



Universiteit  
Leiden  
The Netherlands

## Reply to Watchko and Maisels: exchange transfusion in Rh haemolytic disease

Ree, I.M.C.; Besuden, C.F.J.; Wintjens, V.E.H.J.; Verweij, J.J.T.; Oepkes, D.; Haas, M. de; Lopriore, E.

### Citation

Ree, I. M. C., Besuden, C. F. J., Wintjens, V. E. H. J., Verweij, J. J. T., Oepkes, D., Haas, M. de, & Lopriore, E. (2021). Reply to Watchko and Maisels: exchange transfusion in Rh haemolytic disease. *Vox Sanguinis*, 117(1), 147-148. doi:10.1111/vox.13163

Version: Publisher's Version

License: [Creative Commons CC BY-NC-ND 4.0 license](https://creativecommons.org/licenses/by-nc-nd/4.0/)

Downloaded from: <https://hdl.handle.net/1887/3270930>

**Note:** To cite this publication please use the final published version (if applicable).

## Reply to Watchko and Maisels: Exchange transfusion in Rh haemolytic disease

We thank Dr. Watchko and Dr. Maisels for their knowledgeable comments on our study and would like to address their questions.

One concern was the use of a higher bilirubin threshold for phototherapy and exchange transfusion (ET) at our centre for infants with haemolytic disease of the foetus and newborn (HDFN). The relationship between haemolysis and high bilirubin levels is clear, but it is unclear whether infants with HDFN have a higher risk of bilirubin neurotoxicity compared to infants without HDFN in case of similar levels of serum bilirubin [1]. We have no reason to assume that HDFN alters the blood–brain barrier and therefore do not assume that the studied, term infants have a higher risk of bilirubin encephalopathy. Moreover, the maximum bilirubin level at birth remained stable for group II (2005–2015) and III (2015–2020), with maximum values of 257 (standard deviation [SD] of 89) and 262  $\mu\text{mol/L}$  (SD of 80) after birth. Local policy also indicates the near-immediate start (within 15 min after birth) of intensive phototherapy for infants with HDFN regardless of the first measured bilirubin after birth or other risk factors and will therefore not delay treatment. Other risk factors include prematurity, asphyxia, suspected infection/sepsis and low albumin levels.

With regard to phototherapy, technological advancements of the used (LED) lamps very likely also contributed to more effective treatment of hyperbilirubinaemia and decreased use of ET. In our study, the median duration of phototherapy per infant remained stable over the years, with a median of 4 (interquartile range [IQR] 3–5), 5 (IQR 3–6) and 5 (IQR 4–6) days in our three time cohorts. The timing of the start of phototherapy (within 15 min after birth) has not changed over the years.

The relationship between intrauterine transfusion (IUT) and ET(s) was previously reported by our study group [2, 3]. Infants treated with more IUTs required fewer ET(s); the rate of ET dropped from 39% of infants treated with one IUT to 25% after two IUTs and further declined to 8% for infants treated with five IUTs (infants born 2005–2018) [2]. The decline in ET rate in this study does not show a similar trend as the IUT rate has declined in the three time cohorts in our study from a median of three IUTs (IQR 2–4) in group I to two (IQR 2–4) and two (IQR 1–3) in group II and III.

The standard blood product in the Netherlands for neonatal ET has a haematocrit of around 0.50; no additional albumin is transfused prior to or during the procedure. As stated, the product

consists of a two donor combination of washed red blood cells and adult plasma.

We hope to have clarified the raised issues and welcome all further thoughts on our study.

### ACKNOWLEDGEMENTS

The authors received no specific funding for this work.

### CONFLICT OF INTEREST

There are no conflicts of interest to report.

Isabelle M. C. Ree<sup>1,2</sup> 

Carolin F. J. Besuden<sup>1</sup>

Vivianne E. H. J. Wintjens<sup>1</sup>

Joanne (E.) J. T. Verweij<sup>3</sup>

Dick Oepkes<sup>3</sup>

Masja de Haas<sup>2,4,5</sup>

Enrico Lopriore<sup>1</sup>

<sup>1</sup>Department of Paediatrics, Division of Neonatology, Leiden University Medical Centre, Leiden, The Netherlands

<sup>2</sup>Centre for Clinical Transfusion Research, Sanquin, Leiden, The Netherlands

<sup>3</sup>Department of Obstetrics, Division of Foetal Medicine, Leiden University Medical Centre, Leiden, The Netherlands

<sup>4</sup>Department of Immunohaematology and Blood Transfusion, Leiden University Medical Centre, Leiden, The Netherlands

<sup>5</sup>Department of Immunohaematology Diagnostics, Sanquin, Leiden, The Netherlands

### Correspondence

Isabelle Ree, Department of Paediatrics, Leiden University Medical Centre, J6-S, Albinusdreef 2, 2333 ZA, Leiden, The Netherlands.

Email: i.m.c.ree@lumc.nl

Alsabelle M.C. Ree and Carolin F.J. Besuden contributed equally to this study.

## ORCID

Isabelle M. C. Ree  <https://orcid.org/0000-0003-3608-3509>

## REFERENCES

1. Kaplan M, Bromiker R, Hammerman C. Hyperbilirubinemia, hemolysis, and increased bilirubin neurotoxicity. *Semin Perinatol.* 2014;38:429–37.
2. Ree IMC, Lopriore E, Zwiers C, Böhringer S, Janssen MWM, Oepkes D, et al. Suppression of compensatory erythropoiesis in hemolytic disease of the fetus and newborn due to intrauterine transfusions. *Am J Obstet Gynecol.* 2020;223:119.e1–119.e10.
3. Ree IMC, de Haas M, Milderburg RA, Zwiers C, Oepkes D, Bom JG, et al. Predicting anaemia and transfusion dependency in severe alloimmune haemolytic disease of the fetus and newborn in the first 3 months after birth. *Br J Hematol.* 2019;186:565–7.