

1 **ROP prevention, screening and treatment programmes - progress update from Latin America**

2

3 Juan Carlos Silva, MD, MPH
4 Pan-American Health Organization-PAHO,
5 Calle 66 # 11 – 50,
6 Bogotá,
7 Colombia

8

9 Andrea Zin, MD, MSc, PhD
10 Clinical Research Unit,
11 Fernandes Figueira Institute,
12 Av Rui Barbosa 716,
13 Flamengo, Rio de Janeiro 22250-020,
14 Brazil

15

16 Clare Gilbert MB ChB., FRCOphth., MD. MSc
17 Department of Clinical Research,
18 London School of Hygiene & Tropical Medicine,
19 Keppel Street
20 London WC1E 7HT
21 United Kingdom

22

23 **Corresponding author**

24 Juan Carlos Silva
25 Pan-American Health Organization-PAHO,
26 Calle 66 # 11 – 50,
27 Bogotá,
28 Colombia
29 Email: juasilva@paho.org

30

1 **Abstract**

2 Retinopathy of prematurity (ROP) is the main cause of blindness and visual impairment in Latin
3 America and prevention, detection and treatment have been a priority in the Region in the last two
4 decades. There is progress in the number of eligible babies screened and treated in at least half of the
5 countries with strong regulations on ROP, but screening is not yet available in all provinces in most.
6 More effort is needed to increase national protocols and legislation, strengthening of services and
7 human resources to cover all national provinces, telemedicine might be a promising strategy.

8

9 **Introduction**

10 Retinopathy of prematurity (ROP) is a major cause of blindness and visual impairment in children in
11 Latin America being responsible for 16.6% of blindness in Chile, 11.8% in Sao Paulo, Brazil and
12 34.7% in Guadalajara, Mexico.¹⁻³ Studies from a number of countries of preterm infants with a birth
13 weight of < 1,500 g, found the incidence of any stage ROP varied from 10% to 71%, and the
14 incidence of treatable ROP varied from 1.2% to 19%.⁴

15

16 An earlier study in Latin America and the Caribbean showed that a higher proportion of eligible
17 babies were screened in countries with government policies mandating and financing screening,
18 which had clinical guidelines and where ROP was included in national health information systems.⁵ In
19 a worldwide survey of ROP, nine Latin American countries responded that they had national ROP
20 protocols,⁶ but information was not provided on the availability of ophthalmologists to screen, nor
21 whether countries had a national ROP screening programme.

22 **Methods**

23 This paper summarizes the recent published data on the situation of ROP legislation, guidelines and
24 programme development in Latin America. To obtain additional information, the Pan American
25 Health Organization (PAHO) distributed a survey to national members of the Pan American Society
26 of Retinopathy of Prematurity in all 19 countries in the region in 2018, to obtain information on the
27 number of ophthalmologists performing diagnostic examination, the number of NICUs implementing
28 ROP programmes at subnational level and the use of telemedicine in the screening.

1 Regional Response

2 The control of blindness from ROP has been a regional priority since the nineties when ROP was
3 identified as an important cause blindness in children,⁷ and was included in PAHO Action Plans for
4 the Prevention of Blindness in 2009-2013⁸ and 2014-2019.⁹ Ministers of Health across the region
5 approved these plans which are being implemented with support from PAHO, the International
6 Agency for the Prevention of Blindness, international non-government organizations, worldwide
7 experts and the Pan American Society of Retinopathy of Prematurity. In 2017, PAHO worked with a
8 multi-disciplinary, international group to develop evidence-based guidelines for ROP to serve as a
9 regional standard.¹⁰

10 National response

11 Normative

12 In the earlier study in ten Latin America countries, a higher proportion of eligible preterm newborns
13 were being screened and treated in countries with (i) policies on financing mechanisms for screening
14 and treatment, (ii) national ROP guidelines (iii) legislation mandating eye screening and (iv)
15 management systems for data collection and analysis.⁵ In Argentina, crucial factors which led to
16 scaling up services for ROP included a dedicated group of professionals, the Ministry of Health, a
17 national ROP committee, international cooperation for norms development and programme
18 implementation, and external funding for a limited period of time.¹¹ Several countries in the region
19 have national clinical guidelines which were developed using the GRADE methodology, in Argentina
20 in 2016,¹² Mexico Ministry of Health in 2010¹³ and Social Security in 2015¹⁴. Chile has an expert
21 review guidelines,¹⁵ whilst guidelines from Costa Rica, Cuba, El Salvador and Nicaragua exist but are
22 not publicly available on the internet. The criteria for screening in these guidelines vary from <1,500
23 to <2,000 g, and from <32 to <34 weeks gestational age, and recommend screening preterm infants at
24 any birth weight who have risk factors such as sepsis and transfusion of blood products.

25 Plans and Programmes

26 Nine of the nineteen countries collected and submitted the information requested in this survey
27 whereas Chile and Argentina routinely collect data. In Chile, the programme is established in all

1 regions, and in eight provinces screening is by imaging and telemedicine with ophthalmologist
 2 readers, which has proved to be effective.¹⁶ In Argentina in 2018, 114 neonatal intensive care units
 3 (NICUs), comprising most of the government sector, and 119 ophthalmologists have been
 4 implementing screening and treatment in all provinces since 2016.¹⁷
 5 Over 60% of provinces/regions in Bolivia, Costa Rica and Nicaragua have at least one NICU doing
 6 ROP screening, and in about half of provinces/regions in Paraguay and Peru. El Salvador and
 7 Guatemala have less than 50% of provinces/regions implementing a programme (Table 1).

8

9 **Table 1. Policies mandating screening and provinces/regions with at least one hospital doing**
 10 **screening**

Country	Policies mandating the exam or covering costs	Number of provinces/regions	Provinces/regions with screening (N, %)
Argentina	Yes	24	24 (100%)
Bolivia	No	9	6 (67%)
Chile	Yes	16	16 (100%)
Costa Rica	Yes	7	5 (71%)
El Salvador	Yes	14	5 (36%)
Guatemala	No	22	9 (41%)
Nicaragua	No	17	12 (71)
Paraguay	No	18	9 (50)
Peru	No	25	13 (52)

11

12 Nine countries had information on the number of NICUs with ROP screening and the number of
 13 ophthalmologists doing screening. The number of preterm babies per ophthalmologist screener varies
 14 and there is estimated to be one ophthalmologist per 108 preterm babies in Argentina and one per 250
 15 preterm babies in Chile (Table 2). Both these countries have programmes which cover all provinces /
 16 regions, and Chile has a highly developed telemedicine program. In Guatemala there are estimated to
 17 be 474 babies per ophthalmologists doing screening, and if each baby is examined 3 times this would
 18 result in a workload of 27 preterm babies per ophthalmologists per week.. This workload would be

1 manageable if each ophthalmologist dedicates most of their time in ROP screening and treatment and
 2 if the babies are in very few large NICUs nearby, but this is rarely the case.

3

4 **Table 2: Number of NICUs with ROP screening, number of ophthalmologists doing screening,**
 5 **number of eligible preterm babies per ophthalmologist doing screening**

Country	NICUs w screening (N)	Ophthalmols screening (N)	Preterm births, 2015 (N)		Access to NICU (80%)	Survive NICU care (80%)	Eligible preterms/ screener (N)
			<37 wks ²	<34 wks ³			
Argentina ¹	114	119	60.900	20.097	16.078	12.862	108
Bolivia	9	10	22.100	7.293	5.834	4.668	467
Chile ¹	24	14	16.600	5.478	4.382	3.506	250
Costa Rica ¹	9	11	9.300	3.069	2.455	1.964	179
El Salvador	7	8	13.400	4.422	3.538	2.830	354
Guatemala	9	16	35.900	11.847	9.478	7.582	474
Nicaragua	14	18	11.100	3.663	2.930	2.344	130
Paraguay ¹	11	28	10.500	3.465	2.772	2.218	79
Peru ¹	31	35	45.700	15.081	12.065	9.652	276

1 Countries using digital imaging with remote reading

2 Preterm births in 2015 www.everywomaneverychild.org/wp.../Country-data-for-WPD-2016-Excel-doc.xlsx

3 33% estimate

6 A tele-education system for ROP was effective in improving the diagnostic accuracy and management
 7 of ROP by ophthalmologists-in-training in Mexico.¹⁸

8 Monitoring

9 The regional ROP guidelines recommend collecting data on the number of preterm newborns who are
 10 eligible for screening, the number examined, the number with ROP of any stage, the number with
 11 indications for treatment and number treated, to construct the following indicators, (1) coverage of
 12 screening, (2) proportion of preterm infants with any stage of ROP, and (3) treatment coverage for
 13 ROP.¹⁰

14 Since 2004 the national programme in Argentina has collected annual data and produces a report, with
 15 the indicators from 98 hospitals in 24 provinces reported in 2016. From 2004 to 2016, the mean birth

1 weight of newborns with any ROP decreased by 200 g and the mean GA reduced by 3 weeks. In
2 addition, the proportion of babies treated for ROP with BW >1500 g or >32 weeks of gestational age
3 was 30% in 2004, which rose to a peak of 40% in 2006 and progressively declined to 17 % (44/ 253)
4 in 2016.¹⁷ These changes almost certainly reflect improvements in neonatal care. Costa Rica has
5 documented risk factors for ROP, and the results of ROP screening and treatment between 2010 and
6 2014,¹⁹ and Colombia uses the national health information system (ASIS) to collect data on the
7 number of live born babies with birth weight <1,999 g with a diagnosis of ROP; in 2014 it was
8 8.3%.²⁰ Some countries have assessed their ROP programme in a selected group of NICUs; in 7 units
9 in Rio de Janeiro, Brazil in 2010 to identify the characteristics of preterm infants at risk of ROP,²¹ in 6
10 units in Lima, Peru in 2012,²² and in 32 units in Mexico in 2015.²³

11 **Comment**

12

13 Latin America has experienced a strong regional direction, advocacy and support from international
14 allies and experts, the Ministries of Health are introducing protocols and regulations and making
15 important financial allocations for the incorporation of ROP services into neonatal care. Almost half
16 of the countries have an inventory on the number of NICUs and ophthalmologists implementing a
17 ROP programme at the subnational level and are using telemedicine in the screening. Argentina and
18 Chile have a national steering committee that plan and implement activities to strengthening services
19 and human resources to cover all national provinces/regions and collect, analyses and uses data. The
20 countries with less development in ROP programs would require developing protocols and regulations
21 and stewardship for services and human resources development at the subnational level. Further
22 research is needed on adherence to national or international guidelines in regions/provinces.

23

24 It is not possible to recommend the ideal number of ophthalmologists who should screen for ROP, as
25 this is influenced not only by the number of preterm infants who survive but also by the distribution
26 of NICUs and of ophthalmologists. Assuming each preterm infant needs an average of three
27 examinations, the workload in terms of the number of preterm infants who need to be screened each
28 week would very high in some countries in the region and more ophthalmologist need to be trained in

1 ROP screening. Much of this shortfall could be addressed by neonatology-led screening using wide-
2 field imaging whereby members of the neonatal team take the images which are either interpreted by
3 the team, which would require rigorous quality assurance systems, or are remotely interpreted by ROP
4 experts.²⁴ Almost real-time image analysis using artificial intelligence is also on the horizon²⁵. These
5 advances in combination have great potential to increase access to high quality, reliable screening.

6

7

8 **Funding:** This study received no specific grant from any funding agency in the public, commercial or
9 not-for-profit sectors.

10

11 **Conflict of interest statement:** The authors have no conflicts of interest to declare.

12

13

14

15

References

1. Gilbert CE, Canovas R, Hagan M, Rao S, Foster A. Causes of childhood blindness: Results from west Africa, south India and Chile. *Eye* (1993) 7, 184-188
2. Haddad MA, Sei M, Sampaio MW, Kara-José N. Causes of visual impairment in children: a study of 3,210 cases. *J Pediatr Ophthalmol Strabismus*. 2007 Jul-Aug;44(4):232-40
3. Zepeda-Romero LC, Barrera-de-Leon JC, Camacho-Choza C, Gonzalez Bernal C, Camarena-Garcia E, Diaz-A; Latorre C et al. Retinopathy of prematurity as a major cause of severe visual impairment and blindness in children in schools for the blind in Guadalajara city, Mexico *Br J Ophthalmol* 2011;95:1502e1505. doi:10.1136/bjophthalmol-2011-300015
4. Carrion JZ, Fortes Filho JB, Tartarella MB, Zin A, Jornada ID Jr. Prevalence of retinopathy of prematurity in Latin America. *Clin Ophthalmol* 2011;5:1687-95.
5. Arnesen L, Durán P, Silva J, Brumana L. A multi-country, cross-sectional observational study of retinopathy of prematurity in Latin America and the Caribbean. *Rev Panam Salud Publica*. 2016 Jun;39(6):322-329
6. Mora JS, Waite C, Gilbert CE, Breidenstein B, Sloper JJ. A worldwide survey of retinopathy of prematurity screening. 2018 Jan;102(1):9-13. doi: 10.1136/bjophthalmol-2017-310709.
7. Gilbert C, Rahi J, Eckstein M, O'Sullivan J, Foster A. Retinopathy of prematurity in middle-income countries. *The Lancet* Volume 350, Issue 9070, 5 July 1997, Pages 12-14
8. Pan American Health Organization. Plan of Action for the Prevention of Blindness and Visual Impairment 2009-2013. Proceedings of the 49th Directing Council of PAHO, 61th Session of the Regional Committee of WHO for the Americas. 28 September-2 October 2009. Washington: PAHO, 2009. [http://www1.paho.org/hq/dmdocuments/2010/CD49-19-e\[1\].pdf](http://www1.paho.org/hq/dmdocuments/2010/CD49-19-e[1].pdf)
9. Pan American Health Organization. Plan of Action for the Prevention of Blindness and Visual Impairment 2014-2019. Proceedings of the 53rd Directing Council of PAHO, 66th Session of the Regional Committee of WHO for the Americas; 29 September-3 October 2014. Washington: PAHO, 2014. https://www.paho.org/hq/index.php?option=com_content&view=article&id=9774:2014-53rd-directing-council&Itemid=40507&lang=en#official
10. Pan American Health Organization. Guía de práctica clínica para el manejo de la retinopatía de la prematuridad. Washington D.C: USA, 2017 <http://iris.paho.org/xmlui/handle/123456789/34948>
11. Hariharan L, Gilbert CE, Quinn GE, Barg FK, Lomuto C, Quiroga A et al. Reducing blindness from retinopathy of prematurity (ROP) in Argentina through collaboration, advocacy and policy implementation. *Health Policy and Management*. 2018 Jun 1;33(5):654-665. doi: 10.1093/heapol/czy004
12. Ministerio de Salud de Argentina & Grupo ROP. Guía de Práctica Clínica para la prevención, diagnóstico y tratamiento de la Retinopatía del Prematuro (ROP) Buenos Aires, Argentina: Ministerio de Salud; 2016. <http://www.msal.gob.ar/images/stories/bes/graficos/0000000723cnt-guia-rop-2016.pdf>
13. Secretaria de Salud de México. Detección, diagnóstico y tratamiento de retinopatía del prematuro, México DF: Secretaria de Salud 2010. <http://sgm.issste.gob.mx/medica/medicadocumentacion/guiasautorizadas/ofthalmolog%C3%ACa/I>

1 MSS-281-10-etinopat%C3%ADa%20del%20prematuro/IMSS-281-
2 10GER%20Retinopat%C3%ADa%20del%20Prematuro.pdf
3

- 4 14. Instituto Mexicano del Seguro Social. Detección, diagnóstico y tratamiento de retinopatía del
5 prematuro en el segundo y tercer nivel de atención. México: Instituto Mexicano del Seguro
6 Social; 2015. ISBN: 978-607-8392-49-
7 a. <http://www.imss.gob.mx/sites/all/statics/guiasclinicas/281GER.pdf>
8
- 9 15. Ministerio de Salud de Chile. Guia Clinica Retinopatía del Prematuro. Ministerio de salud de
10 Chile. Santiago: Minsal, 2010.
11 <https://www.minsal.cl/portal/url/item/721fc45c973b9016e04001011f0113bf.pdf>
12
- 13 16. Ossandon D, Zanolli M, Stevenson R, Agurto R, Ortiz P, Dotan G. A national telemedicine
14 network for retinopathy of prematurity screening. *J AAPOS*. 2018 Apr;22(2):124-127. doi:
15 10.1016/j.jaapos.2017.11.005.
16
- 17 17. Alda E, Lomuto CC, Benítez AM, Bouzas L, Brussa M, Cattaino A, et al. Results of the National
18 Program for the Prevention of Blindness in Childhood by Retinopathy of Prematurity in
19 Argentina (2004-2016). *Arch Argent Pediatr*. 2018 Dec 1;116(6):386-393. doi:
20 10.5546/aap.2018.eng.386.
21 <https://www.sap.org.ar/docs/publicaciones/archivosarg/2018/v116n6a05e.pdf>
22
- 23 18. Patel SN, Martinez MA, Berrones D, Swan R, Ryan MC, Jonas KE, et al. Assessment of a Tele-
24 education System to Enhance Retinopathy of Prematurity Training by International
25 Ophthalmologists-in-Training in Mexico. *Ophthalmology*. 2017 Jul;124(7):953-961. doi:
26 10.1016/j.opthta.2017.02.014
27
- 28 19. Tabarez-Carvajal AC, Montes-Cantillo M, Unkrich KH, Trivedi RH, Peterseim MMW.
29 Retinopathy of prematurity: screening and treatment in Costa Rica. *Br J Ophthalmol*. 2017
30 Dec;101(12):1709-1713. doi: 10.1136/bjophthalmol-2016-310
31
- 32 20. Ministerio de Salud y Protección Social de Colombia. Programa Nacional de Atención Integral
33 en Salud Visual 2016-2022. Page 40. Ministerio de Salud y Protección Social Bogotá. 2016.
34 [https://www.minsalud.gov.co/sites/rid/Lists/BibliotecaDigital/RIDE/VS/PP/ENT/programa-nal-](https://www.minsalud.gov.co/sites/rid/Lists/BibliotecaDigital/RIDE/VS/PP/ENT/programa-nal-salud-visual-2016.pdf)
35 [salud-visual-2016.pdf](https://www.minsalud.gov.co/sites/rid/Lists/BibliotecaDigital/RIDE/VS/PP/ENT/programa-nal-salud-visual-2016.pdf)
36
- 37 21. Zin AA, Moreira ME, Bunce C, Darlow BA, Gilbert CE. Criteria and workload implications
38 retinopathy of prematurity in 7 neonatal units in Rio de Janeiro: screening criteria and workload
39 implications. *Pediatrics*. 2010 Aug;126(2):e410-7. doi: 10.1542/peds.2010-0090.
40
- 41 22. Gordillo L, Villanueva AM, Quinn GE. A practical method for reducing blindness due to
42 retinopathy of prematurity in a developing country. *J Perinat Med*. 2012 Sep 4;40(5):577-82. doi:
43 10.1515/jpm-2011-0225
44
- 45 23. Zepeda C, Gilbert C. Limitations in ROP Programs in 32 Neonatal Intensive Care Units in Five
46 States in Mexico. *BioMed Research International*. Volume 2015, Article ID 712624, 8 pages.
47 <https://www.hindawi.com/journals/bmri/2015/712624/>
48
- 49 24. Gilbert C, Wormald R, Fielder A, Deorari A, Zepeda-Romero LC, Quinn G, et al. Potential for a
50 paradigm change in the detection of retinopathy of prematurity requiring treatment. *Arch Dis*
51 *Child Fetal Neonatal Ed* January 2016 Vol 101 No 1
52

- 1 25. Wang J, Ju R, Chen Y, Zhang L, Hu J, Wu Y, et al . Automated retinopathy of prematurity
- 2 screening using deep neural networks. *EBioMedicine*. 2018 Sep;35:361-368. doi:
- 3 10.1016/j.ebiom.2018.08.033. Epub 2018 Aug 27.