

JOHNS HOPKINS ALL CHILDREN'S HOSPITAL

Screening Examination for Retinopathy of Prematurity Clinical Pathway



JOHNS HOPKINS
All Children's Hospital



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Owners: Sandra Brooks MD

This pathway is intended as a guide for physicians, physician assistants, nurse practitioners and other healthcare providers. It should be adapted to the care of specific patient based on the patient's individualized circumstances and the practitioner's professional judgment.

Johns Hopkins All Children's Hospital

Screening Examination for Retinopathy of Prematurity Clinical Pathway

SCOPE

This Clinical Practice Guideline (CPG) applies to:

All Children's Hospital, Inc., and

All Children's Health System, Inc.

o West Coast Neonatology, Inc.

Guideline Panel

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Executive Summary

Retinopathy of prematurity (ROP) is a serious disorder of the developing retinal blood vessels in the low birth weight preterm infant and is a leading cause of childhood preventable blindness.

The pathophysiology of ROP is characterized by two phases. Phase I ROP is due to vaso-obliteration beginning immediately after birth secondary to a marked decrease in vascular endothelial growth factor (VEGF) and insulin-like growth factor-1 (IGF-1). Phase II begins around 33 weeks' postmenstrual age at which time VEGF levels increase, especially if there is retinal hypoxia with increasing retinal metabolism and demand for oxygen leading to abnormal vasoproliferation.

- A. Preterm infants with low birth weight neonates are considered high risk for ROP
- B. ROP progression is sequential in nature, it is imperative that at-risk infants receive their initial retinal exam at the appropriate post-menstrual age and have timely follow-up exams. This will ensure detection of early changes of ROP and allow indicated intervention before irreversible retinal damage occurs.
- C. Timely initial screening and follow-up examinations and prompt treatment of at-risk neonates has proven benefits of reducing unfavorable structural and visual outcomes.

Published Data and Levels of Evidence

- A. Preterm and low birth weight infants are born with an immature retina and retinal vasculature. Lower gestational age at birth and lower birth weight directly correlate with increased immaturity of the retina and retinal vasculature
- B. Neonates born with gestational age (GA) at birth of 30 weeks and less or birth weight (BW) of 1500g or less have immature retinal tissue at high risk of developing in a pathologic manner and progressing to ROP, retinal detachment and blindness
- C. Hyperoxia and oxygen saturations consistently outside the target range of 90%-95% at less than 36 weeks GA are known risk factors
- D. Several large multi-center trials have demonstrated a significantly decreased incidence of poor visual outcomes with early ROP detection and treatment
- E. The 2018 revised Consensus Statement of American Academy of Pediatrics, American Academy of Ophthalmology, American Association for Pediatric Ophthalmology and Strabismus, and American Association of Certified Orthoptists has proposed the schedule for the timing of retinal examinations in this patient population and is intended to permit early detection of ROP before it progresses to permanent disease while minimizing the number of potentially traumatic examinations.
- F. A small number of case reports have raised the possibility of ROP occurrence in newborns of older GA and larger BW. The cases presented were very few and had a common denominator of fetal distress and stormy neonatal course

Clinical Practice Guideline

- A. All preterm infants with gestational age (GA) at birth of 30 weeks and less or birth weight (BW) of 1500g and less are considered at risk for ROP and should undergo retinal screening examinations. (*Evidence level 1A; Strong recommendation, high quality evidence*)
- B. The onset of serious ROP correlates better with postmenstrual age (gestational age at birth plus chronologic age) than postnatal age. That is the more preterm an infant is at birth, the longer the time to develop serious ROP. Thus, the initiation of acute-phase ROP screening should be based on the infant's POSTMENSTRUAL AGE, and should be performed between 31 and 34 weeks. (*Evidence level 1A; Strong recommendation, high quality evidence*)

ROP Screening Initiation

	Age at initial examination, weeks	Age at initial examination, weeks
GA at Birth (weeks)	<i>Postmenstrual</i>	<i>Chronologic</i>
22	31	9
23	31	8
24	31	7
25	31	6
26	31	5
27	31	4
28	32	4
29	33	4
30	34	4
Older GA with high risk (consider based on severity of comorbidities)		4

- C. The proposed schedule provides for the detection of ROP that is potentially damaging to the retina with 99% confidence, usually before any required treatment.
- D. For infants born at 22 and 23 weeks, the schedule should be considered tentative rather than evidence-based because of the small number of survivors in these postmenstrual-age categories.
- E. For those infants ≤ 24 weeks GA at birth, may consider an earlier timing at 6 weeks chronological age, based on severity of comorbidities, likelihood of a meaningful examination and tolerance to an eye exam. Those infants are at higher risk for developing a posterior ROP, a usually more aggressive form of ROP. (*Evidence level 1C; Strong recommendation, moderate quality evidence*)
- F. It is important to note that despite appropriate timing of examinations and treatment, a small number of infants at risk progress to poor outcomes.
- G. It is recommended to minimize the discomfort and systemic effects of an eye exam by considering the use of pacifiers, oral sucrose when applicable and so forth.
- H. Using multiple dilating drops SHOULD BE AVOIDED because poor pupillary dilation can occur in advanced ROP, and administering multiple doses of dilating drops can adversely affect the systemic status of the infant.
- I. Additional infants with GA at birth between 31 and 34 weeks or BW between 1500 Grams and 2000 Grams with a prolonged unstable, critically ill clinical course as determined by the treating team / Neonatologist may be considered high risk for ROP and should undergo screening ROP exam. (*Evidence level 2C; Weak recommendation, low quality evidence*)
- J. For infants deemed clinically unfit for initial retinal exam (as determined by the neonatologist), the exam may be rescheduled within the next week (7 days). However, documentation of the instability of the patient and the postponing of the eye exam is paramount and should be entered daily until the next eye exam is successfully performed.

Follow-Up Exams during the NICU stay

- a) Timing of NICU follow-up exams will be recommended by the examining ophthalmologist in accordance with retinal findings classified according to the “International classification of retinopathy of prematurity revisited” and the 2018 American Academy of Pediatrics Policy statement. (see figure)

Follow-Up Exams after discharge home

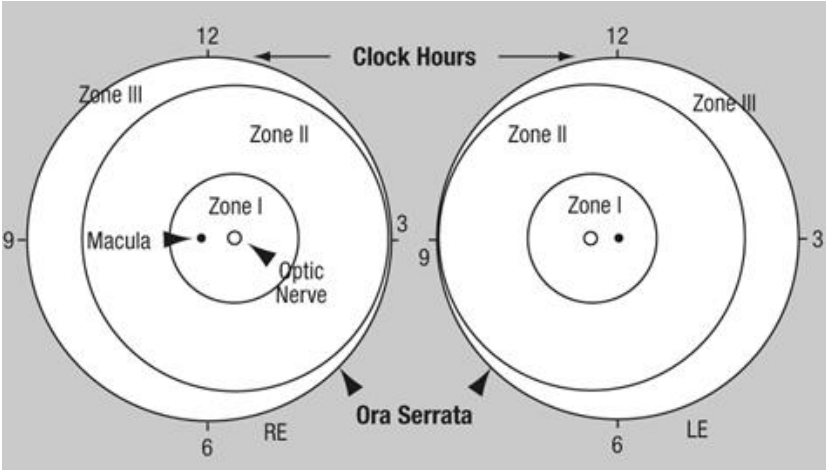
- a) Infants discharged from the NICU who require on-going retinal exams will continue to follow-up with ophthalmology on an outpatient basis. Prior to discharge, the schedule will be recommended by the examining ophthalmologist according to the retinal exam findings. Parents and Primary care providers for these infants must be made aware of, and agree to strictly follow, this outpatient schedule.

- b) All infants who developed ROP (regardless of need for treatment), are at high risk of other visual disorders including but not limited to amblyopia, strabismus, cataract, and high refractive errors. Pediatricians / outpatient physicians who care for these infants should be made aware of these potential risks. Additionally, despite resolution of ROP, these infants should have follow-up with ophthalmology within 4-6 months of discharge from the NICU.

Glossary

BW - Birth Weight
GA - Gestational Age
NICU - Neonatal Intensive Care Unit
PMA - Post Menstrual Age
ROP - Retinopathy of Prematurity

Figure



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References

Atkins D, Best D, Briss PA, Eccles M, Falck-Ytter Y, Flottorp S, Guyatt GH, Harbour RT, Haugh MC, Henry D, Hill S, Jaeschke R, Leng G, Liberti A, Magrini N, Mason J, Middleton P, Mrukowicz J, O'Connell D, Oxman AD, Phillips B, Schünemann HJ, Edejer T, Varonen H, Vist GE, Williams JW Jr, Zaza S; GRADE Working Group. Grading quality of evidence and strength of recommendations. *BMJ*. 2004 Jun 19;328(7454):1490

Ebell MH, Siwek J, Weiss BD, Woolf SH, Susman J, Ewigman B, Bowman M. Strength of recommendation taxonomy (SORT): a patient-centered approach to grading evidence in the medical literature. *Am Fam Physician*. 2004 Feb 1;69(3):548-56..

International Committee for the Classification of Retinopathy of Prematurity. The International Classification of Retinopathy of Prematurity revisited. *Arch Ophthalmol*. 2005 Jul;123(7):991-9. doi: 10.1001/archophth.123.7.991. PMID: 16009843

Early Treatment For Retinopathy Of Prematurity Cooperative Group. Revised indications for the treatment of retinopathy of prematurity: results of the early treatment for retinopathy of prematurity randomized trial. *Arch Ophthalmol*. 2003 Dec;121(12):1684-94.

Early Treatment for Retinopathy of Prematurity Cooperative Group, Good WV, Hardy RJ, Dobson V, Palmer EA, Phelps DL, Tung B, Redford M. Final visual acuity results in the early treatment for retinopathy of prematurity study. *Arch Ophthalmol*. 2010 Jun;128(6):663-71.

Fierson WM; American Academy of Pediatrics Section on Ophthalmology; American Academy of Ophthalmology; American Association for Pediatric Ophthalmology and Strabismus; American Association of Certified Orthoptists. Screening Examination of Premature Infants for Retinopathy of Prematurity. *Pediatrics*. 2018; 142(6):e20183061. *Pediatrics*. 2019 Mar;143(3):e20183810.

Good WV; Early Treatment for Retinopathy of Prematurity Cooperative Group. Final results of the Early Treatment for Retinopathy of Prematurity (ETROP) randomized trial. *Trans Am Ophthalmol Soc*. 2004;102:233-48; discussion 248-50.

Good WV, Hardy RJ, Dobson V, et al; Early Treatment for Retinopathy of Prematurity Cooperative Group. Final visual acuity results in the early treatment for retinopathy of prematurity study. *Arch Ophthalmol*. 2010;128(6):663–671pmid

Hutchinson AK, Saunders RA, O'Neil JW, Lovering A, Wilson ME. Timing of initial screening examinations for retinopathy of prematurity. *Arch Ophthalmol*. 1998 May;116(5):608-12.

International Committee for the Classification of Retinopathy of Prematurity. The International Classification of Retinopathy of Prematurity revisited. *Arch Ophthalmol*. 2005 Jul;123(7):991-9. doi: 10.1001/archophth.123.7.991. PMID: 16009843

Padhi, T., Rath, S., Jalali, S. *et al*. Larger and near-term baby retinopathy: a rare case series. *Eye* **29**, 286–289 (2015)

Palmer, E. A., Flynn, J. T., Hardy, R. J., Phelps, D. L., Phillips, C. L., Schaffer, D. B., & Tung, B. (1991). Incidence and early course of retinopathy of prematurity. the cryotherapy for retinopathy of prematurity cooperative group. *Ophthalmology*, *98*(11), 1628-1640.

Palmer EA. Results of U.S. randomized clinical trial of cryotherapy for ROP (CRYO-ROP). *Doc Ophthalmol.* 1990 Mar;74(3):245-51

Palmer EA, Hardy RJ, Dobson V, Phelps DL, Quinn GE, Summers CG, Krom CP, Tung B; Cryotherapy for Retinopathy of Prematurity Cooperative Group. 15-year outcomes following threshold retinopathy of prematurity: final results from the multicenter trial of cryotherapy for retinopathy of prematurity. *Arch Ophthalmol.* 2005 Mar;123(3):311-8.v

Reynolds JD, Dobson V, Quinn GE, Fielder AR, Palmer EA, Saunders RA, Hardy RJ, Phelps DL, Baker JD, Trese MT, Schaffer D, Tung B; CRYO-ROP and LIGHT-ROP Cooperative Study Groups. Evidence-based screening criteria for retinopathy of prematurity: natural history data from the CRYO-ROP and LIGHT-ROP studies. *Arch Ophthalmol.* 2002 Nov;120(11):1470-6.

Saugstad OD, Aune D: Optimal oxygenation of extremely low birth weight infants: a meta-analysis and systematic review of the oxygen saturation target studies. *Neonatology* 2014;105:55-63

Schmidt B, Whyte RK, Asztalos EV, et al; Canadian Oxygen Trial (COT) Group: Effects of targeting higher vs lower arterial oxygen saturations on death or disability in extremely preterm infants: a randomized clinical trial. *JAMA* 2013;309:2111-2120

Shah PK, Narendran V, Kalpana N. Aggressive posterior retinopathy of prematurity in large preterm babies in South India. *Arch Dis Child Fetal Neonatal Ed.* 2012 Sep;97(5):F371-5

SUPPORT Study Group of the Eunice Kennedy Shriver NICHD Neonatal Research Network, Carlo WA, Finer NN, Walsh MC, et al: Target ranges of oxygen saturation in extremely preterm infants. *N Engl J Med* 2010;362:1959-1969.

The BOOST II United Kingdom, Australia, and New Zealand Collaborative Groups: Oxygen saturation and outcomes in preterm infants. *N Engl J Med* 2013;368:2094-2104.

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Screening Examination for Retinopathy of Prematurity
Clinical Pathway

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Disclaimer

Clinical Pathways are intended to assist physicians, physician assistants, nurse practitioners and other health care providers in clinical decision-making by describing a range of generally acceptable approaches for the diagnosis, management, or prevention of specific diseases or conditions. The ultimate judgment regarding care of a particular patient must be made by the physician in light of the individual circumstances presented by the patient.

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