 2003 Sep;29(9):1547-54. Epub 2003 Aug 07.
709. Caverni V, Rastrelli M, Aufieri R, Agostino R. Can dedicated ambulances improve the efficiency of the neonatal emergency transport service? *J Matern Fetal Neonatal Med*. 2004 Feb;15(2):126-8.
710. Broughton SJ, Berry A, Jacob S, et al. The mortality index for neonatal transportation score: a new mortality prediction model for retrieved neonates. *Pediatrics*. 2004 Oct;114(4):e424-8.
711. Carreras Gonzalez E, Carreras Gonzalez G, Fraga Rodriguez G, et al. Critical care helicopter transport. Report of 224 cases. *An Pediatr (Barc)*. 2003 Dec;59(6):529-34.
712. Young T, Torner JC, Sihler KC, et al. Factors associated with mode of transport to acute care hospitals in rural communities. *J Emerg Med*. 2003 Feb;24(2):189-98.
713. Comette L. Contemporary neonatal transport: problems and solutions. *Arch Dis Child Fetal Neonatal Ed*. 2004 May;89(3):F212-4.

714. Das UG, Leuthner SR. Preparing the neonate for transport. *Pediatr Clin North Am.* 2004 Jun;51(3):581-98, vii.
715. Gill AB, Bottomley L, Chatfield S, Wood C. Perinatal transport: problems in neonatal intensive care capacity. *Arch Dis Child Fetal Neonatal Ed.* 2004 May;89(3):F220-3.
716. Karlsen K. The STABLE Program: Manual del participante, Park City, UT, USA 2001.
717. Respondek-Liberska M, Sysa A, Gadzinowski J. The cost of newborns transportation to the referral centers in comparison to the cost of the transport in-utero. *Ginekol Pol.* 2004 Apr;75(4):326-31.
718. Sontheimer D, Fischer CB, Buch KE. Kangaroo transport instead of incubator transport. *Pediatrics.* 2004 Apr;113(4):920-3.
719. Wagner C. Management of umbilical lines in neonatal transport. *Air Med J.* 2004 Jul-Aug;23(4):10-3.
720. Mullane D, Byrne H, Clarke TA, et al. Neonatal transportation: the effects of a national neonatal transportation programme. *Ir J Med Sci.* 2004 Apr-Jun;173(2):105-8.
721. Meberg A, Hansen TW. Quality evaluation of neonatal transports. *Tidsskr Nor Laegeforen.* 2005 Sep 22;125(18):2474-6.
722. McNamara P, Whyte H. Neonatal Transport. In: Residents Handbook of Neonatology. Perlman M, Moore AM, Kirpalani HM Eds. 3<sup>rd</sup> Ed. BC Decker Inc, Hamilton, 2007 pp 45-57.
723. Ramnarayan P. Measuring the performance of an inter-hospital transport service. *Arch Dis Child.* 2009 Jun;94(6):414-6.
- 8. Treating neonatal sepsis with a single daily dose of gentamicin**
724. Kiatchoosakun P, Kosalaraksa P, Jirapraditha J, et al. Once-daily gentamicin dosing of 4 Mg/Kg/dose in neonates. *J Med Assoc Thai.* 2005 Jul;88(7):934-8.
725. Nestaas E, Bangstad HJ, Sandvik L, Wathne KO. Aminoglycoside extended interval dosing in neonates is safe and effective: a meta-analysis. *Arch Dis Child Fetal Neonatal Ed.* 2005 Jul;90(4):F294-300.
726. Rao SC, Ahmed M, Hagan R. One dose per day compared to multiple doses per day of gentamicin for treatment of suspected or proven sepsis in neonates. *Cochrane Database Syst Rev.* 2006 Jan 25;(1):CD005091.
727. Darmstadt GL, Hossain MM, Jana AK, et al. Determination of extended-interval gentamicin dosing for neonatal patients in developing countries. *Pediatr Infect Dis J.* 2007 Jun;26(6):501-7.
728. Darmstadt GL, Miller-Bell M, Batra M, et al. Extended-interval dosing of gentamicin for treatment of neonatal sepsis in developed and developing countries. *J Health Popul Nutr.* 2008 Jun;26(2):163-82.
729. Serane TV, Zenguya S, Penford G, et al. Once daily dose gentamicin in neonates - is our dosing correct? *Acta Paediatr.* 2009 Jul;98(7):1100-5.
730. Hoff DS, Wilcox RA, Tolleson LM, et al. Pharmacokinetic outcomes of a simplified, weight-based, extended-interval gentamicin dosing protocol in critically ill neonates. *Pharmacotherapy.* 2009 Nov;29(11):1297-305.
731. Hossain MM, Chowdhury NA, Shirin M, et al. Simplified dosing of gentamicin for treatment of sepsis in Bangladeshi neonates. *J Health Popul Nutr.* 2009 Oct;27(5):640-5
- 9. Treating respiratory distress syndrome (RDS) with natural surfactant**
732. Halliday HL. History of surfactant from 1980. *Biol Neonate.* 2005;87(4):317-22.
733. Zimmermann LJ, Janssen DJ, Tibboel D, et al. Surfactant metabolism in the neonate. *Biol Neonate.* 2005;87(4):296-307
734. Mugford M. Cost effectiveness of prevention and treatment of neonatal respiratory distress (RDS) with exogenous surfactant: what has changed in the last three decades? *Early Hum Dev.* 2006 Feb;82(2):105-15.
735. Stevens TP, Harrington EW, Blennow M, Soll RF. Early surfactant administration with brief ventilation vs. selective surfactant and continued mechanical ventilation for preterm infants with or at risk for respiratory distress syndrome. *Cochrane Database Syst Rev.* 2007 Oct 17;(4):CD003063.
736. Halliday HL. Surfactants: past, present and future. *J Perinatol.* 2008 May;28 Suppl 1:S47-56.
737. Soll R, Ozek E. Multiple versus single doses of exogenous surfactant for the prevention or treatment of neonatal respiratory distress syndrome. *Cochrane Database Syst Rev.* 2009 Jan 21;(1):CD000141.
738. Seger N, Soll R. Animal derived surfactant extract for treatment of respiratory distress syndrome. *Cochrane Database Syst Rev.* 2009 Apr 15;(2):CD007836.
739. Pfister RH, Soll R, Wiswell TE. Protein-containing synthetic surfactant versus protein-free synthetic surfactant for the prevention and treatment of respiratory distress syndrome. *Cochrane Database Syst Rev.* 2009 Oct 7;(4):CD006180.
740. Sweet DG, Halliday HL. The use of surfactants in 2009. *Arch Dis Child Educ Pract Ed.* 2009 Jun;94(3):78-83.
741. Carnielli VP, Zimmermann LJ, Hamvas A, Cogo PE. Pulmonary surfactant kinetics of the newborn infant: novel insights from studies with stable isotopes. *J Perinatol.* 2009 May;29 Suppl 2:S29-37.
- 10. Neonatal screening to detect congenital hypothyroidism, galactosemia, phenylketonuria, and cystic fibrosis**
742. Velázquez A, et al. Tamiz neonatal para hipotiroidismo congénito y fenilcetonuria. *Sal Pub Mex* 1994; 36: 249-256.
743. Walter JH. Arguments for early screening: A clinician's perspective. *Europ J Pediatr* 2003; 162 (suppl 1): S2-S4.
744. Schweitzer-Krantz S. Early diagnosis of inherited metabolic disorders towards improving outcome: The controversial issue of galactosaemia. *Europ J Pediatr* 2003; 162 (suppl 1): S50- S53.

745. Wilcken B. Ethical issues in newborn screening and the impact of new technologies. *Euro J Pediatr* 2003; 162 (suppl 1): S62-S66.
746. Bonham JR et al. Screening for cystic fibrosis: the practice and the debate. *Euro J Pediatr* 2003; 162 (suppl 1): S42-S45.
747. Springer-Verlag Heidelberg. Quality control requirements in neonatal screening. *Euro J Pediatr* Vol 2003; 162 (suppl 1): S54-S56.
748. Liebl B et al. Data required for the evaluation of newborn screening programmes. *Euro J Pediatr* 2003; 162 (suppl 1): S57-S61.
749. Liebl B et al. Data required for the evaluation of newborn screening programmes. *Euro J Pediatr* 2003; 162 (suppl 1): S57-S61.
750. Röschinger W et al. Advances in analytical mass spectrometry to improve screening for inherited metabolic diseases. *Euro J Pediatr* 2003; 162 (suppl 1): S67-S76.
751. Banta SA et al. Tandem mass spectrometry in newborn screening: A primer for neonatal and perinatal nurses. *J Perinat Neonat Nurs* 2004; 18 (1): 41-59.
752. Barba JR. Tamiz neonatal: una estrategia en la medicina preventiva. *Rev Mex Patol Clin*, 2004; Vol. 51,(Núm. 3): 130-144
753. Tylek-Lemanska D, Kumorowicz-Kopiec M, Starzyk J. Screening for congenital hypothyroidism: the value of retesting after four weeks in neonates with low and very low birth weight. *J Med Screen*. 2005;12(4):166-9.
754. Büyükgeliz A. Newborn screening for congenital hypothyroidism. *J Pediatr Endocrinol Metab*. 2006 Nov;19(11):1291-8.
755. Kempers MJ, Lanting CI, van Heijst AF, et al. Neonatal screening for congenital hypothyroidism based on thyroxine, thyrotropin, and thyroxine-binding globulin measurement: potentials and pitfalls. *J Clin Endocrinol Metab*. 2006 Sep;91(9):3370-6.
756. Grüters A, Krude H. Update on the management of congenital hypothyroidism. *Horm Res*. 2007;68 Suppl 5: 107-11.
757. Comeau AM, Accurso FJ, White TB, et al. Guidelines for implementation of cystic fibrosis newborn screening programs: Cystic Fibrosis Foundation workshop report. *Pediatrics*. 2007 Feb;119(2):e495-518.
758. Borrago GJC. Newborn screening in Latin America at the beginning of the 21st century. *J Inher Metab Dis* 2007;30(4):466-481.
759. Korada M, Kibirige M, Turner S, et al. The implementation of revised guidelines and the performance of a screening programme for congenital hypothyroidism. *J Med Screen*. 2008;15(1):5-8.
760. Castellani C, Southern KW, Brownlee K, et al. European best practice guidelines for cystic fibrosis neonatal screening. *J Cyst Fibros*. 2009 May;8(3):153-73.
- 11. Detecting and preventing tuberculosis and hepatitis B**
761. Hanekom WA. The immune response to BCG vaccination of newborns. *Ann NY Acad Sci*. 2005 Dec;1062:69-78.
762. Davids V, Hanekom WA, Mansoor N, et al. The effect of bacille Calmette-Guérin vaccine strain and route of administration on induced immune responses in vaccinated infants. *J Infect Dis*. 2006 Feb 15;193(4):531-6.
763. Lee C, Gong Y, Brok J, Boxall EH, Gluud C. Effect of hepatitis B immunisation in newborn infants of mothers positive for hepatitis B surface antigen: systematic review and meta-analysis. *BMJ*. 2006 Feb 11;332(7537):328-36.
764. Hawkrige A, Hatherill M, Little F, et al. Efficacy of percutaneous versus intradermal BCG in the prevention of tuberculosis in South African infants: randomised trial. *BMJ*. 2008 Nov 13;337:a2052. doi: 10.1136/bmj.a2052.
765. Pickering LK, Baker CJ, Freed GL, et al. Immunization programs for infants, children, adolescents, and adults: clinical practice guidelines by the Infectious Diseases Society of America. *Clin Infect Dis*. 2009 Sep 15;49(6):817-40. Advisory Committee on Immunization Practices. The Recommended Immunization Schedules for Persons Aged 0 through 18 Years. AAP/AAFP/CDC, 2010.
- 12. Detecting and treating retinopathy of prematurity in the newborn (ROP)**
766. Karna P, Muttineni J, Angell L, Karmaus W. Retinopathy of prematurity and risk factors: a prospective cohort study. *BMC Pediatr*. 2005 Jun 28;5(1):18.
767. Huang ZN, Yin DM, Huang DR, Liang SX. Screening analysis of retinopathy of prematurity and treatment of threshold retinopathy of prematurity. *Zhonghua Yan Ke Za Zhi*. 2006 Jun;42(6):496-500.
768. Cervantes-Munguia R, Espinosa-Lopez L, Gomez-Contreras P, et al. Retinopathy of prematurity and oxidative stress. *Anales de Pediatría* 2006;64(2):126-131.
769. DiBiasie A. Evidence-based review of retinopathy of prematurity prevention in VLBW and ELBW infants. *Neonatal Netw*. 2006 Nov-Dec;25(6):393-403.
770. Harrell SN, Brandon DH. Retinopathy of prematurity: the disease process, classifications, screening, treatment, and outcomes. *Neonatal Netw*. 2007 Nov-Dec;26(6):371-8.
771. Ahmed AS, Muslina H, Anwar KS, et al. Retinopathy of Prematurity in Bangladeshi Neonates. *J Trop Pediatr*. 2008 May 25.
772. Jandeck C, Kellner U, Lorenz B, Seiberth V. Guidelines for ophthalmologic screening of premature infants. *Ophthalmologe*. 2008 Oct;105(10):955-63.
773. Chawla D, Agarwal R, Deorari AK, Paul VK. Retinopathy of prematurity. *Indian J Pediatr*. 2008 Jan;75(1):73-6.
774. Askin DF, Diehl-Jones W. Retinopathy of prematurity. *Crit Care Nurs Clin North Am*. 2009 Jun;21(2):213-33.

775. Fortes Filho JB, Eckert GU, Procianoy L, et al. Incidence and risk factors for retinopathy of prematurity in very low and in extremely low birth weight infants in a unit-based approach in southern Brazil. *Eye (Lond)*. 2009 Jan;23(1):25-30.
776. Carlo WA, Finer NN, Walsh MC, et al. Target ranges of oxygen saturation in extremely preterm infants. *N Engl J Med*. 2010 May 27;362(21):1959-69.
777. Kiyota M, Hiraoka M, Watanabe T. Optimum timing of initial examination for retinopathy of prematurity. *Nippon Ganka Gakkai Zasshi*. 2010 Apr;114(4):356-61.
778. Christiansen SP, Dobson V, Quinn GE, et al. (Early Treatment for Retinopathy of Prematurity Cooperative Group). Progression of type 2 to type 1 retinopathy of prematurity in the Early Treatment for Retinopathy of Prematurity Study. *Arch Ophthalmol*. 2010 Apr;128(4):461-5.
779. Wu C, Vanderveen DK, Hellström A, et al. Longitudinal postnatal weight measurements for the prediction of retinopathy of prematurity. *Arch Ophthalmol*. 2010 Apr;128(4):443-7.
- 13. Early home visits for newborn care**
780. Bang AT, Bang RA, Baitule SB, et al. Effect of home-based neonatal care and management of sepsis on neonatal mortality: field trial in rural India. *Lancet* 1999 Dec 4;354(9194):1955-61.
781. Baqui AH, El-Arifeen S, Darmstadt GL, et al. Projahnmo Study Group. Effect of community-based newborn-care intervention package implemented through two service delivery strategies in Sylhet district, Bangladesh: a cluster-randomised controlled trial. *Lancet* 2008 Jun 7;371(9628):1936-44.
782. Kumar V, Mohaty S, Kumar A, et al. Saksham Study Group. Effect of community-based behaviour change management on neonatal mortality in Shivgarh, Uttar Pradesh, India: a cluster-randomised controlled trial. *Lancet* 2008 Sep 27;372(9644):1151-62.
783. Bhutta ZA, Memon ZA, Soofi S, et al. Implementing community-based perinatal care: results from a pilot study in rural Pakistan. *Bull World Health Organ* 2008 Jun;86(6):452-9.
784. WHO and UNICEF Joint Statement. Home visits for the care of the newborn child: complementary strategy to improve newborn survival. 2008.
785. Mannan I, Rahman SM, Sania A, et al. Can early postpartum home visits by trained community health workers improve breastfeeding of newborns? *J Perinatol*. 2008 Sep;28(9):632-40.
786. Ogbuanu CA, Jones CA, McTigue JF, et al. A program evaluation of postpartum/newborn home visitation services in aiken county, South Carolina. *Public Health Nurs*. 2009 Jan-Feb;26(1):39-47.
- 14. Supine sleeping position and sudden infant death syndrome (SIDS)**
787. Willinger M, Hoffman HJ, Hartford RB. Infant sleep position and risk for sudden infant death syndrome: report of meeting held January 13 and 14, 1994, National Institutes of Health, Bethesda, MD. *Pediatrics*. 1994 May;93(5):814-9.
788. [No authors listed]. Positioning and sudden infant death syndrome (SIDS): update. American Academy of Pediatrics Task Force on Infant Positioning and SIDS. *Pediatrics*. 1996 Dec;98(6 Pt 1):1216-8.
789. [No authors listed]. Changing concepts of sudden infant death syndrome: implications for infant sleeping environment and sleep position. American Academy of Pediatrics. Task Force on Infant Sleep Position and Sudden Infant Death Syndrome. *Pediatrics*. 2000 Mar;105(3 Pt 1):650-6.
790. Moon RY, Oden RP, Grady KC. Back to Sleep: an educational intervention with women, infants, and children program clients. *Pediatrics*. 2004 Mar;113(3 Pt 1):542-7.
791. Gilbert R, Salanti G, Harden M, See S. Infant sleeping position and the sudden infant death syndrome: systematic review of observational studies and historical review of recommendations from 1940 to 2002. *Int J Epidemiol*. 2005 Aug;34(4):874-87.
792. Colson ER, Levenson S, Rybin D, et al. Barriers to following the supine sleep recommendation among mothers at four centers for the Women, Infants, and Children Program. *Pediatrics*. 2006 Aug;118(2):e243-50.
793. Moon RY, Calabrese T, Aird L. Reducing the risk of sudden infant death syndrome in child care and changing provider practices: lessons learned from a demonstration project. *Pediatrics*. 2008 Oct;122(4):788-98.
794. Dwyer T, Ponsonby AL. Sudden infant death syndrome and prone sleeping position. *Ann Epidemiol*. 2009 Apr;19(4):245-9.
- 15. Exclusive breastfeeding**
795. Kramer MS, Kakuma R. Optimal duration of exclusive breastfeeding. *Cochrane Database Syst Rev*. 2002;(1):CD003517.
796. Kramer MS, Kakuma R. The optimal duration of exclusive breastfeeding: a systematic review. *Adv Exp Med Biol*. 2004;554:63-77.
797. Fairbank L, O'Meara S, Renfrew MJ, et al. A systematic review to evaluate the effectiveness of interventions to promote the initiation of breastfeeding. *Health Technol Assess*. 2000;4(25):1-171.
798. Owen CG, Martin RM, Whincup PH, et al. Effect of infant feeding on the risk of obesity across the life course: a quantitative review of published evidence. *Pediatrics*. 2005 May;115(5):1367-77.
799. Ip S, Chung M, Raman G, et al. Breastfeeding and maternal and infant health outcomes in developed countries. *Evid Rep Technol Assess (Full Rep)*. 2007 Apr;(153):1-186.

800. Flint A, New K, Davies MW. Cup feeding versus other forms of supplemental enteral feeding for newborn infants unable to fully breastfeed. *Cochrane Database Syst Rev*. 2007 Apr 18;(2):CD005092.
801. Academy of Breastfeeding Medicine Board of Directors. Position on breastfeeding. *Breastfeed Med*. 2008 Dec;3(4):267-70.
802. Boccolini CS, Carvalho ML, Oliveira MI, et al. Factors that affect time between birth and first breastfeeding. *Cad Saude Publica*. 2008 Nov;24(11):2681-94.
803. U.S. Preventive Services Task Force. Primary care interventions to promote breastfeeding: U.S. Preventive Services Task Force recommendation statement. *Ann Intern Med*. 2008 Oct 21;149(8):560-4.
804. Collins CT, Makrides M, Gillis J, McPhee AJ. Avoidance of bottles during the establishment of breast feeds in preterm infants. *Cochrane Database Syst Rev*. 2008 Oct 8;(4):CD005252.
805. Rosen IM, Krueger MV, Carney LM, Graham JA. Prenatal breastfeeding education and breastfeeding outcomes. *MCN Am J Matern Child Nurs*. 2008 Sep-Oct;33(5):315-9.
806. Owen CG, Whincup PH, Kaye SJ, et al. Does initial breastfeeding lead to lower blood cholesterol in adult life? A quantitative review of the evidence. *Am J Clin Nutr*. 2008 Aug;88(2):305-14.
807. Ehlayel MS, Bener A. Duration of breast-feeding and the risk of childhood allergic diseases in a developing country. *Allergy Asthma Proc*. 2008 Jul-Aug;29(4):386-91.
808. Renfrew MJ, Craig D, Dyson L, et al. Breastfeeding promotion for infants in neonatal units: a systematic review and economic analysis. *Health Technol Assess*. 2009 Aug;13(40):1-146, iii-iv.
809. Kim Y. Effects of a breast-feeding empowerment program on exclusive breast-feeding. *J Korean Acad Nurs*. 2009 Apr;39(2):279-87.
810. Academy of Breastfeeding Medicine Protocol Committee. Clinical protocol number #19: breastfeeding promotion in the prenatal setting. *Breastfeed Med*. 2009 Mar;4(1):43-5.
811. Bartick M, Stuebe A, Shealy KR, et al. Closing the quality gap: promoting evidence-based breastfeeding care in the hospital. *Pediatrics*. 2009 Oct;124(4):e793-802.
812. Petrova A, Ayers C, Stechna S, et al. Effectiveness of exclusive breastfeeding promotion in low-income mothers: a randomized controlled study. *Breastfeed Med*. 2009 Jun;4(2):63-9.
813. Hoddinott P, Britten J, Prescott GJ, et al. Effectiveness of policy to provide breastfeeding groups (BIG) for pregnant and breastfeeding mothers in primary care: cluster randomised controlled trial. *BMJ*. 2009 Jan 30;338:a3026. doi: 10.1136/bmj.a3026.
814. Racine EF, Frick KD, Strobino D, et al. How motivation influences breastfeeding duration among low-income women. *J Hum Lact*. 2009 May;25(2):173-81.
815. Manganaro R, Marseglia L, Mamì C, et al. Effects of hospital policies and practices on initiation and duration of breastfeeding. *Child Care Health Dev*. 2009 Jan;35(1):106-11.
816. Daelmans B, Mangasaryan N, Martines J, et al. Strengthening actions to improve feeding of infants and young children 6 to 23 months of age: summary of a recent World Health Organization/UNICEF technical meeting, Geneva, 6-9 October 2008. *Food Nutr Bull*. 2009 Jun;30(2 Suppl):S236-8.
817. Lutter Chessa. El inicio temprano de la lactancia materna: la clave para supervivencia y desarrollo. OPS/AECID. 2010.

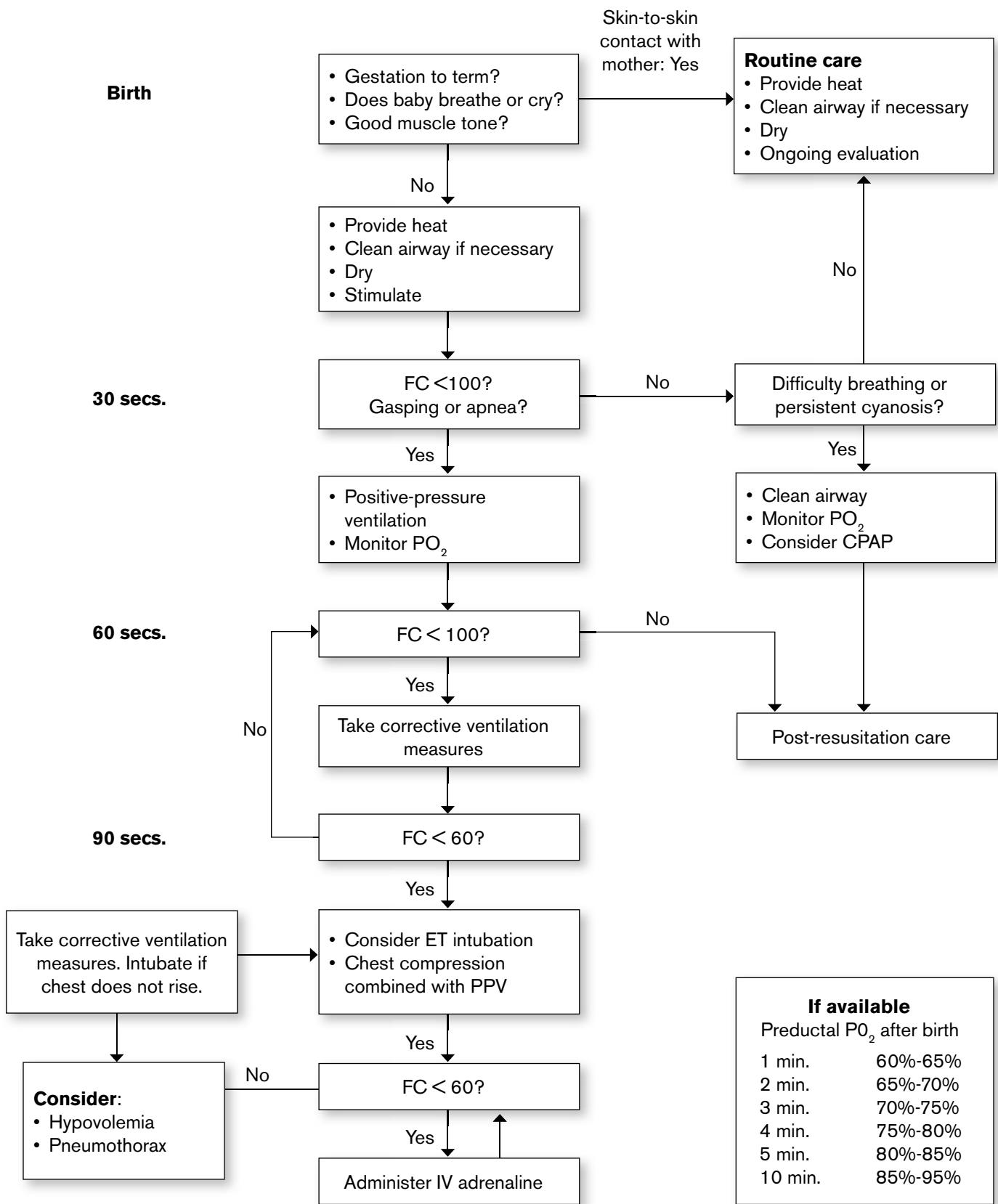


# VIII. Annexes

## **Recommendations of the World Health Organization (WHO)**

- > Each woman has a fundamental right to receive proper prenatal care, and plays a central role in all aspects of this care, including participation in the planning, carrying out and evaluation of the care; and social, emotional and psychological factors are decisive in the understanding and implementation of proper prenatal care.
- > Information about birth practices in hospitals (rates of cesarean sections, etc.) should be given to the public served by the hospitals.
- > There is no justification in any specific geographic region to have more than 10-15% cesarean section births.
- > There is no evidence that routine electronic fetal monitoring during labor has a positive effect on the outcome of pregnancy.
- > There is no indication for pubic shaving or a pre-delivery enema.
- > Pregnant women should not be put in a lithotomy (flat on the back) position during labor or delivery. They should be encouraged to walk during labor and each woman must freely decide which position to adopt during delivery.
- > The systematic use of episiotomy (incision to enlarge the vaginal opening) is not justified.
- > Birth should not be induced (started artificially) for convenience and the induction of labor should be reserved for specific medical indications. No geographic region should have rates of induced labor over 10%.
- > Artificial early rupture of the membranes, as a routine process, is not scientifically justified.
- > The healthy newborn must remain with the mother whenever both their conditions permit it. No process of observation of the healthy newborn justifies a separation from the mother.
- > The immediate beginning of breastfeeding should be promoted, even before the mother leaves the delivery room.

### Neonatal resuscitation flowchart (AAP 2010)



# Evaluate and determine risk before pregnancy

ASK	DETERMINE OBSERVE
<ul style="list-style-type: none"> <li>• How old are you?</li> <li>• Do you have a stable partner?</li> <li>• Do you have sexual relations?</li> <li>• Do you and your partner use some method of family planning?</li> <li>• Have you had vaginal discharge?</li> <li>• Do you use alcohol, tobacco, or drugs?</li> <li>• Have you had contact with insecticides or chemicals?</li> <li>• If there have been previous pregnancies, investigate history of: perinatal deaths, low birthweight, premature birth, abortions, congenital malformations of the neural tube (CMFTNs)</li> <li>• Have you had any chronic disease?</li> <li>• Have you been the victim of any type of violence?</li> </ul>	<ul style="list-style-type: none"> <li>• Weight</li> <li>• Size</li> <li>• BMI</li> <li>• Hb</li> <li>• STI</li> <li>• Discharge</li> <li>• Pallor</li> <li>• Oral cavity (pain, bleeding, inflammation, halitosis, caries)</li> <li>• Vaccination series</li> <li>• Have you had contact with insecticides or chemicals?</li> <li>• If there have been previous pregnancies, investigate history of: perinatal deaths, low birthweight, premature birth, abortions, congenital malformations of the neural tube (CMFTNs)</li> <li>• Previous birth defects</li> <li>• Multiple sexual partners</li> <li>• Not using family planning</li> <li>• STI under treatment</li> </ul>

EVALUATE	CATEGORIZE	TREAT
Classify	<p><b>One of the following signs:</b></p> <ul style="list-style-type: none"> <li>• &lt; 19 years old</li> <li>• BMI &lt; 18.5 or &gt; 29.9</li> <li>• Hb &lt; 7 g/dl or intense pallor</li> <li>• Untreated sexually transmitted infections (STI)</li> <li>• Previous disease brought under control</li> <li>• Alcohol, tobacco, or drug use</li> <li>• High risk for malformations of the neural tube</li> <li>• 35 or older</li> <li>• BMI &gt; 25.0</li> <li>• Hb between 7 and 12 g/dl or moderate pallor</li> <li>• Previous disease brought under control</li> <li>• Oral health problems</li> <li>• Exposed to chemicals or insecticides</li> <li>• Perinatal death, low birthweight, premature birth, previous abortion</li> <li>• Previous birth defects</li> <li>• Multiple sexual partners</li> <li>• Not using family planning</li> <li>• STI under treatment</li> </ul>	<p><b>PREGNANCY NOT ADVISED, or delay advised</b></p> <ol style="list-style-type: none"> <li>1. Transfer if necessary, or treat if able.</li> <li>2. Control existing disease.</li> <li>3. Provide nutritional counseling.</li> <li>4. Seek cause and treat anemia with iron.</li> <li>5. Congenital anomalies and neural tube defects: folic acid, 4-5 mg orally daily.</li> <li>6. Deparasitize with albendazole.</li> <li>7. Provide family planning.</li> <li>8. If positive VDRL test:, administer benzathine benzylpenicillin, 2.4 mill IU, IM.</li> </ol>
		<p><b>IN A POSITION TO BECOME PREGNANT BUT WITH RISK FACTORS</b></p> <ol style="list-style-type: none"> <li>1. Nutritional counseling and adequate diet.</li> <li>2. Give iron.</li> <li>3. Folic acid, 1.0 mg daily orally 3 months before pregnancy.</li> <li>4. Deparasitize with albendazole.</li> <li>5. Provide family planning.</li> <li>6. Personal hygiene.</li> <li>7. Oral prophylaxis and treatment.</li> <li>8. Counsel on prevention of breast and cervical cancer (HPV vaccination).</li> </ol>
		<p><b>IN A POSITION TO BECOME PREGNANT</b></p> <ol style="list-style-type: none"> <li>1. Give folic acid, 1.0 mg/day orally 3 months before pregnancy.</li> <li>2. Family planning.</li> <li>3. Personal hygiene.</li> <li>4. Oral hygiene.</li> <li>5. Counsel on prevention of breast and cervical cancer (HPV vaccination).</li> <li>6. Counsel on healthy lifestyle: nutrition, exercise, prevention of exposure to toxic substances and infection.</li> </ol>

**Risk factors for malformations of the neural tube:**  
 Exposure to anticonvulsive drugs, maternal diabetes, low socioeconomic level, maternal malnutrition, deficiency of folic acid, maternal hyperthermia, genetic factors, agricultural work, exposure to pesticides, occupational type, contact with pesticides during pregnancy.

# Evaluate and determine risk during pregnancy

DETERMINE SIGNS AND SYMPTOMS OF DANGER	
ASK	DETERMINE
<ul style="list-style-type: none"> <li>How old are you?</li> <li>Have you had a prenatal check-up? How many times?</li> <li>When was your last delivery?</li> <li>Have deliveries been natural or via caesarean section?</li> <li>How many pregnancies have you had?</li> <li>Have you had premature or low birthweight children?</li> <li>Have you had children with malformations?</li> <li>Have you had a child die before birth or during the first week of life?</li> <li>Do you feel the baby move?</li> <li>Have you had a fever?</li> <li>Have you had vaginal discharge?</li> <li>Do you have any disease? What?</li> <li>Have they given you any drugs? What?</li> <li>Have you lost consciousness?</li> <li>Do you smoke, drink, or use drugs?</li> </ul>	<ul style="list-style-type: none"> <li>Probable delivery date</li> <li>Gestational age</li> <li>Weight</li> <li>Size</li> <li>BMI</li> <li>Uterine height</li> <li>Multiple gestation</li> <li>Previous caesarean section</li> <li>Blood pressure</li> <li>Intense palmar pallor</li> <li>Swelling in face, hands, and/or legs</li> <li>Signs of systemic and/or sexually transmitted disease</li> <li>Immunization with tetanus toxoid</li> <li>Condition of oral cavity (bleeding, inflammations)</li> <li>Moderate palmar pallor and/or Hb 8-10 g/dL</li> <li>Teratogenic drug intake</li> <li>Alcoholism, smoking, or drug addiction</li> <li>Controlled hypertension</li> <li>Inadequate weight gain</li> <li>Abnormal presentation</li> <li>Multiple gestation</li> <li>Mother Rh negative</li> <li>Positive VDRL, HIV, or hepatitis B test</li> <li>Oral health problems</li> <li>Not immunized with tetanus toxoid</li> </ul>

EVALUATE	CLASSIFY	TREAT
<p><b>Presence of any of the following signs:</b></p> <ul style="list-style-type: none"> <li>Pregnancy over 41 weeks</li> <li>Diminished or absent fetal movement</li> <li>Severe systemic disease</li> <li>Urinary infection with fever</li> <li>Uncontrolled diabetes</li> <li>Uncontrolled hypertension and/or convulsions, blurred vision, loss of consciousness, or intense cephalgia</li> <li>Severe palmar pallor and/or Hb &lt; 7 g/dL</li> <li>PROM before 37 weeks</li> </ul>	<b>PREGNANCY WITH IMMINENT RISK</b>	<ol style="list-style-type: none"> <li><b>Refer URGENTLY to facility offering more complex services.</b></li> <li>Treat hypertension.</li> <li>In case of PPROM, administer initial dose of erythromycin, 250 mg orally.</li> <li>If GA age is between 26 and 36 weeks, administer initial dose of prenatal steroids.</li> </ol>
<p><b>One of the following signs:</b></p> <ul style="list-style-type: none"> <li>Under 19 years or over 35 years old</li> <li>First pregnancy or large multiple fetuses</li> <li>Without prenatal check-up</li> <li>Period between pregnancies &lt; 2 years</li> <li>Uterine height does not correlate with gestational age</li> <li>Previous caesarean section</li> <li>Background of premature, low birthweight, and/or malformed children</li> <li>Background of early miscarriage, fetal or neonatal death</li> <li>Controlled systemic disease</li> <li>Urinary infection without fever</li> <li>Moderate palmar pallor and/or Hb 8-10 g/dL</li> <li>Vaginal discharge</li> <li>Teratogenic drug intake</li> <li>Alcoholism, smoking, or drug addiction</li> <li>Controlled hypertension</li> <li>Inadequate weight gain</li> <li>Abnormal presentation</li> <li>Multiple gestation</li> <li>Mother Rh negative</li> <li>Positive VDRL, HIV, or hepatitis B test</li> <li>Oral health problems</li> <li>Not immunized with tetanus toxoid</li> </ul>	<b>HIGH-RISK PREGNANCY</b>	<ol style="list-style-type: none"> <li><b>Refer for consultation by specialist.</b></li> <li>In case of multiple gestation, refer before 30 weeks of gestation. If VDRL test is positive, initiate treatment with benzathine penicillin.</li> <li>Advise the mother to continue with the treatment instituted.</li> <li>Administer iron, folic acid, and multivitamins.</li> <li>If there is a history of hypertension, administer calcium + aspirin (100 mg/day) supplement.</li> <li>Administer tetanus toxoid.</li> <li>In areas of high prevalence, administer albendazole.</li> <li>Provide HIV/AIDS-STI counseling.</li> <li>Decide on reconsultation.</li> <li>Provide counseling on nutrition, pregnancy care, and breastfeeding.</li> <li>Teach warning signs.</li> <li>Plan for referral with the family as risk factors and response capacity dictate.</li> </ol>
	<b>LOW-RISK PREGNANCY</b>	<ul style="list-style-type: none"> <li>Teach warning signs.</li> <li>Plan with the family for delivery in the health care facility.</li> <li>Monitor throughout pregnancy.</li> <li>Provide counseling on nutrition, pregnancy care, puerperium, breastfeeding, and vaccines for the infant.</li> <li>Provide HIV/AIDS counseling.</li> <li>Advise the mother to continue with the treatment instituted.</li> <li>Administer iron, folic acid, and multivitamins.</li> <li>Administer tetanus toxoid.</li> </ul>

## Evaluate and determine risk during delivery

DETERMINE SIGNS AND SYMPTOMS OF DANGER		ASK	DETERMINE	EVALUATE	CLASSIFY	TREAT
		<ul style="list-style-type: none"> <li>When was your last menstruation?</li> <li>Have you had labor pains?</li> <li>Have you had vaginal bleeding?</li> <li>Have you had vaginal discharge? What color?</li> <li>Have you had severe headache?</li> <li>Have you had blurred vision?</li> <li>Have you had convulsions?</li> </ul>	<ul style="list-style-type: none"> <li>Blood pressure</li> <li>Temperature</li> <li>Presence of contractions within 10 minutes</li> <li>Fetal heart rate</li> <li>Cervical dilation and appearance</li> <li>Swelling in face, hands, and/or legs</li> <li>Vaginal bleeding now or earlier</li> </ul>	Classify	<p><b>Presence of any of the following signs:</b></p> <ul style="list-style-type: none"> <li>Labor in progress at less than 37 weeks</li> <li>Vaginal bleeding</li> <li>Premature rupture of membranes (PROM) &gt; 12 hours</li> <li>Uncontrolled hypertension and/or convulsions</li> <li>blurred vision, loss of consciousness, or intense headache</li> <li>Changes in fetal heart rate (FHR) (&lt;120 or &gt;160/minute)</li> <li>Abnormal fetal presentation in labor</li> <li>Swelling in face, hands, or legs</li> </ul>	<b>PREGNANCY WITH IMMINENT RISK</b> <ol style="list-style-type: none"> <li>Refer URGENTLY to a facility that offers more complex care, with the patient lying on her left side.</li> <li>Prevent hypotension.</li> <li>Treat hypertension.</li> <li>In cases of premature labor, inhibit contractions with nifedipine and administer corticoids (betamethasone or dexamethasone).</li> <li>In cases of PROM, administer the first dose of the appropriate antibiotic.</li> <li>In cases of PPROM, administer the first dose of erythromycin, 250 mg orally.</li> <li>If possible, administer oxygen.</li> </ol>
				<p><b>One of the following signs:</b></p> <ul style="list-style-type: none"> <li>Mother Rh negative</li> <li>Positive VDRL, HIV, or hepatitis B test</li> </ul>	<b>HIGH-RISK DELIVERY</b> <ol style="list-style-type: none"> <li><b>Refer for consultation by specialist.</b></li> <li>If VDRL test is positive, begin benzathine penicillin treatment.</li> <li>Provide HIV/AIDS-STI counseling</li> <li>Decide on reconsultation.</li> <li>Provide counseling on nutrition, pregnancy care, oral hygiene, and breastfeeding.</li> <li>Teach warning signs.</li> </ol>	<b>LOW-RISK DELIVERY</b> <ul style="list-style-type: none"> <li>Indicate that the patient should walk freely during labor, and provide for oral rehydration.</li> <li>Conduct normal delivery with active birth and late clamping of the cord.</li> <li>Counsel on warning signs during puerperium.</li> <li>Provide HIV/AIDS counseling.</li> </ul>

### Determine:

- Hb, Ht, VDRL test, HIV, hepatitis B
- Blood typing, Coombs test

ASK ALL MOTHERS WHETHER THEY HAVE THEIR PERINATAL CARD

## BMI: Body Mass Index

Height (m)	Weight in kg										93	96	99	101
	45	48	51	54	57	60	63	66	69	72				
1.50	20	21	23	24	25	27	28	29	31	32	33	35	36	37
1.53	19	21	22	23	24	26	27	28	29	31	32	33	35	36
1.56	18	20	21	22	23	25	26	27	28	30	31	32	33	35
1.59	18	19	20	21	23	24	25	26	27	28	30	31	32	33
1.62	17	18	19	21	22	23	24	25	26	27	29	30	31	32
1.65	17	18	19	20	21	22	23	24	25	26	28	29	31	32
1.68	16	17	18	19	20	21	22	23	24	26	27	28	29	30
1.71	15	16	17	18	19	21	22	23	24	25	26	27	28	29
1.74	15	16	17	18	19	20	21	22	23	24	25	26	27	28
1.77	14	15	16	17	18	19	20	21	22	23	24	25	26	27
1.80	14	15	16	17	18	19	19	20	21	22	23	24	25	26
1.83	13	14	15	16	17	18	19	20	21	21	22	23	24	25
1.86	13	14	15	16	17	18	19	20	21	22	23	24	25	26
1.89	13	13	14	15	16	17	18	18	19	20	21	22	23	24
1.92	12	13	14	15	15	16	17	18	19	20	20	21	22	23
1.95	12	13	13	14	15	16	17	17	18	19	20	21	22	23
1.98	11	12	13	14	15	15	16	17	18	18	19	20	21	22

BMI = WEIGHT/HEIGHT<sup>2</sup>

Underweight:

Normal:

Overweight:

Obese:

BMI < 18.5

BMI of 18.5–24.9

BMI of 25.0–29.9

BMI ≥ 30.0

Source: World Health Organization - WHO 1998





**Pan American  
Health  
Organization**



Regional Office of the  
World Health Organization

525 Twenty-third Street, N.W.  
Washington, D.C. 20037

[www.paho.org](http://www.paho.org)



**Texas Children's Hospital®**



**USAID**  
FROM THE AMERICAN PEOPLE

ISBN: 978-92-75-11630-2



9 789275 116302