# WHO recommendations Drug treatment for severe hypertension in pregnancy





# WHO recommendations **Drug treatment for severe hypertension in pregnancy**



WHO recommendations: drug treatment for severe hypertension in pregnancy

ISBN 978-92-4-155043-7

#### © World Health Organization 2018

Some rights reserved. This work is available under the Creative Commons Attribution-NonCommercial-ShareAlike 3.0 IGO licence (CC BY-NC-SA 3.0 IGO; <a href="https://creativecommons.org/licenses/by-nc-sa/3.0/igo">https://creativecommons.org/licenses/by-nc-sa/3.0/igo</a>).

Under the terms of this licence, you may copy, redistribute and adapt the work for non-commercial purposes, provided the work is appropriately cited, as indicated below. In any use of this work, there should be no suggestion that WHO endorses any specific organization, products or services. The use of the WHO logo is not permitted. If you adapt the work, then you must license your work under the same or equivalent Creative Commons licence. If you create a translation of this work, you should add the following disclaimer along with the suggested citation: "This translation was not created by the World Health Organization (WHO). WHO is not responsible for the content or accuracy of this translation. The original English edition shall be the binding and authentic edition".

Any mediation relating to disputes arising under the licence shall be conducted in accordance with the mediation rules of the World Intellectual Property Organization.

**Suggested citation.** WHO recommendations: drug treatment for severe hypertension in pregnancy. Geneva: World Health Organization; 2018. Licence: <u>CC BY-NC-SA 3.0 IGO</u>.

Cataloguing-in-Publication (CIP) data. CIP data are available at http://apps.who.int/iris.

**Sales, rights and licensing.** To purchase WHO publications, see <a href="http://apps.who.int/bookorders">http://apps.who.int/bookorders</a>. To submit requests for commercial use and queries on rights and licensing, see <a href="http://www.who.int/about/licensing">http://www.who.int/about/licensing</a>.

**Third-party materials.** If you wish to reuse material from this work that is attributed to a third party, such as tables, figures or images, it is your responsibility to determine whether permission is needed for that reuse and to obtain permission from the copyright holder. The risk of claims resulting from infringement of any third-party-owned component in the work rests solely with the user.

**General disclaimers.** 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 WHO concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted and dashed lines on maps represent approximate border lines for which there may not yet be full agreement.

The mention of specific companies or of certain manufacturers' products does not imply that they are endorsed or recommended by WHO 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 WHO 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 WHO be liable for damages arising from its use.

## Acknowledgements

The Department of Reproductive Health and Research of the World Health Organization (WHO) gratefully acknowledges the contributions of many individuals and organizations to the updating of these recommendations. Work on this update was coordinated by Olufemi Oladapo, Joshua Vogel and A. Metin Gülmezoglu of the WHO Department of Reproductive Health and Research.

WHO extends its sincere thanks to Edgardo Abalos, Ebun Adejuyigbe, Shabina Ariff, Jemima Dennis-Antwi, Luz Maria De-Regil, Christine East, Lynn Freedman, Pisake Lumbiganon, Anita Maepioh, James Neilson, Hiromi Obara, Rachel Plachcinski, Zahida Qureshi, Kathleen Rasmussen, Niveen Abu Rmeileh and Eleni Tsigas who served as members of the Guideline Development Group (GDG), and to Zahida Qureshi (Chair) and Jim Neilson (Vice-Chair) for leading the meeting. We also thank José Guilherme Cecatti, Sylvia Deganus, M Jeeva Sankar, Hayfaa Wahabi, Jack Moodley, Jane Sandall, Ola Shaker and Nguyen Xuan Hoi who were members of the External Review Group. WHO also gratefully acknowledges the contribution of the members of the Executive Guideline Steering Group.

Anna Cuthbert, Leanne Jones, Frances Kellie and Myfanwy Williams reviewed the scientific evidence, prepared the Grading of Recommendations, Assessment, Development, and Evaluation (GRADE) tables and drafted the narrative summary of evidence. Joshua Vogel and Olufemi Oladapo revised the narrative summaries and double-checked the corresponding GRADE tables and prepared the Evidence-to-Decision frameworks. Joshua Vogel, Olufemi Oladapo, A. Metin Gülmezoglu, Ana Pilar Betrán, Özge Tuncalp and Mercedes Bonet commented on the draft document before it was reviewed by participants at the WHO Guideline Development Group meeting. The External Review Group peer reviewed the final document.

We acknowledge the various organizations that were represented by observers, including Deborah Armbruster and Mary-Ellen Stanton (United States Agency for International Development), Kathleen Hill (Maternal and Child Survival Program/ Jhpiego), Jerker Liljestrand (Bill & Melinda Gates Foundation), Lesley Page (International Confederation of Midwives), Gerard Visser (International Federation of Gynaecology and Obstetrics) and Charlotte Warren (Ending Eclampsia Project, Population Council). We also appreciate the contributions of WHO Regional Office staff – Nino Berdzuli, Bremen De Mucio, Chandani Anoma Jayathilaka, Ramez Khairi Mahaini, Léopold Ouedraogo and Howard Sobel.

The United States Agency for International Development and the Department of Reproductive Health and Research provided financial support for this work. The views of the funding bodies have not influenced the content of these recommendations.

### **Abbreviations**

BMGF Bill & Melinda Gates Foundation

CI Confidence interval
CS Caesarean section
DOI Declaration of Interest

FHR Fetal heart rate

FIGO International Federation of Gynaecology and Obstetrics
FWC Family, Women's and Children's Health (a WHO cluster)

GDG Guideline Development Group
GRC Guideline Review Committee

GRADE Grading of Recommendations, Assessment, Development, and Evaluation

GREAT Guideline development, Research priorities, Evidence synthesis, Applicability of

evidence, Transfer of knowledge (a WHO project)

GSG Executive Guideline Steering Group

HELLP Haemolysis, elevated liver enzymes, low platelet

ICM International Confederation of Midwives

LMIC Low and middle-income country

MCA [WHO Department of] Maternal, Newborn, Child and Adolescent Health

MCSP Maternal and Child Survival Programme

MPA Maternal and Perinatal Health and Preventing Unsafe Abortion (a team in

WHO's Department of Reproductive Health and Research)

MPH Maternal and perinatal health

NNT Number needed to treat

PICO Population (P), intervention (I), comparison (C), outcome (O)
RHR [WHO Department of] Reproductive Health and Research

RR Relative risk

SDG Sustainable Development Goals

UN United Nations

UNFPA United Nations Population Fund

USAID United States Agency for International Development

WHO World Health Organization



# **Contents**

Acknowledgements	III
Abbreviations	iv
Executive Summary	1
1. Background	3
2. Methods	4
3. Recommendations and supporting evidence	9
4. Dissemination and implementation of the recommendations	10
5. Research implications	11
6. Applicability issues	11
7. Updating the recommendations	11
References	12
Annex 1. External experts and WHO staff involved in the preparation of the guideline	13
Annex 2. Priority outcomes for decision-making	17
Annex 3. Summary and management of declared interests from GDG members	18
Annex 4. Evidence-to-decision framework	20
Annex 5. GRADE Tables	38

## **Executive Summary**

#### Introduction

Hypertensive disorders of pregnancy are an important cause of severe morbidity, long-term disability and death among both pregnant women and their babies, and account for approximately 14% of all maternal deaths worldwide. Improving care for women around the time of childbirth is a necessary step towards achievement of the health targets of the Sustainable Development Goals (SDGs). Efforts to prevent and reduce morbidity and mortality during pregnancy and childbirth could also help address the profound inequities in maternal and perinatal health globally. To achieve these goals, healthcare providers, health managers, policy makers and other stakeholders need up-to-date and evidence-based recommendations to inform clinical policies and practices.

In 2017, the Executive Guideline Steering Group (GSG) on WHO maternal and perinatal health recommendations prioritized the updating of the existing WHO recommendations on antihypertensive drugs for severe hypertension in pregnancy in response to important new evidence on these interventions. These recommendations are a revalidation of the previous recommendations issued in 2011 in the WHO recommendations on prevention and treatment of pre-eclampsia and eclampsia.

#### **Target audience**

The primary audience of these recommendations includes healthcare providers who are responsible for developing national and local health protocols (particularly those related to hypertensive disorders of pregnancy) and those directly providing care to pregnant women and their newborns, including midwives, nurses, general medical practitioners, obstetricians, managers of maternal and child health programmes, and relevant staff in ministries of health, in all settings.

### **Guideline development methods**

The updating of these recommendations was guided by standardized operating procedures in accordance with the process described in the *WHO handbook for guideline development.* The recommendations were initially developed using this process, namely:

- (i) identification of the priority question and critical outcomes;
- (ii) retrieval of evidence;
- (iii) assessment and synthesis of evidence;
- (iv) formulation of the recommendations; and
- (v) planning for the dissemination, implementation, impact evaluation and updating of the recommendations.

The scientific evidence supporting the recommendations was synthesized using the Grading of Recommendations, Assessment, Development, and Evaluation (GRADE) approach. The systematic review was used to prepare evidence profiles for the prioritized question. WHO convened an online meeting on 2 May 2018 where an international group of experts – the Guideline Development Group (GDG) – reviewed and approved the recommendations.

#### The recommendations

The GDG reviewed the balance between the desirable and undesirable effects and the overall quality of supporting evidence, values and preferences of stakeholders, resource requirements and cost-effectiveness, acceptability, feasibility and equity. The GDG revalidated the WHO recommendations published in 2011 with minor revisions to the remarks and implementation considerations.

To ensure that the recommendations are correctly understood and applied in practice, guideline users should refer to the remarks, as well as to the evidence summary, if there is any doubt as to the basis of the recommendations and how best to implement them.

#### Table 1: WHO recommendations: drug treatment for severe hypertension in pregnancy.

1. Women with severe hypertension during pregnancy should receive treatment with antihypertensive drugs (strong recommendation, very low certainty evidence)

#### **Remarks**

- The guideline development group considered that there is no clinical uncertainty over whether treatment of severe hypertension during pregnancy is beneficial. This recommendation was made based on expert opinion; the group considered that most maternal deaths related to hypertensive disorders are associated with complications of uncontrolled severe high blood pressure. Based on that, the group agreed that antihypertensive treatment should be recommended in all cases of severe hypertension.
- 2. The choice and route of administration of an antihypertensive drug for severe hypertension during pregnancy, in preference to others, should be based primarily on the prescribing clinician's experience with that particular drug, its cost and local availability (conditional recommendation, very low certainty evidence)

#### **Remarks**

- In terms of the choice and route of administration of an antihypertensive drug for severe hypertension during pregnancy, the guideline development group noted that not only is the evidence base for this recommendation limited, but also some antihypertensive drugs may not be feasible options in many settings.
- The group acknowledged that hydralazine, alpha methyldopa, beta blockers (including labetalol) and nifedipine have been extensively used, and therefore, these agents would seem to be reasonable choices until further evidence becomes available.
- The group noted that there was no evidence to suggest that nifedipine interacts adversely with
  magnesium sulfate. In addition, the group considered that the use of angiotensin- converting enzyme
  inhibitors, angiotensin receptor blockers and sodium nitroprusside should be avoided due to safety
  concerns.

预览已结束,完整报告链接和二维码如下:

https://www.yunbaogao.cn/report/index/report?reportId=5 25449

