

CERPO

Centro de Referencia Perinatal Oriente

Facultad de Medicina, Universidad de Chile



Seminario N° 71

Embarazo Gemelar

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Juan Guillermo Rodríguez, Dra. Daniela Cisternas O.

10 de Mayo de 2021.-

Introducción



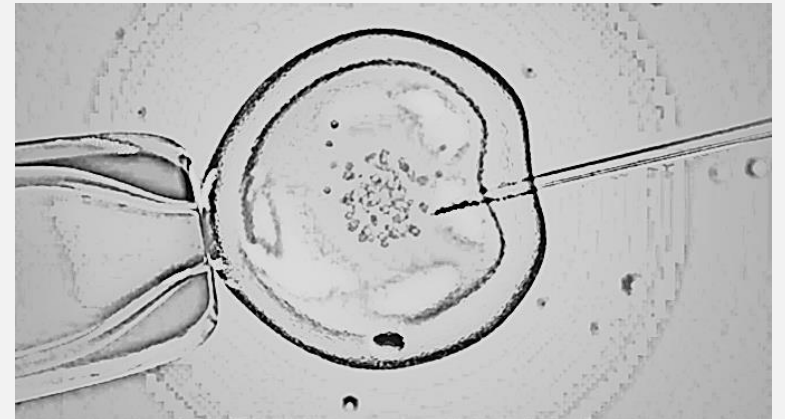
~ 3% en EEUU¹.

En aumento por técnicas de reproducción asistida¹⁻².

Incidencia varía según la cigocidad.

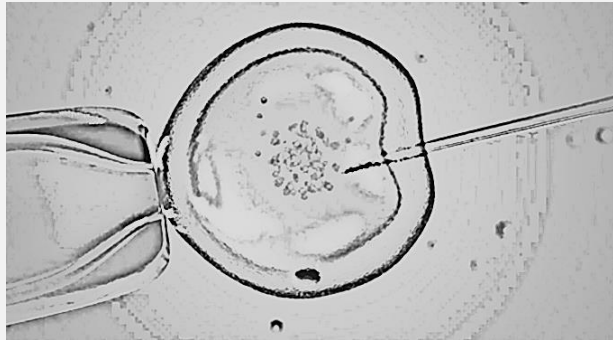
En Chile ha aumentado su incidencia en los últimos 9 años en un 11%².

Incremento en el riesgo de complicaciones maternas y fetales¹.



1. Kiekebusch G, Valdes E, Parra M. Serie guías clínicas: manejo del embarazo Gemelar. Rev Hosp Clin Univ Chile 2016; 27: 246 - 58
2. Fernández C, Poblete J. Prevención de Parto Prematuro en Gemelar: ¿Qué hay de nuevo?. Rev chil obstet ginecol 2017; 82(1): 70 - 76

Factores de riesgo



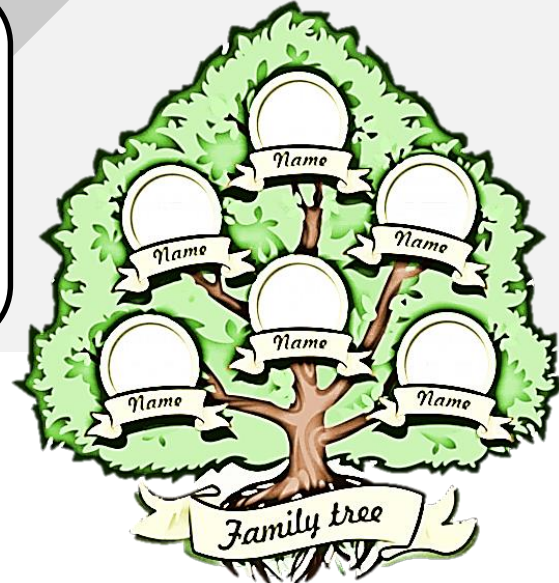
Técnicas de fertilización asistida

Edad Materna



Raza

Historia Familiar

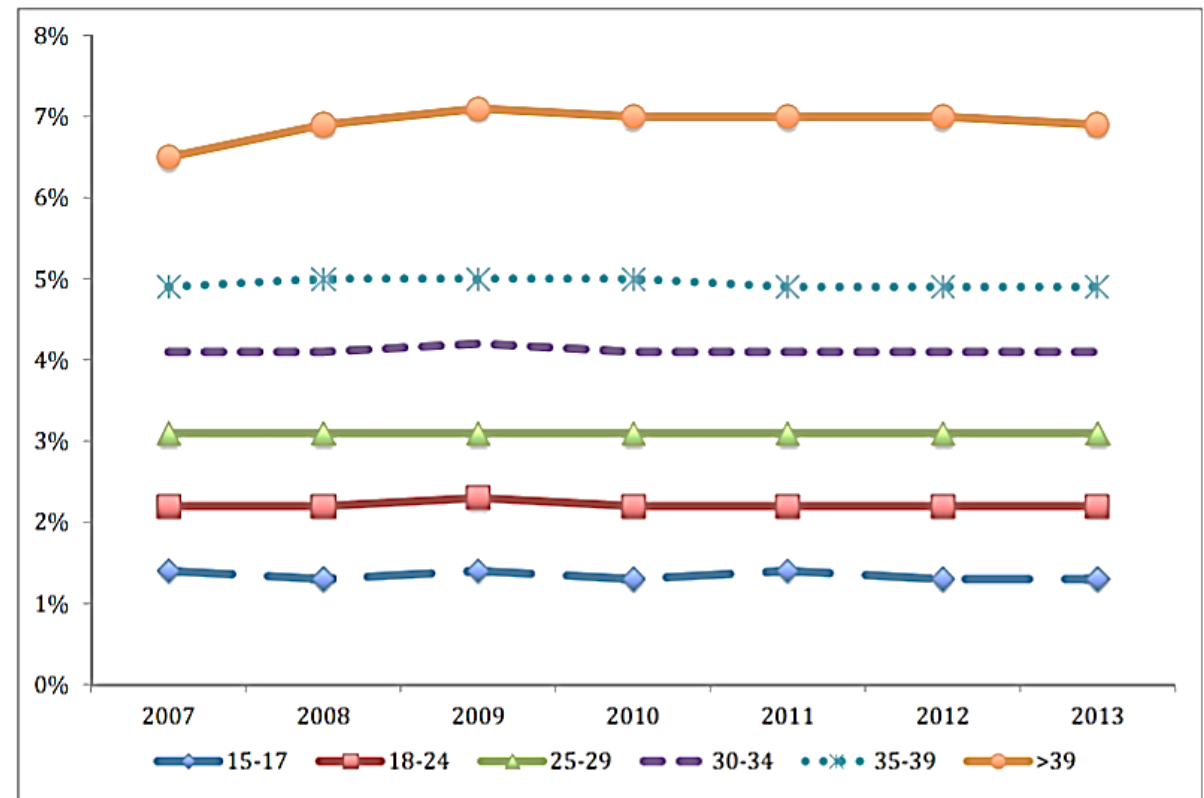


Embarazo múltiple y edad materna



Si bien, la mayoría de los embarazos múltiples se concentran en pacientes entre 25 y 34 años, proporcionalmente el riesgo es mayor a medida que aumenta la edad materna.

Figure 1. Twins as proportion of all neonates by year of birth and maternal age category



Legend: The figure demonstrates the proportion of all live-born neonates for each maternal age category that are twins by year of birth.



Clasificación



- Kiekebusch G, Valdes E, Parra M. Serie guias clinicas: manejo del embarazo Gemelar. Rev Hosp Clin Univ Chile 2016; 27: 246 - 58
- Cunningham F, Leveno KJ, Bloom SL, et al. Williams Obstetrics. 22th edition. International Edition, McGraw-Hall Companies Inc. Chapter 39. Multifetal Gestation.

Diagnóstico y Determinación de la corionicidad

Ecografía 11-14 semanas

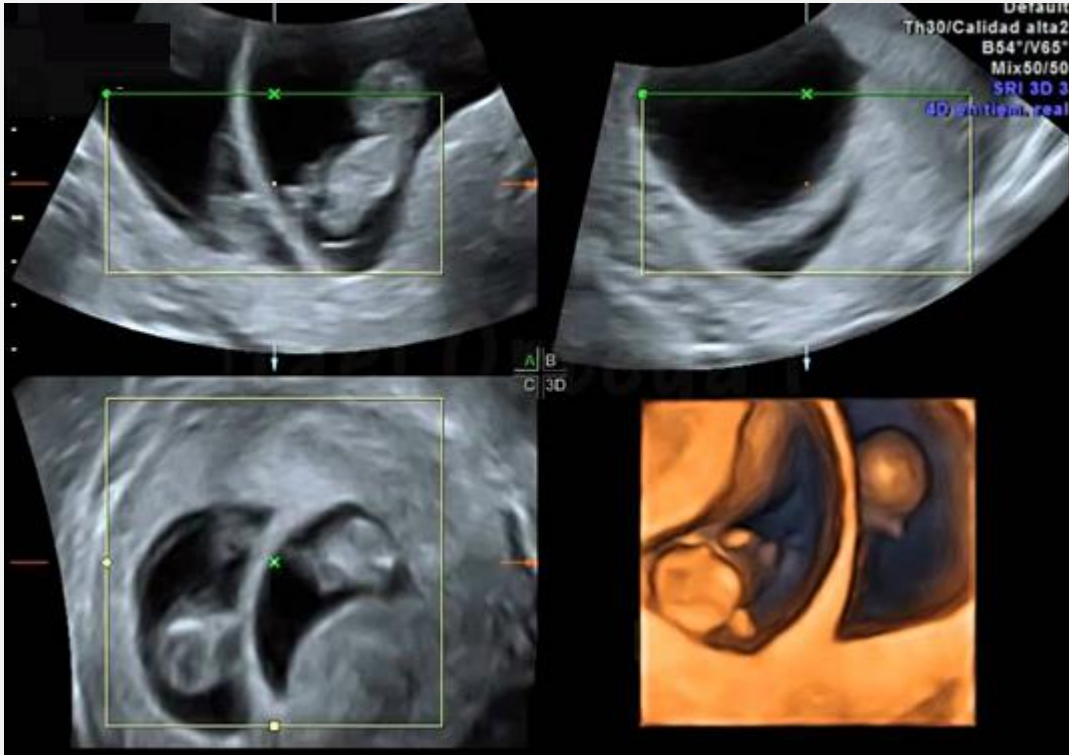
- Determinar edad gestacional (LCN gemelo más grande)
- Corionicidad
- Riesgo de aneuploidías
- Nomenclatura de gemelos

Eco tardía

- Circunferencia craneana gemelo más grande
- Corionicidad



Diagnóstico y Determinación de la corionicidad



Complicaciones maternas

Cambios Hemodinámicos	Síndrome hipertensivo del embarazo <ul style="list-style-type: none"> •HTA gestacional •Preeclampsia •HELLP 	Diabetes Gestacional
Hiperemesis gravídica	Edema Pulmonar	Anemia
	Infección Urinaria	

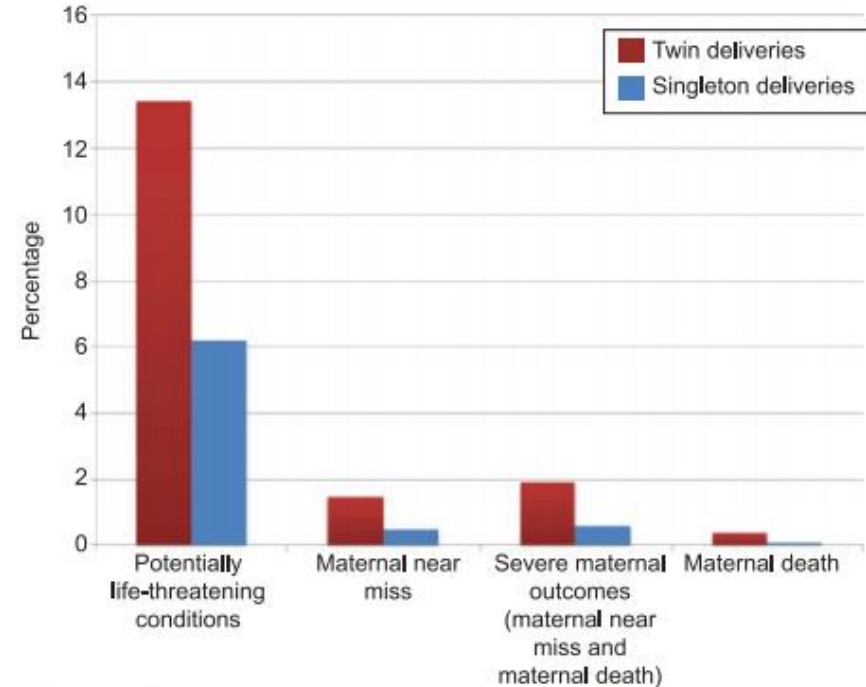


Fig. 3. Proportion of potentially life-threatening conditions, maternal near miss, severe maternal outcomes, and maternal death comparatively among twin and singleton deliveries.

Santana. Twin Pregnancy and Severe Maternal Outcomes. Obstet Gynecol 2016.

Table 1. Potentially Life-Threatening Condition, Maternal Near Miss, and Maternal Death for Twin and Singleton Deliveries by Human Development Index, Country, and Region: World Health Organization Multicountry Survey, 2010–2011

HDI and Country	Twin Deliveries				Singleton Deliveries				P
	NC (%)	PLTC (%)	MNM (%)	MD (%)	NC (%)	PLTC (%)	MNM (%)	MD (%)	
Very high HDI	206 (79.8)	51 (19.8)	0	1 (0.4)	15,138 (89.0)	1,785 (10.5)	76 (0.4)	7 (<0.1)	*
Argentina	130 (85.5)	21 (13.8)	0	1 (0.7)	8,891 (92.4)	686 (7.1)	41 (0.4)	7 (0.1)	.019
Japan	24 (63.2)	14 (36.8)	0	0	2,858 (81.7)	620 (17.7)	21 (0.6)	0	.274
Qatar	52 (76.5)	16 (23.5)	0	0	3,389 (87.3)	479 (12.3)	14 (0.4)	0	*
High HDI	783 (84.5)	129 (13.9)	14 (1.5)	1 (0.1)	62,073 (92.7)	4,546 (6.8)	309 (0.5)	16 (<0.1)	<.001
Brazil	68 (84.0)	13 (16.0)	0	0	6,484 (93.0)	474 (6.8)	13 (0.2)	1 (<0.1)	.287
Ecuador	185 (93.4)	13 (6.6)	0	0	8,601 (86.1)	1,366 (13.7)	20 (0.2)	5 (0.1)	.573
Lebanon	89 (79.5)	18 (16.1)	4 (3.6)	1 (0.9)	3,745 (95.2)	172 (4.4)	14 (0.4)	1 (<0.1)	<.001
Mexico	134 (77.5)	32 (18.5)	7 (4.0)	0	12,123 (92.5)	863 (6.6)	120 (0.9)	2 (<0.1)	<.001
Peru	150 (79.8)	36 (19.1)	2 (1.1)	0	14,015 (93.4)	903 (6.0)	87 (0.6)	5 (<0.1)	<.001
Sri Lanka	157 (89.7)	17 (9.7)	1 (0.6)	0	17,105 (95.4)	768 (4.3)	55 (0.3)	2 (<0.1)	.008
Medium HDI	1,257 (85.0)	199 (13.5)	18 (1.2)	4 (0.3)	94,905 (92.7)	7,014 (6.9)	348 (0.3)	87 (0.1)	<.001[†]
Region									
Africa	1,219 (84.2)	196 (13.5)	27 (1.9)	5 (0.3)	67,547 (93.3)	4,181 (5.8)	495 (0.7)	139 (0.2)	<.001
Asia	2,078 (85.8)	299 (12.4)	32 (1.3)	12 (0.5)	161,118 (94.2)	9,129 (5.3)	686 (0.4)	131 (0.1)	<.001
Latin America	735 (82.8)	141 (15.9)	11 (1.2)	1 (0.1)	58,412 (90.3)	5,935 (9.2)	314 (0.5)	24 (<0.1)	<.001
Total	4,032 (84.8)	636 (13.4)	70 (1.5)	18 (0.4)	287,077 (93.2)	19,245 (6.2)	1,495 (0.5)	294 (0.1)	<.001

HDI, Human Development Index; NC, no complication; PLTC, potentially life-threatening condition; MNM, maternal near miss; MD, maternal death; OPT, Occupational Palestinian Territory; DR, Democratic Republic. Data are n (%) unless otherwise specified.

Table 5. Estimated Risks of Maternal Near Miss (Organ Dysfunction Conditions) for Twin Deliveries: World Health Organization Multicountry Survey, 2010–2011

Organ Dysfunction Condition	Twin Deliveries	Singleton Deliveries	Total	PR _{adj} (95% CI)*
Cardiovascular dysfunction [†]	42 (0.9)	812 (0.3)	854 (0.3)	3.35 (2.36–4.76)
Respiratory dysfunction [‡]	34 (0.7)	525 (0.2)	559 (0.2)	4.20 (2.89–6.11)
Coagulation or hematologic dysfunction [§]	36 (0.8)	603 (0.2)	639 (0.2)	3.87 (2.70–5.54)
Uterine dysfunction or hysterectomy	13 (0.3)	353 (0.1)	366 (0.1)	2.39 (1.32–4.31)
Neurologic dysfunction [¶]	9 (0.2)	227 (0.1)	236 (0.1)	2.57 (1.33–4.98)
Hepatic dysfunction [§]	11 (0.2)	206 (0.1)	217 (0.1)	3.46 (2.20–5.46)
Renal dysfunction [#]	15 (0.3)	196 (0.1)	211 (0.1)	4.96 (2.92–8.44)
Any organ dysfunction**	85 (1.8)	1,756 (0.6)	1,841 (0.6)	3.14 (2.49–3.96)
Total	4,756	308,111	312,867	

PR_{adj}, prevalence ratio adjusted for cluster design effect; CI, confidence interval.

Data are n (%) unless otherwise specified.

Bold indicates statistical significance (95% CI not including the value 1.0).

* Adjusted for the cluster design effect.

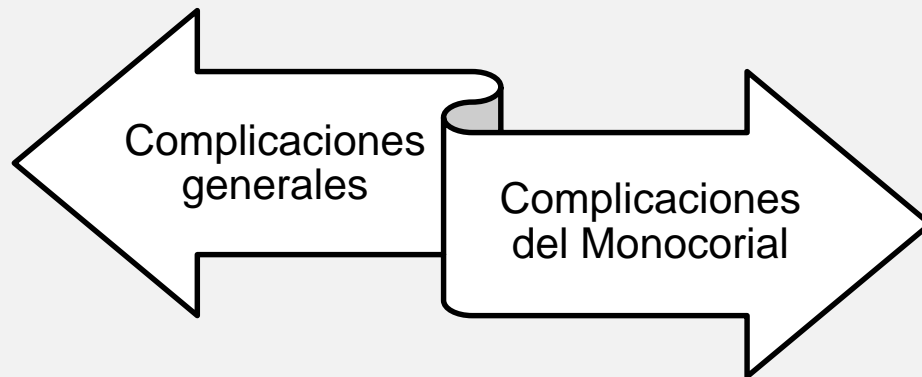
Missing information for [†]220; [‡]198; [§]182; ^{||}178; [¶]180; [#]193; **259 cases.

Resultados maternos por corionicidad (Modificado de Carter et al)



	Bicorial N= 1747	Monocorial N= 554	RR no ajustado (95% IC)	OR Ajustado (95% IC)
Preeclampsia	335 (20,48%)	101 (19,35%)	0,93 (0,73-1,19)	0,92 (0,70-1,20)
Diabetes Gestacional	102 (6,23%)	34 (6,51%)	1,04 (0,70-1,57)	1,08 (0,69-1,67)
Desprendimiento de Placenta	29 (1,77%)	9 (1,72%)	0,97 (0,46-2,07)	0,90 (0,40-1,99)
Placenta Previa	15 (0,92%)	2 (0,38%)	0,42 (0,09-1,83)	0,26 (0,03-1,98)
Parto Prematuro	700 (42,79%)	212 (40,61%)	0,91 (0,75-1,12)	0,90 (0,73-1,11)
PPROM	184 (11,24%)	53 (10,15%)	0,89 (0,65-1,23)	0,83 (0,58-1,18)
Cesárea	1655 (94,74%)	524 (94,58%)	0,97 (0,64-1,48)	1,02 (0,64-1,63)

Complicaciones Fetales



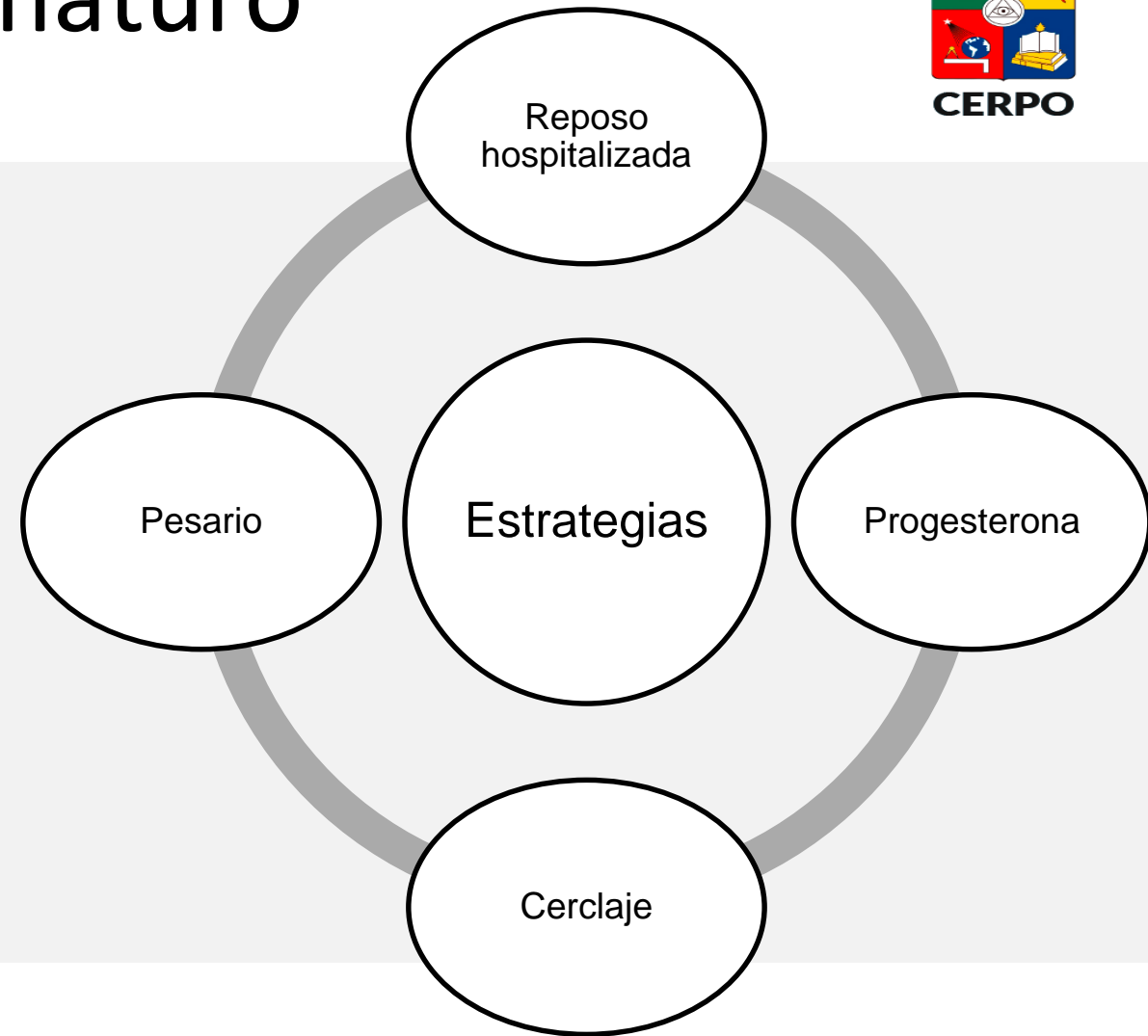
Parto Prematuro

50% Embarazos gemelares

Causa más importante de morbilidad y mortalidad neonatal

EEUU → 23% de los partos <32 semanas

Prevención → cervicometría



Parto Prematuro



EVOLUCIÓN DEL EMBARAZO MÚLTIPLE Y PREMATUREZ. CHILE 2000-2009

Año	NV	Múltiples =2	Múltiples >2	Total múltiples	<37 s (%)	32-36 s (%)
2000	248350	1,66	0,06	1,72	50,56	43,10
2001	245698	1,78	0,05	1,83	51,19	43,78
2002	238678	1,78	0,05	1,83	53,48	45,38
2003	234005	1,77	0,05	1,82	54,29	45,81
2004	229021	1,74	0,05	1,79	57,32	49,66
2005	230366	1,75	0,04	1,79	59,08	49,83
2006	231218	1,77	0,05	1,83	59,49	48,93
2007	239958	1,81	0,04	1,85	60,08	49,63
2008	245661	1,87	0,04	1,91	61,89	52,04
2009	251187	1,84	0,05	1,89	62,47	52,60

NV: nacidos vivos.

Reposo hospitalizada

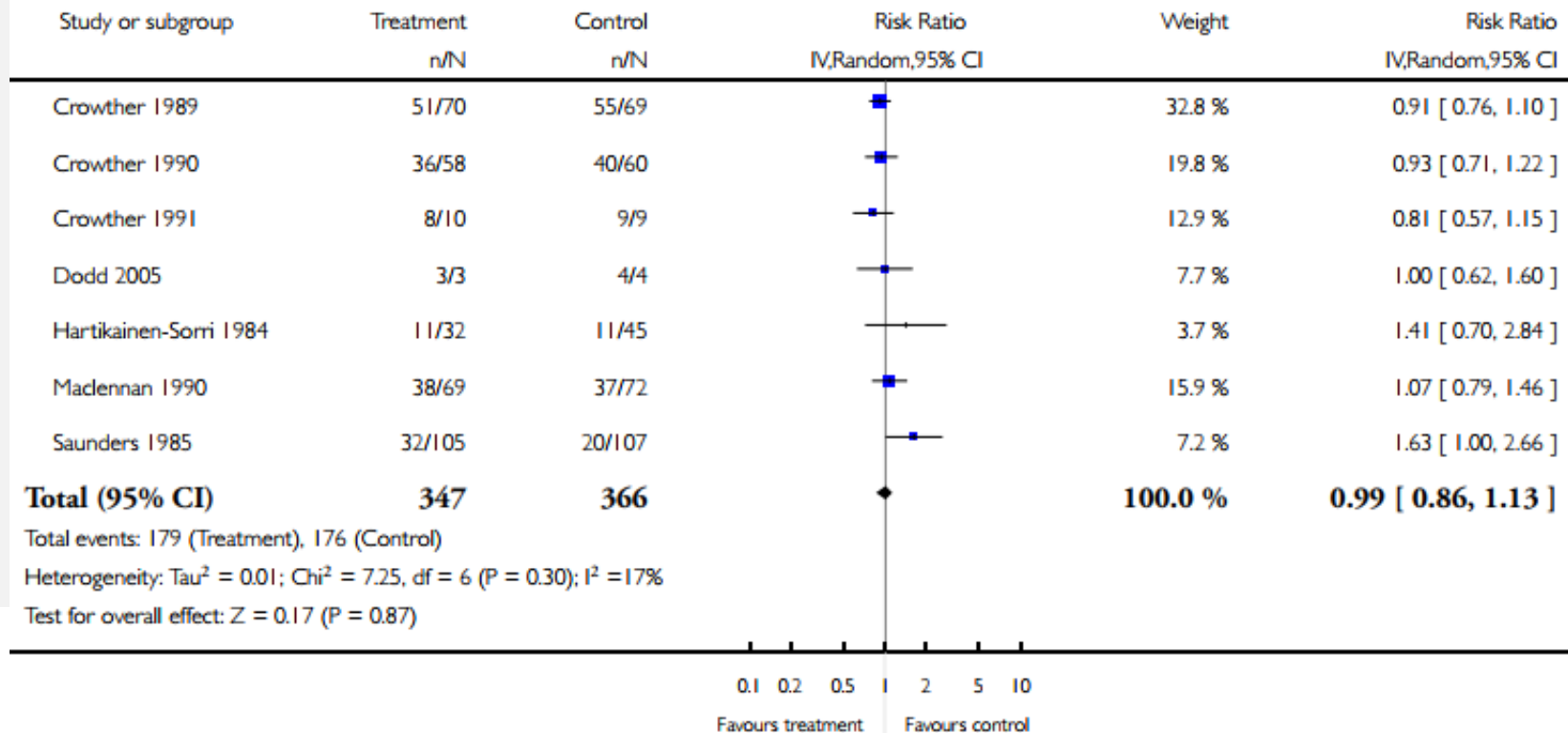


Analysis 1.5. Comparison 1 Hospitalisation for bed rest for women with a multiple pregnancy, Outcome 5 Preterm delivery (< 37 weeks).

Review: Hospitalisation and bed rest for multiple pregnancy

Comparison: 1 Hospitalisation for bed rest for women with a multiple pregnancy

Outcome: 5 Preterm delivery (< 37 weeks)



Progesterona

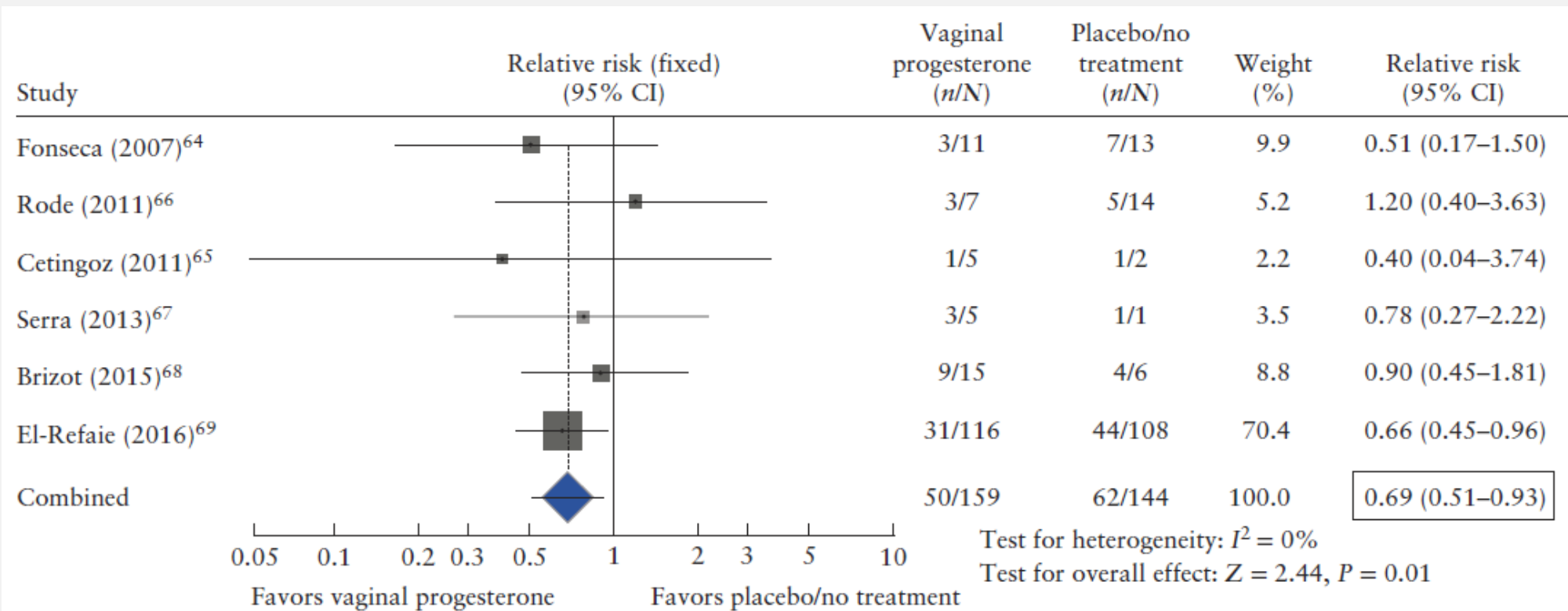


Figure 3 Forest plot of the effect of vaginal progesterone on the risk of preterm birth < 33 weeks' gestation. CI, confidence interval.

Progesterona

Table 3 Effect of vaginal progesterone on the risk of preterm birth

<i>Outcome</i>	<i>Trials</i> (n ^{refs})	<i>Events (n)/Total (N)</i>		<i>Pooled RR</i> (95% CI)	<i>I²</i> (%)	<i>NNT</i> (95% CI)
		<i>Vaginal progesterone</i>	<i>Placebo/no treatment</i>			
Preterm birth < 37 weeks	6 ⁶⁴⁻⁶⁹	137/159	131/144	0.94 (0.86–1.02)	0	—
Preterm birth < 36 weeks	6 ⁶⁴⁻⁶⁹	112/159	110/144	0.92 (0.80–1.05)	0	—
Preterm birth < 35 weeks	6 ⁶⁴⁻⁶⁹	90/159	98/144	0.83 (0.69–0.99)	0	9 (5–147)
Preterm birth < 34 weeks	6 ⁶⁴⁻⁶⁹	63/159	78/144	0.71 (0.56–0.91)	0	6 (4–21)
Preterm birth < 32 weeks	6 ⁶⁴⁻⁶⁹	29/159	46/144	0.51 (0.34–0.77)	0	6 (5–14)
Preterm birth < 30 weeks	6 ⁶⁴⁻⁶⁹	14/159	22/144	0.47 (0.25–0.86)	0	12 (9–47)
Preterm birth < 28 weeks	6 ⁶⁴⁻⁶⁹	9/159	12/144	0.51 (0.24–1.08)	0	—
Spontaneous preterm birth < 33 weeks	6 ⁶⁴⁻⁶⁹	42/159	54/144	0.67 (0.48–0.93)	0	8 (5–38)
Spontaneous preterm birth < 34 weeks	6 ⁶⁴⁻⁶⁹	55/159	69/144	0.71 (0.54–0.93)	0	7 (5–30)

CI, confidence interval; NNT, number needed to treat; refs, reference numbers; RR, relative risk.

Progesterona

Table 4 Effect of vaginal progesterone on the risk of adverse perinatal outcomes

Outcome	Trials (n ^{refs})	Events (n)/Total (N)		Pooled RR (95% CI)		I ² (%)	NNT (95% CI)
		Vaginal progesterone	Placebo/no treatment	Assuming independence between twins	Adjustment for non- independence between twins		
Respiratory distress syndrome	6 ⁶⁴⁻⁶⁹	102/311	131/280	0.67 (0.55–0.82)	0.70 (0.56–0.89)	0	6 (4–16)
Necrotizing enterocolitis	5 ⁶⁴⁻⁶⁸	1/82	0/68	1.00 (0.04–22.43)	1.07 (0.05–22.25)	NA	—
Intraventricular hemorrhage	5 ⁶⁴⁻⁶⁸	2/80	2/68	0.93 (0.15–5.75)	1.47 (0.22–9.63)	0	—
Proven neonatal sepsis	5 ⁶⁴⁻⁶⁸	4/80	7/68	0.44 (0.13–1.46)	0.59 (0.18–1.93)	0	—
Retinopathy of prematurity	5 ⁶⁴⁻⁶⁸	1/80	1/68	0.42 (0.07–2.56)	0.45 (0.08–2.59)	17	—
Fetal death	6 ⁶⁴⁻⁶⁹	9/318	9/288	0.57 (0.23–1.42)	0.68 (0.26–1.84)	0	—
Neonatal death	6 ⁶⁴⁻⁶⁹	34/318	63/288	0.50 (0.34–0.71)	0.53 (0.35–0.81)	25	8 (5–19)
Perinatal death	6 ⁶⁴⁻⁶⁹	43/318	72/288	0.51 (0.36–0.70)	0.58 (0.39–0.84)	24	7 (5–20)
Composite neonatal morbidity/mortality*	5 ⁶⁴⁻⁶⁸	23/84	28/70	0.57 (0.36–0.93)	0.61 (0.34–0.98)	0	6 (3–109)
Birth weight < 1500 g	6 ⁶⁴⁻⁶⁹	48/315	73/280	0.52 (0.38–0.72)	0.53 (0.35–0.80)	17	7 (5–17)
Birth weight < 2500 g	6 ⁶⁴⁻⁶⁹	244/315	223/280	0.97 (0.89–1.06)	0.99 (0.89–1.10)	0	—
Admission to the NICU	6 ⁶⁴⁻⁶⁹	211/315	209/282	0.92 (0.83–1.02)	0.95 (0.84–1.08)	0	—
Mechanical ventilation	6 ⁶⁴⁻⁶⁹	49/311	76/280	0.52 (0.37–0.71)	0.54 (0.36–0.81)	0	7 (5–17)

*Occurrence of any of the following events: respiratory distress syndrome, intraventricular hemorrhage, necrotizing enterocolitis, proven neonatal sepsis or neonatal death. CI, confidence interval; NA, not applicable; NICU, neonatal intensive care unit; NNT, number needed to treat; refs, reference numbers; RR, relative risk.

Cerclaje

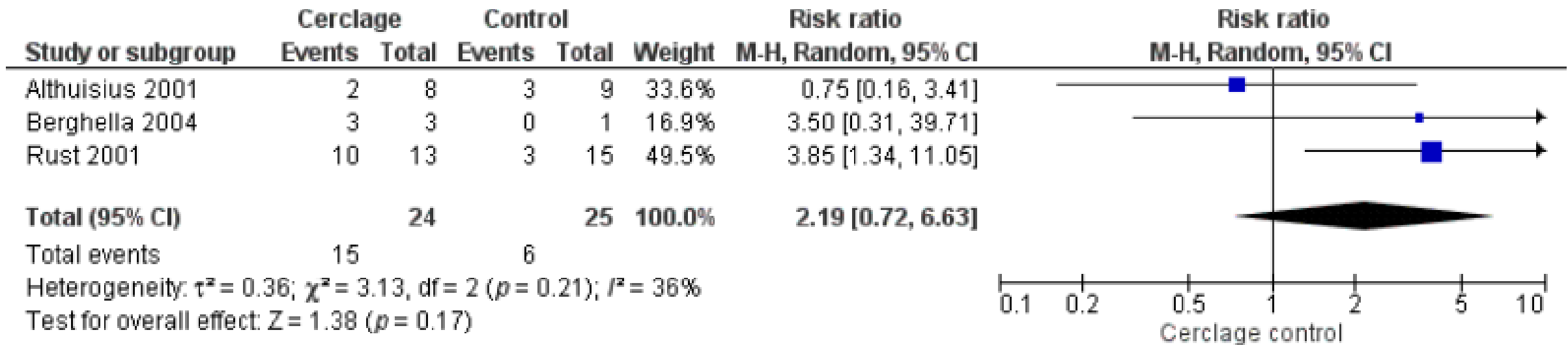


Figure 3. Meta-analysis of cerclage in twins and preterm birth at <34 weeks of gestation.

Does cervical cerclage decrease preterm birth in twin pregnancies with a short cervix?

Tracy M Adams DO, Timothy J Rafael MD, Nadia B Kunzier DO, Supriya Mishra MS, Rose Calixte PhD & Anthony M Vintzileos MD

Cerclaje



Cervicometría

<25

<15

Parto Pretérmino

Outcome Neonatal

Parto Pretérmino

Outcome Neonatal

No significativo

No significativo

Significativo para parto <35 semanas

No significativo

Pregnancy Prolongation (weeks)**	13.1 ± 4.6	8.9 ± 5.1	0.005		
PTB < 37 Weeks (N%)	19 (70.4%)	17 (81.0%)	0.51	0.87 (0.63 – 1.20)	0.88 (0.59 – 1.33)
PTB < 35 Weeks (N%)	10 (37.0%)	15 (71.4%)	0.02	0.52 (0.30 – 0.91)	0.49 (0.26 – 0.93)
PTB < 32 Weeks (N%)	5 (18.5%)	10 (47.6%)	0.06	0.39 (0.16 – 0.97)	0.31 (0.12 – 0.86)
PTB < 28 Weeks (N%)	4 (14.8%)	5 (23.8%)	0.4	0.62 (0.19 – 2.03)	0.30 (0.08 – 1.05)

Pesario



TABLE 3

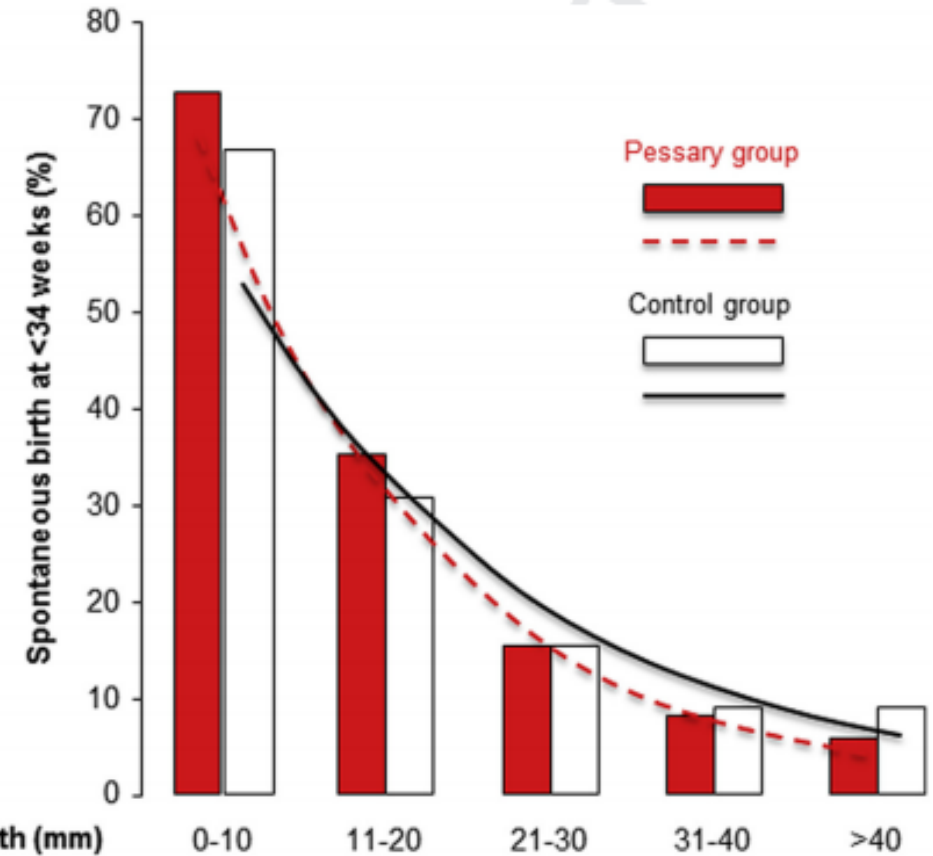
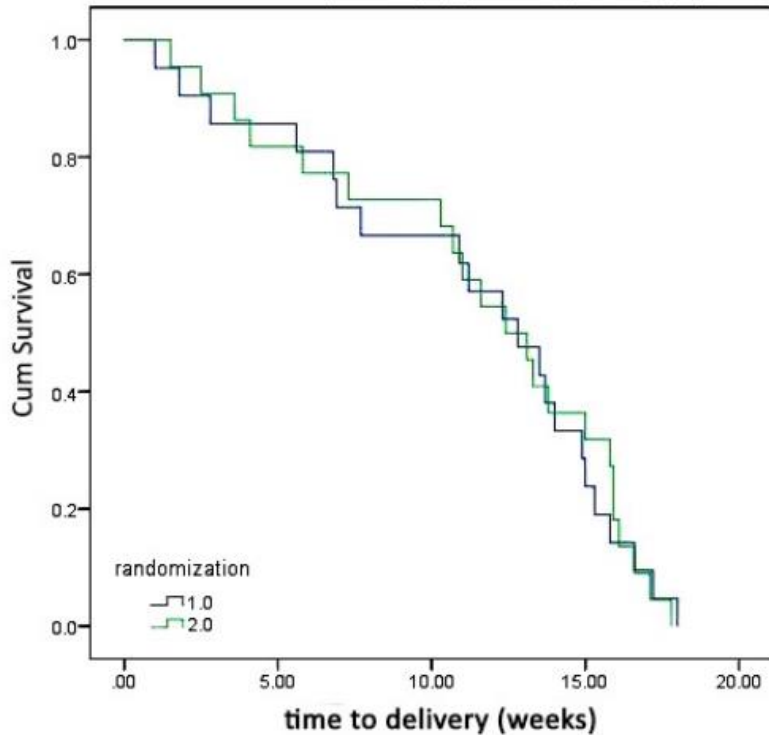
Outcomes according to cervical length at randomization ≤ 25 mm and > 25 mm

Outcome	Pregnancy level			Fetal/neonatal level		
	Pessary group	Control group	RR (95% CI)	Pessary group	Control group	RR (95% CI)
Cervical length ≤ 25 mm (n = 214)	n = 106	n = 108		n = 212	n = 216	
Primary outcome, n (%)						
Spontaneous birth at < 34 wk	33 (31.1)	28 (25.9)	1.201 (0.784–1.839)	—	—	—
Secondary outcomes, n (%)						
Birthweight < 2500 g	82 (77.4)	89 (82.4)	0.939 (0.820–1.074)	149 (70.3)	150 (69.4)	1.012 (0.894–1.146)
Birthweight < 1500 g	24 (22.6)	21 (19.4)	1.164 (0.692–1.960)	45 (21.2)	36 (16.7)	1.274 (0.858–1.891)
Perinatal death	13 (12.3)	6 (5.6)	2.208 (0.872–5.592)	20 (9.4)	12 (5.6)	1.698 (0.852–3.386)
Secondary outcomes in survivors, n (%)	n = 99	n = 102		n = 192	n = 204	
Adverse neonatal event	23 (23.2)	20 (19.6)	1.185 (0.696–2.016)	34 (17.7)	30 (14.7)	1.204 (0.768–1.888)
Neonatal therapy	36 (36.4)	31 (30.4)	1.197 (0.808–1.772)	56 (29.2)	52 (25.5)	1.144 (0.829–1.579)

Pesario



Probability of continued pregnancy among twin mother receiving cervical pessary (2) vs. no pessary (1)



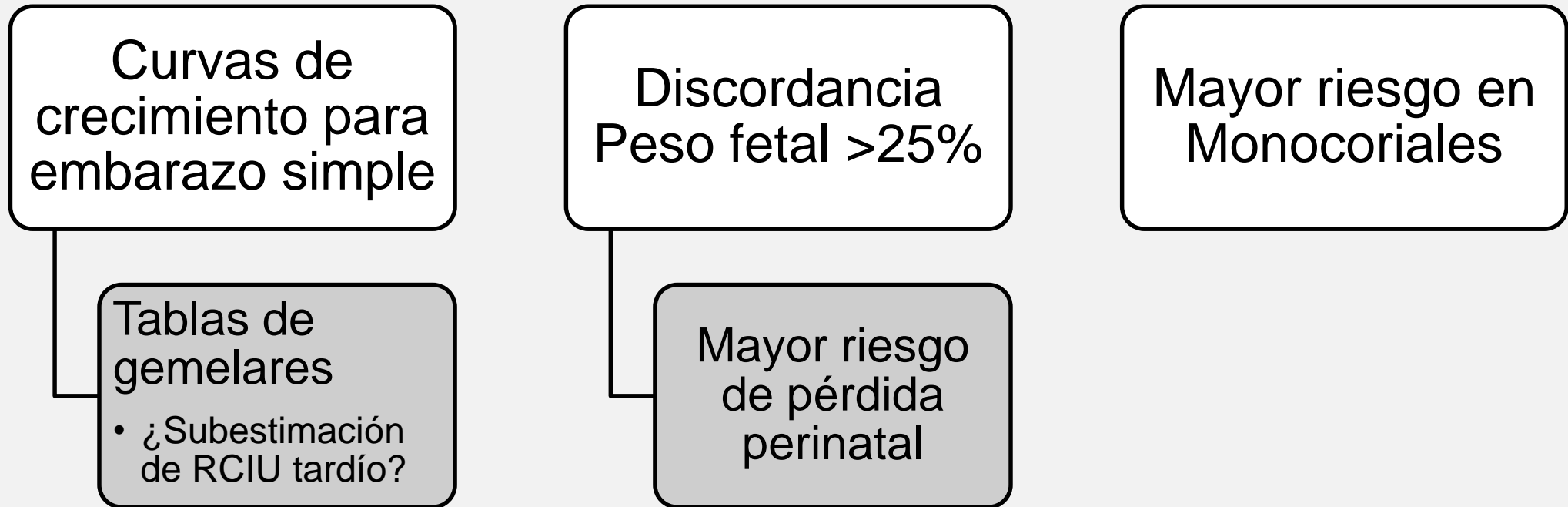
Cervical length (mm)	0-10	11-20	21-30	31-40	>40
Control group	11	45	196	279	58
Pessary group	12	37	211	284	44

Association between cervical length at randomization and rate of spontaneous birth at <34 weeks in pessary (red bars, red interrupted regression curve) and control (white bars, black regression curve) groups.

Nicolaides. RCT of cervical pessary in twin gestations. Am J Obstet Gynecol 2015.

- Nicolaides KH, Syngelaki A, Poon LC, de Paco Matallana C, Plasencia W, Molina FS, Picciarelli G, Tul N, Celik E, Lau TK, Conturso R. Cervical pessary placement for prevention of preterm birth in unselected twin pregnancies: a randomized controlled trial. *Am J Obstet Gynecol.* 2016 Jan;214(1):3.e1-9.
- Berghella V, Dugoff L, Ludmir J. Prevention of preterm birth with pessary in twins (PoPPT): a randomized controlled trial. *Ultrasound Obstet Gynecol.* 2017 May;49(5):567-572.

Restricción de crecimiento Fetal



Restricción de crecimiento Fetal Seguimiento



Bicorial

Similar a embarazo simple
Seguimiento ecográfico y
Doppler cada 2 semanas

Monocorial

Crecimiento cada 2 semanas
Doppler fetal semanal
Evaluar signos de
complicación

- STFF
- SAP

Riesgo de FMIU del gemelo
con RCF antes de las 26
semanas

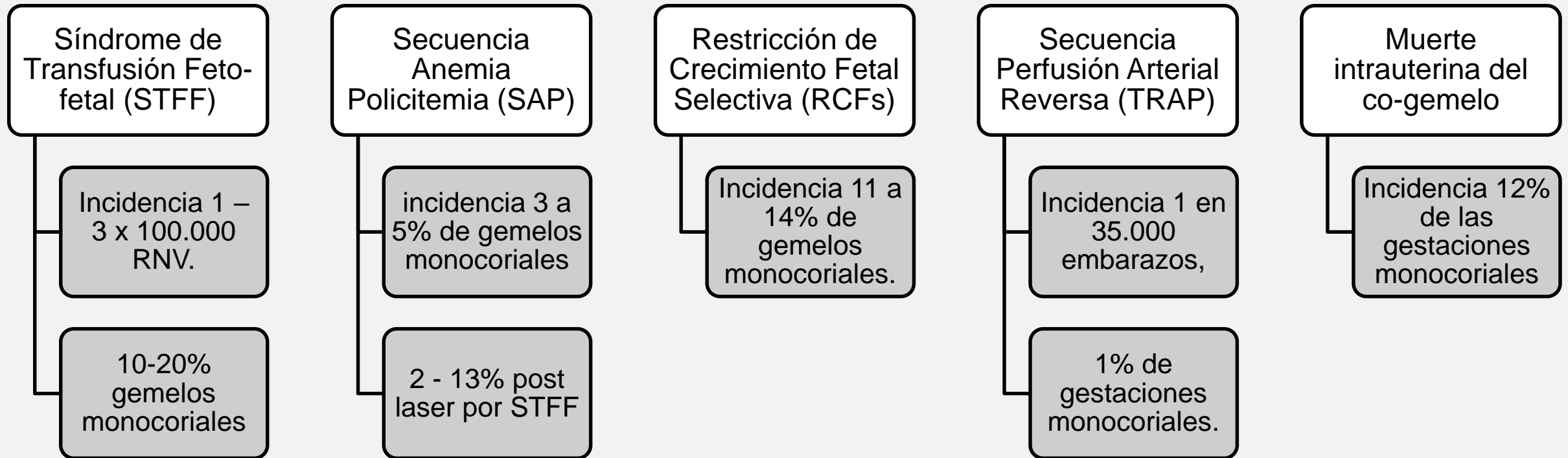
- Evaluar interrupción selectiva

Monocorial Vs Bicorial



	Bicorial N= 1747	Monocorial N= 554	RR no ajustado (95% IC)	OR Ajustado (95% IC)
Restricción de Crecimiento fetal	372 (21,29%)	118 (21,30%)	1,00 (0,79-1,26)	1,04 (0,82-1,32)
Parto <34 semanas	337 (19,79%)	137 (25,75%)	1,41 (1,12-1,77)	1,47 (1,17-1,86)
Parto <28 semanas	41 (2,41%)	30 (5,64%)	2,42 (1,50-3,92)	2,58 (1,58-4,20)
Ingreso a UCI neonatal	334 (21,02%)	164 (27,13%)	1,40 (1,11-1,76)	1,41 (1,12-1,78)
Muerte fetal intrauterina	56 (3,21%)	29 (5,23%)	1,67 (1,05-2,64)	1,81 (1,13-2,82)
Muerte Neonatal	29 (1,66%)	4 (0,72%)	0,43 (0,15-1,23)	0,43 (0,13-1,47)

Complicaciones del embarazo monocorial



- Sepúlveda A. Guía clínica: Complicaciones de embarazo gemelar monocorial. Hospital Clínico Universidad de Chile.
- Djaafri F, Stirnemann J, Mediouni I, Colmant C, Ville Y. Twinetwin transfusion syndrome - What we have learned from clinical trials. Seminars in Fetal & Neonatal Medicine xxx (2017) 1e9.

Flujograma de manejo



Bicorial –
Biamniótico

- Control Ecográfico cada 2 semanas

Monocorial

- Control Ecográfico cada semanal

Eco 11-14

- Corionicidad
- Marcadores de STFF

Eco Tardía

- Marcadores de 2° trimestre de corionicidad

Cuello <25 mm

- Progendo

Cuello <15 mm sintomática

- Evaluar signos de infección para eventual cerclaje

Flujograma de manejo

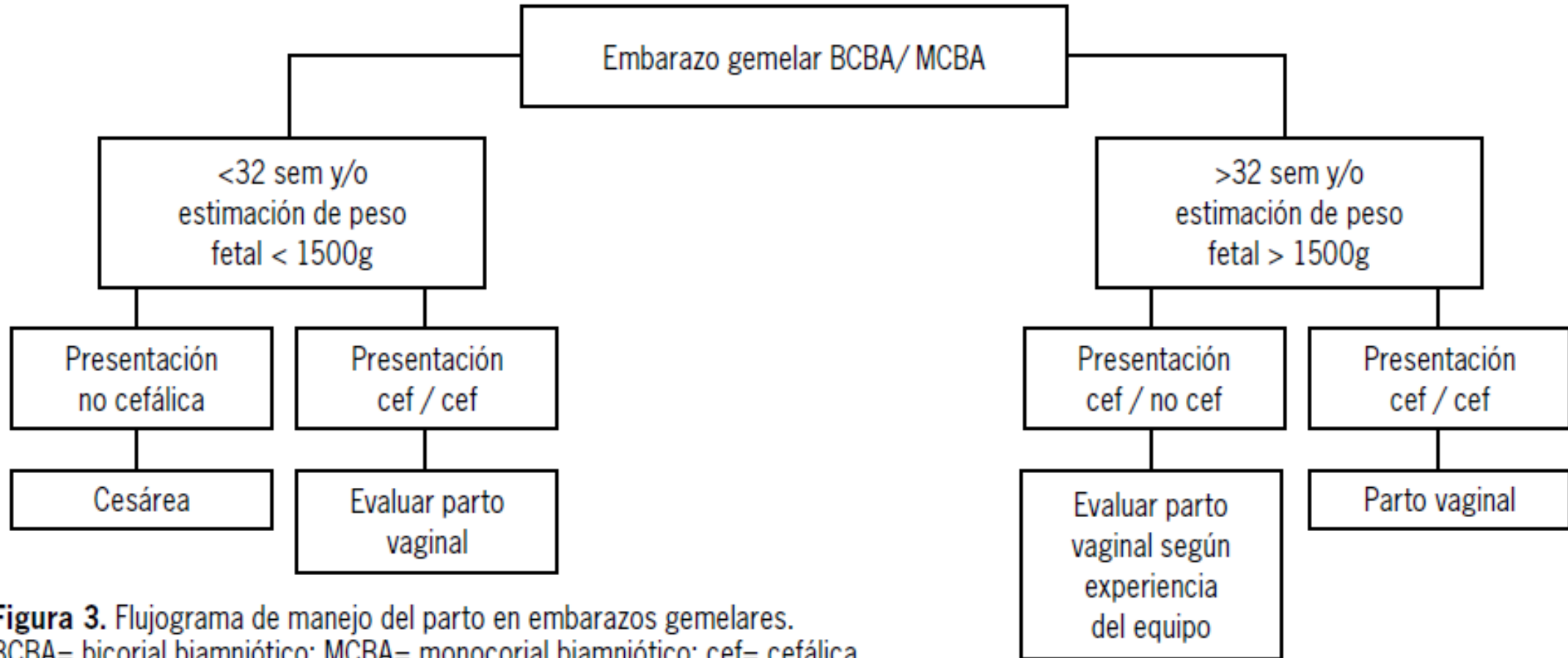


Figura 3. Flujograma de manejo del parto en embarazos gemelares.
BCBA= bicorial biamniótico; MCBA= monocorial biamniótico; cef= cefálica.

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Seminario N° 71

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