



# Vellosidades coriales “raras”

Enfermedad trofoblástica, displasia mesenquimal y tumores

Curso corto sobre patología placentaria

Alfons Nadal

[anadal@clinic.ub.es](mailto:anadal@clinic.ub.es)

Hospital Clínic Universitat de Barcelona

- Mola hidatidiforme completa
- Mola hidatidiforme parcial
- Displasia mesenquimática
- VelloSIDADES dismórficas
- Tumores



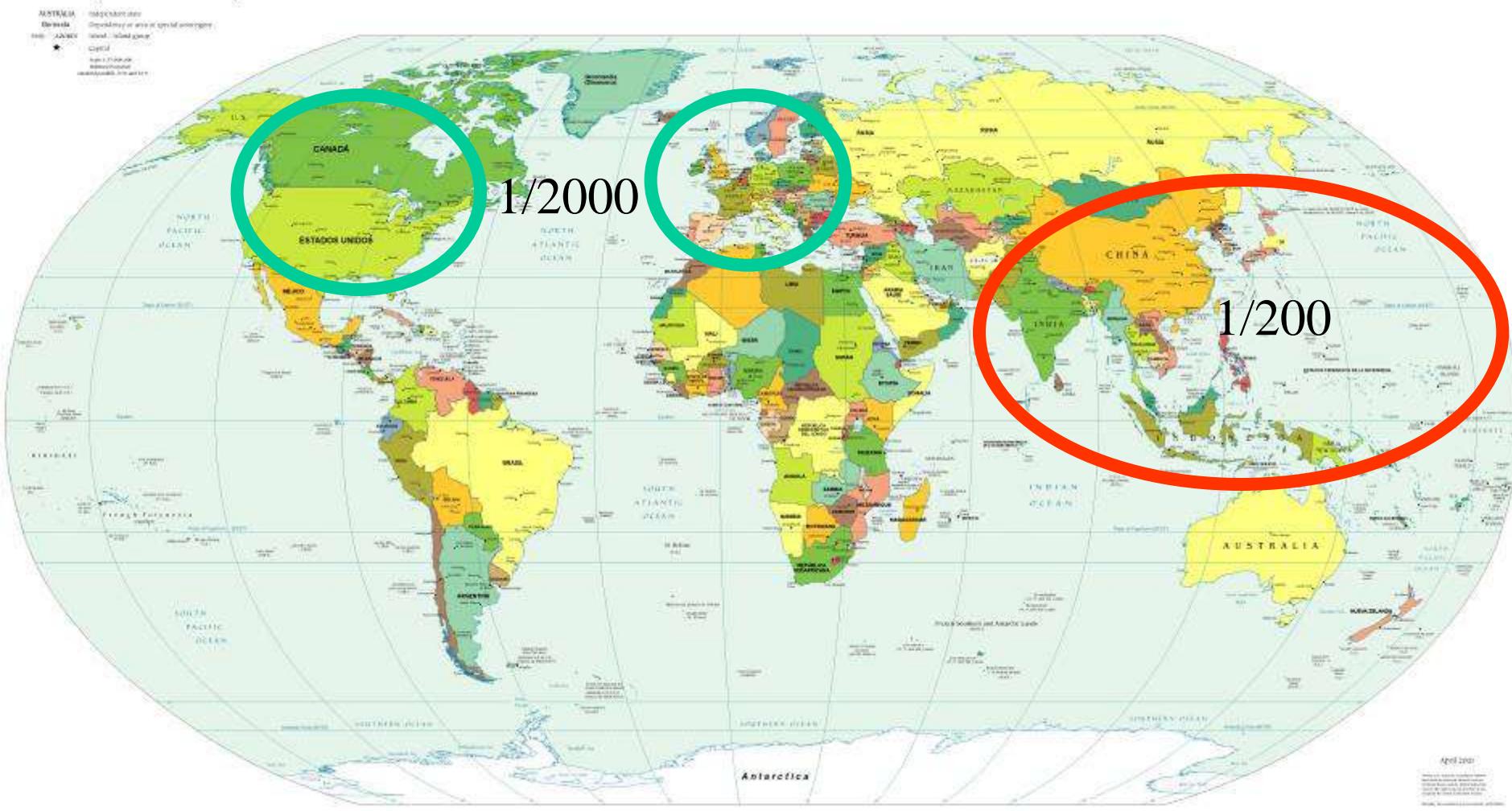
# ¿Mola hidatiforme o mola hidatidiforme?

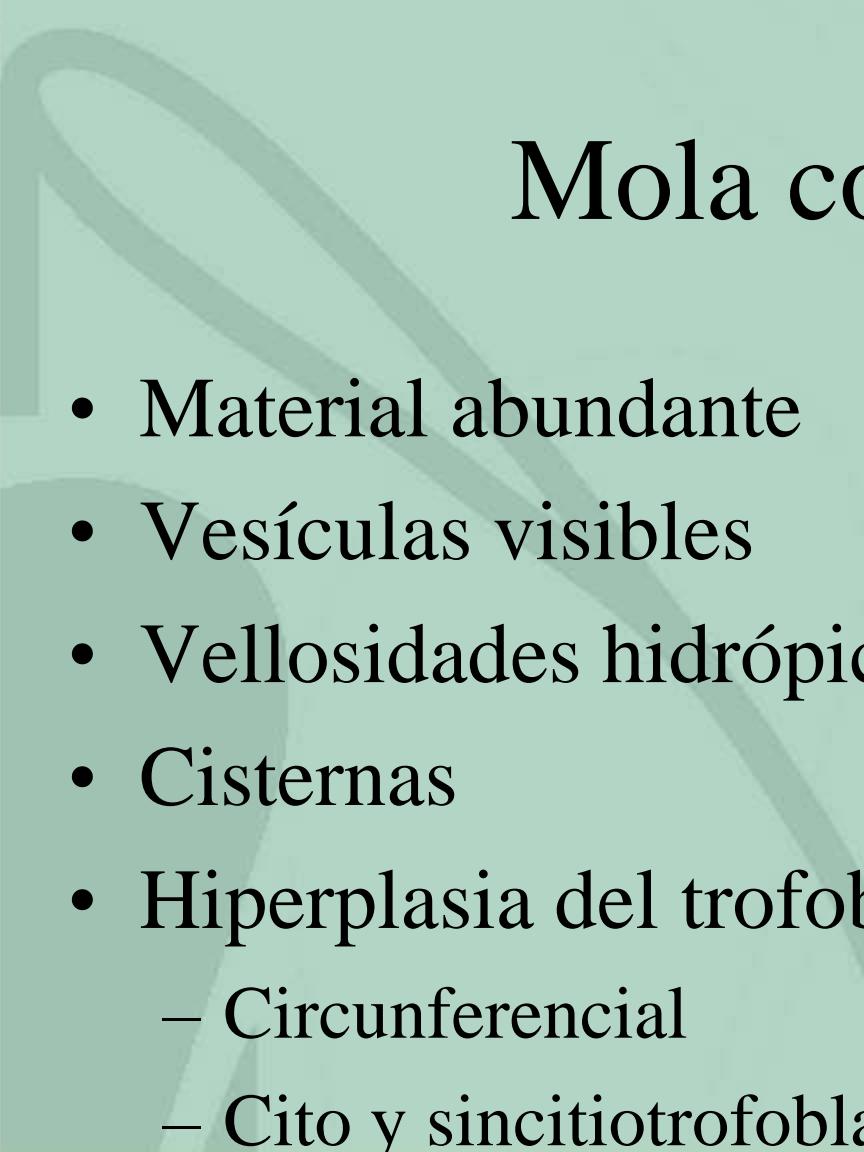
Hidatiforme = hidatidiforme

*“Con forma de hidátide”*



Political Map of the World, April 2001





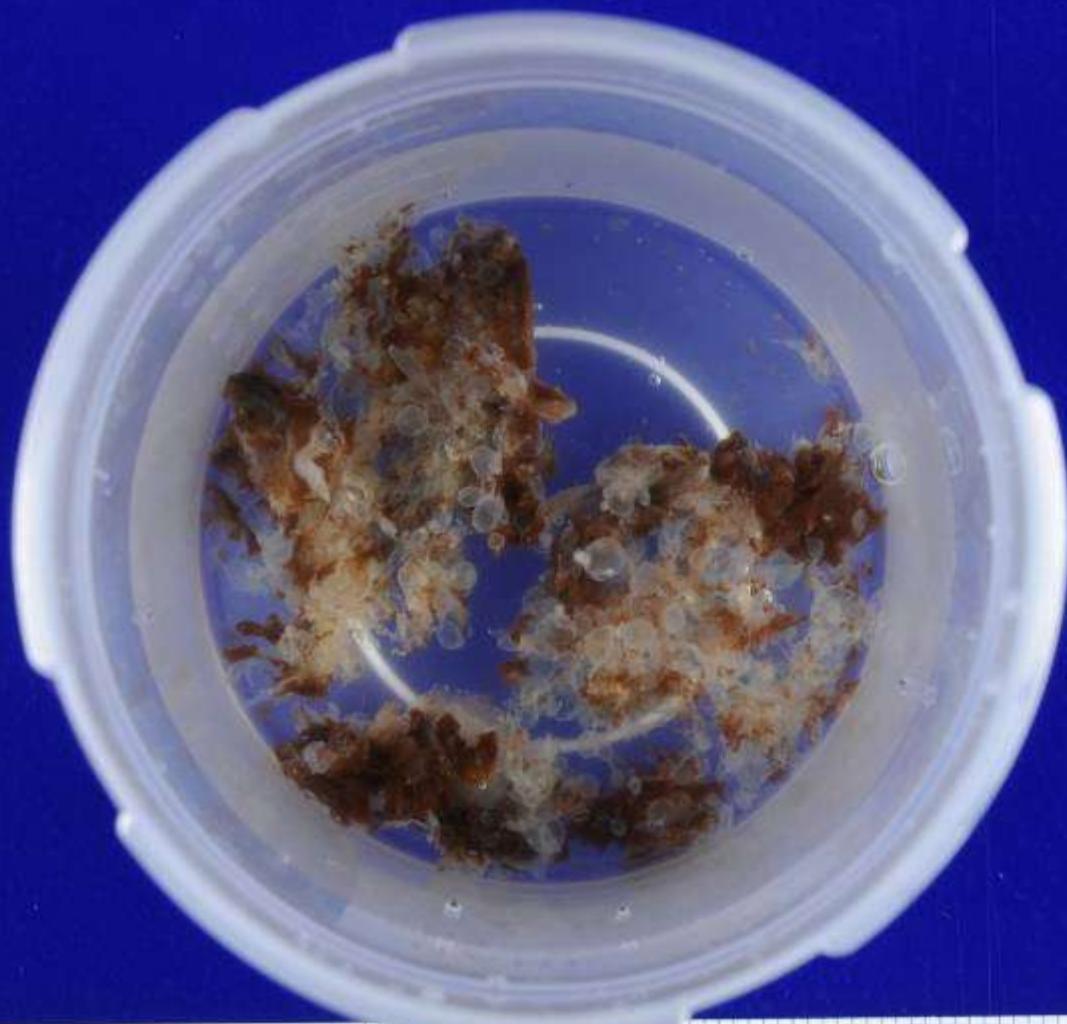
# Mola completa

- Material abundante
- Vesículas visibles
- VelloSIDADES hidrópicas
- Cisternas
- Hiperplasia del trofoblasto
  - Circunferencial
  - Cito y sincitiotrofoblasto



9539 / BII

0 1 2 3 4 5

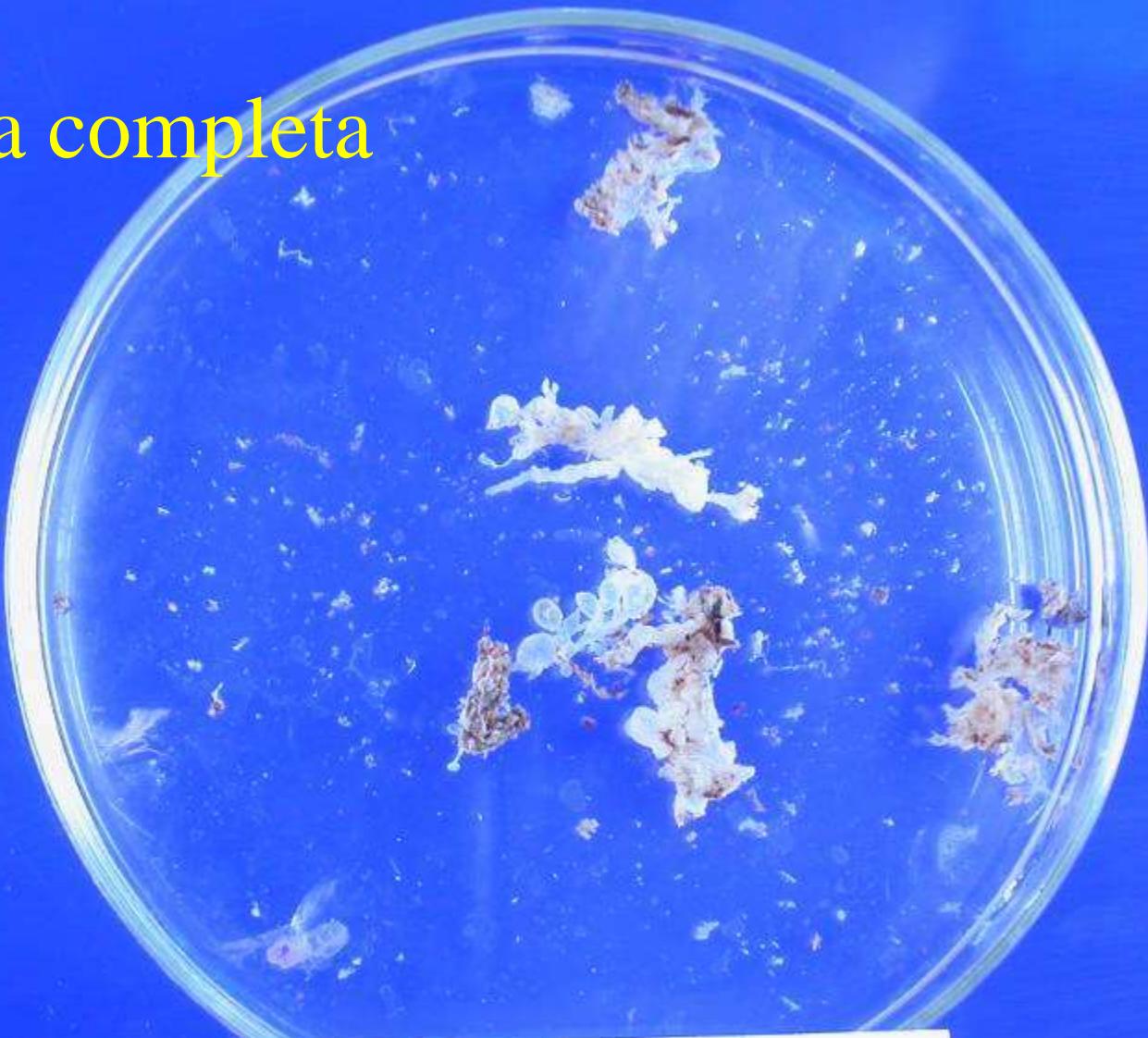


9539 / BII

0 1 2 3 4 5

ANATOMIA PATOLÒGICA. HOSPITAL CLÍNIC. BARCELONA

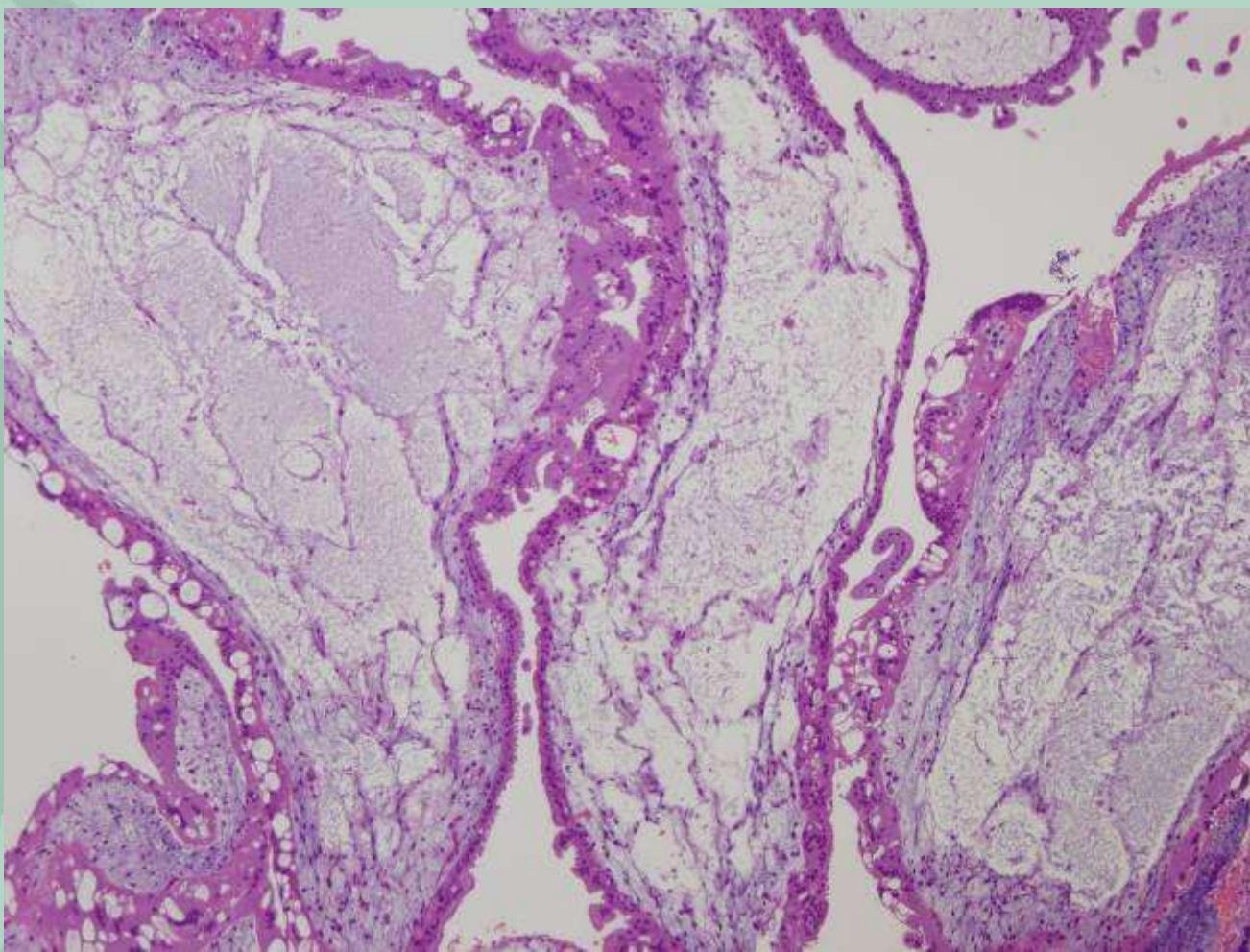
Mola completa

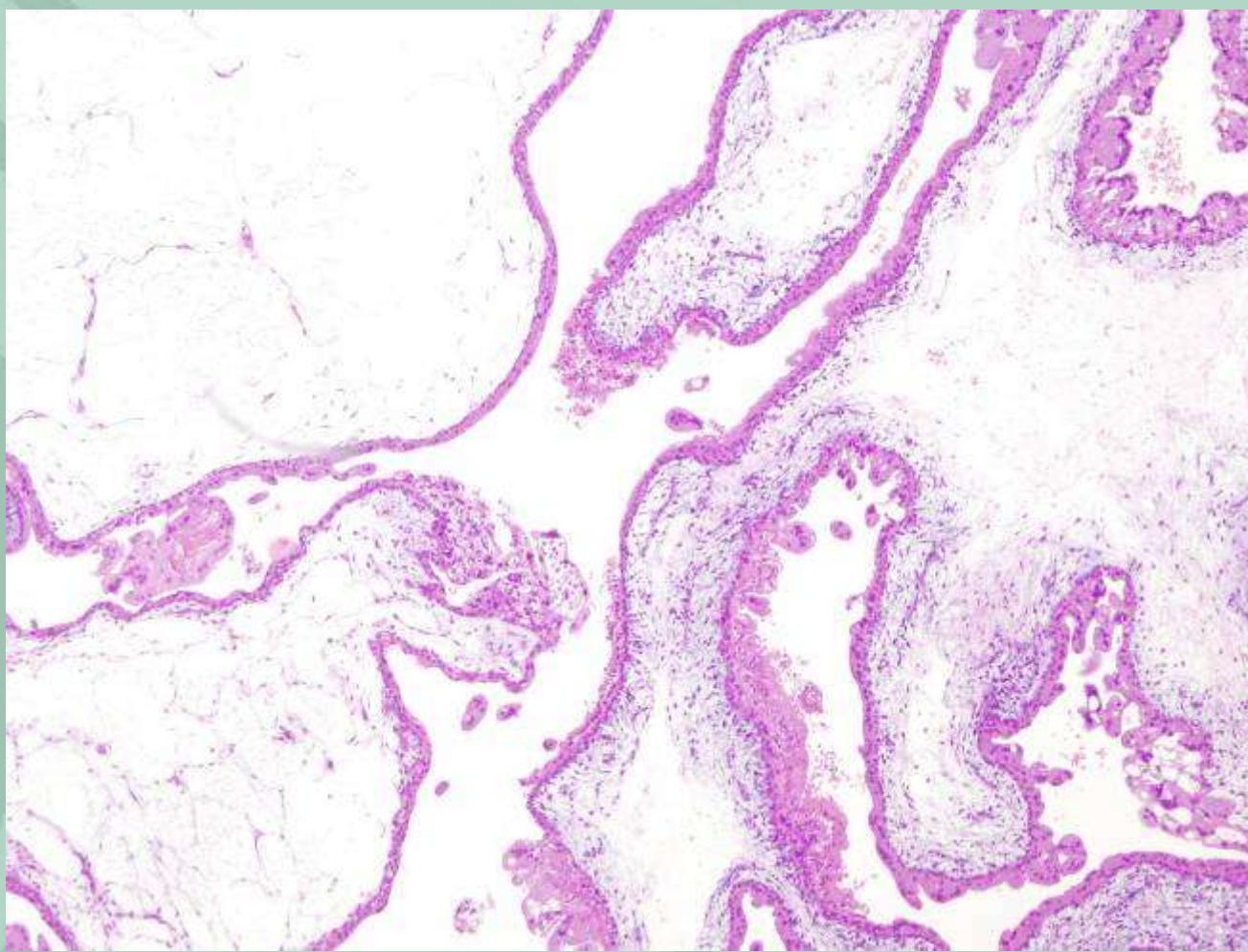


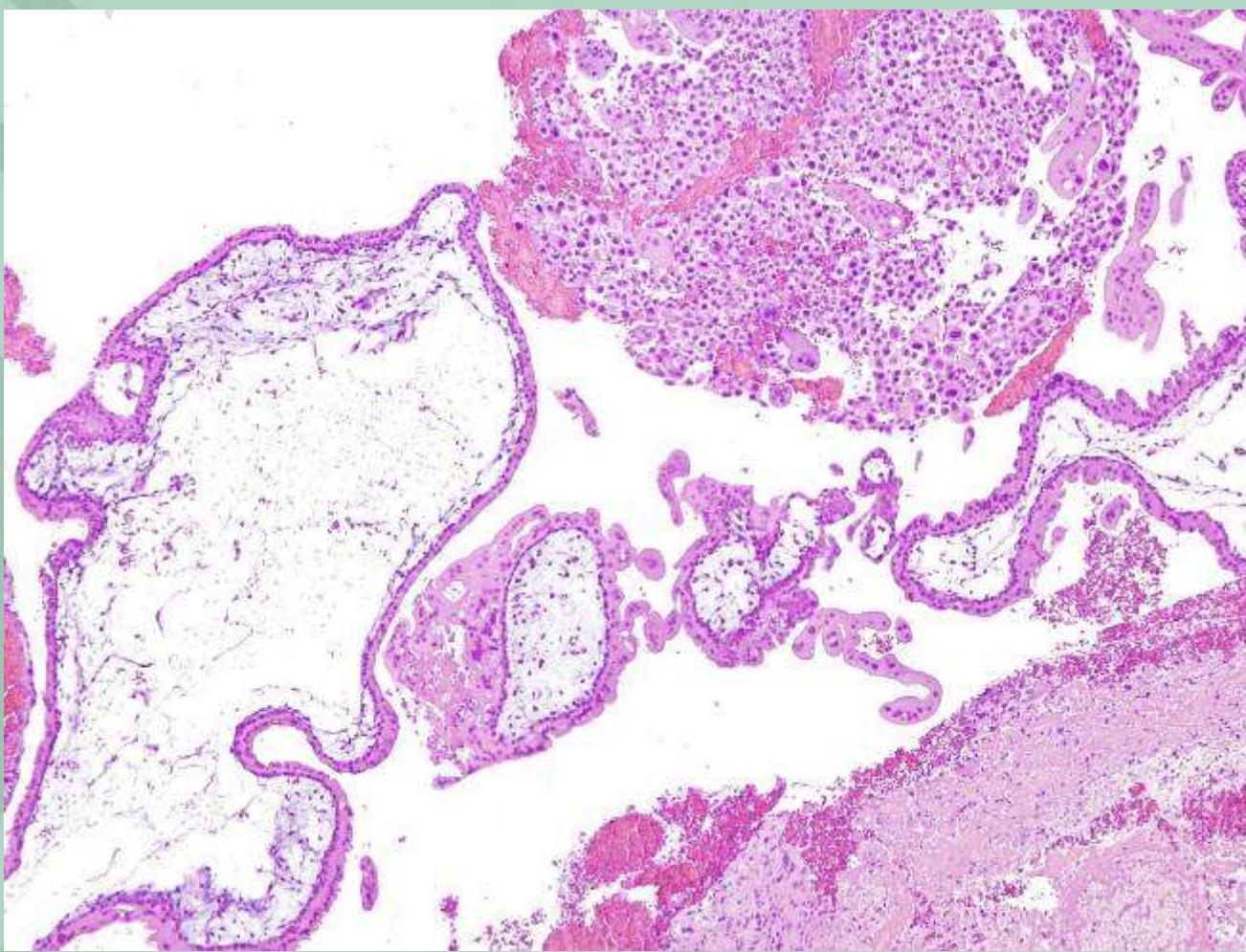
B08.26559



B17083/03





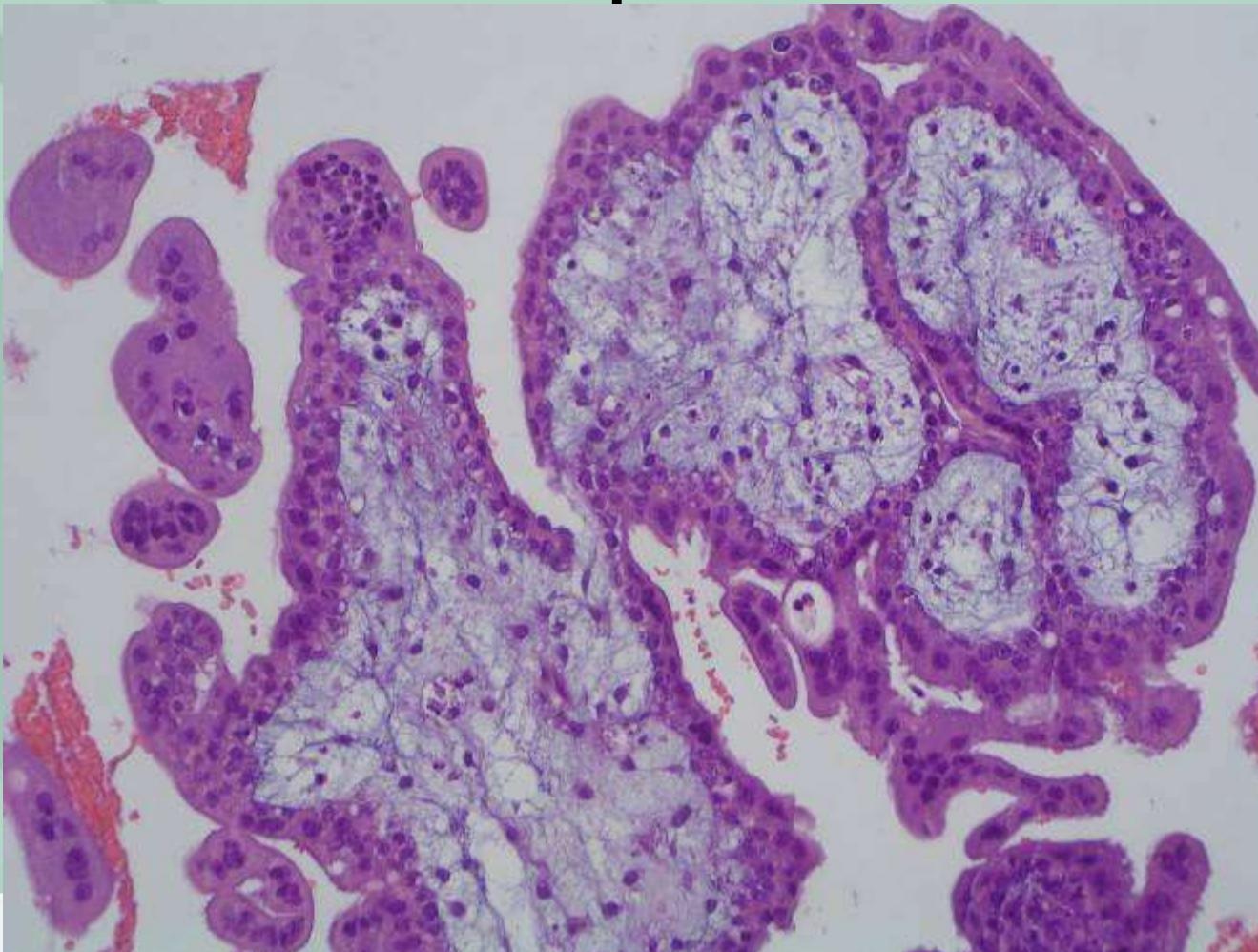




# Mola completa inicial (*early mole*)

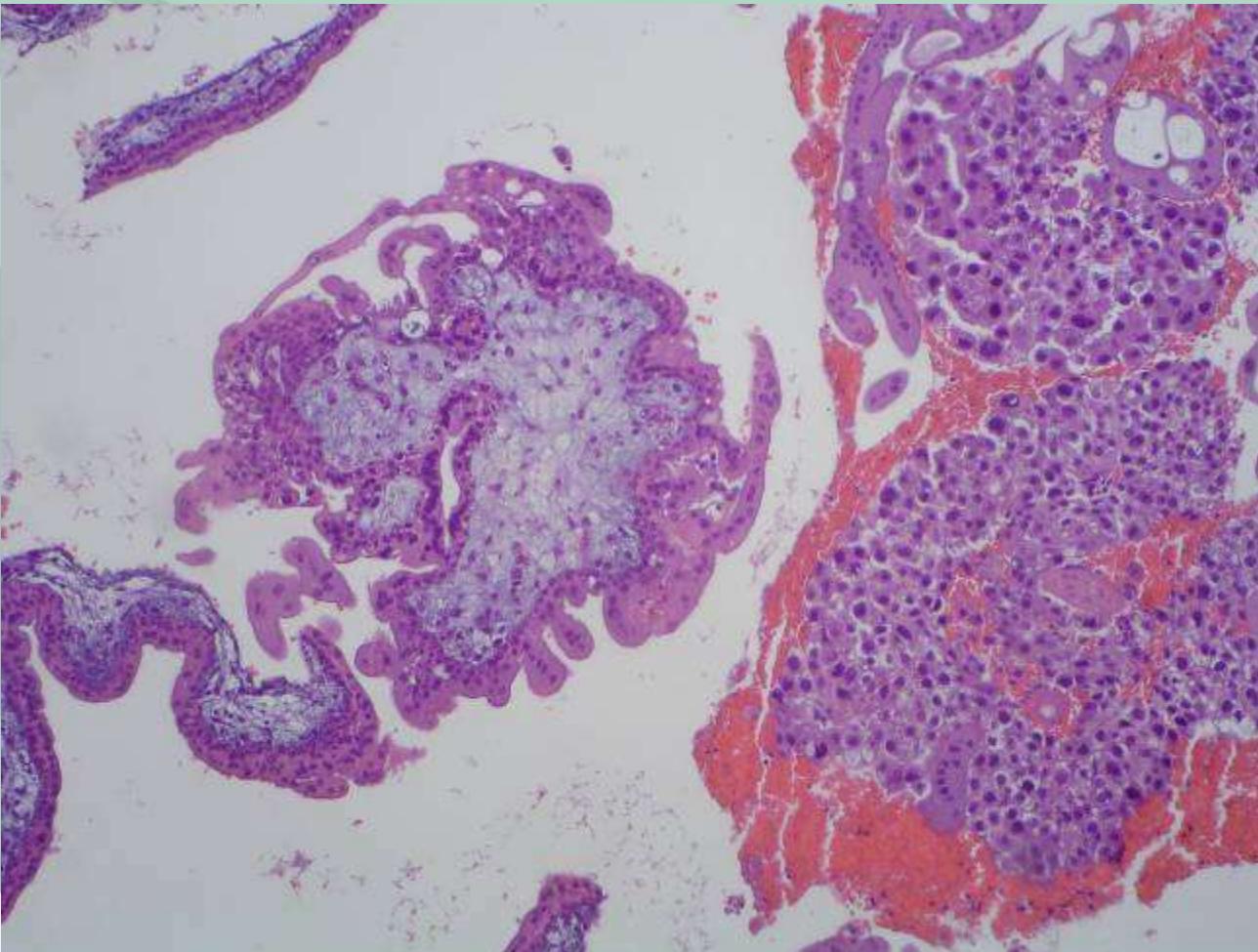
- Edema difuso
- Estroma vellositario azulado
- Hipercelularidad estromal con cariorexis
- Hiperplasia del trofoblasto
- Aspecto de maza

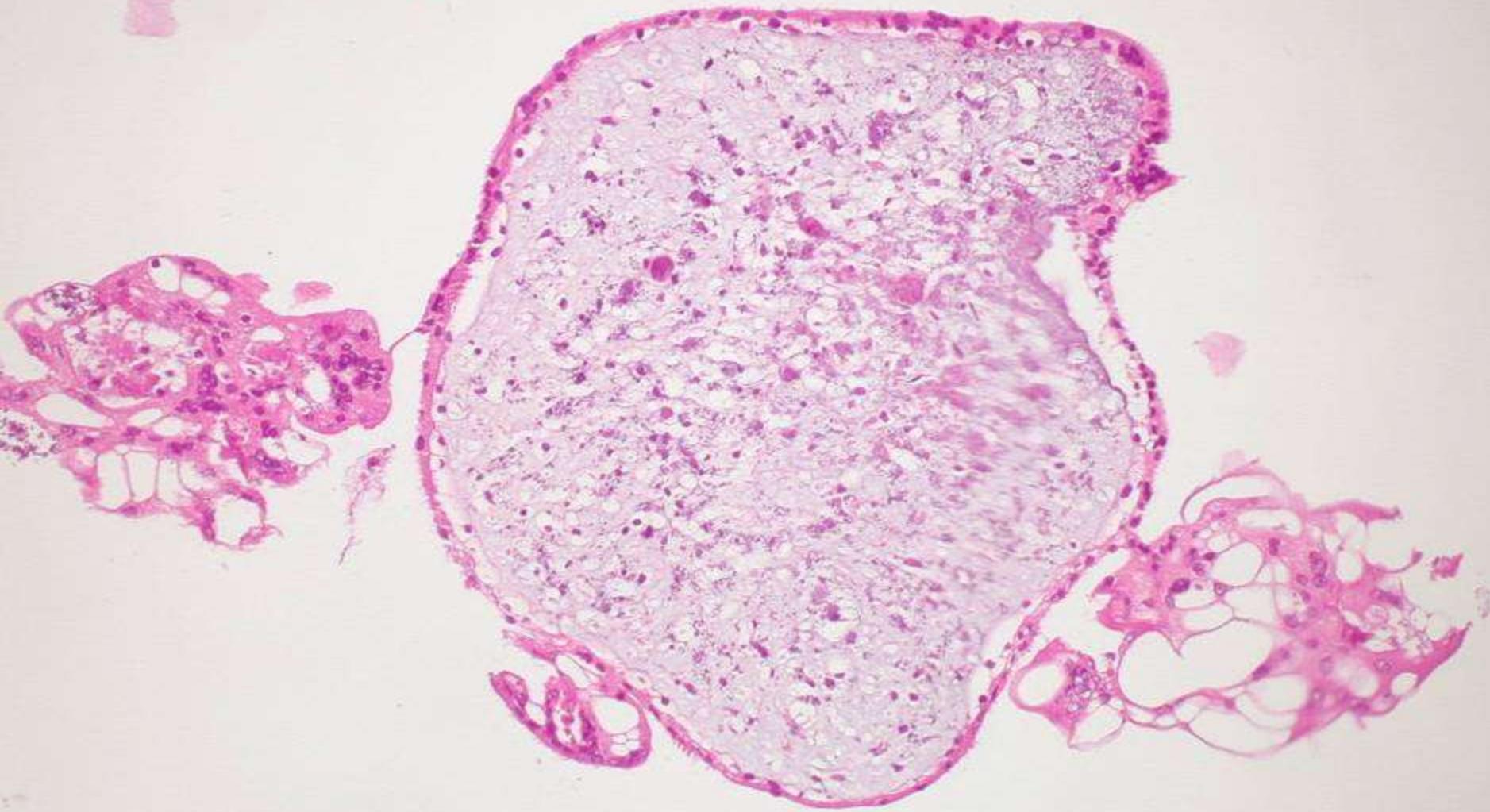
# Mola completa inicial

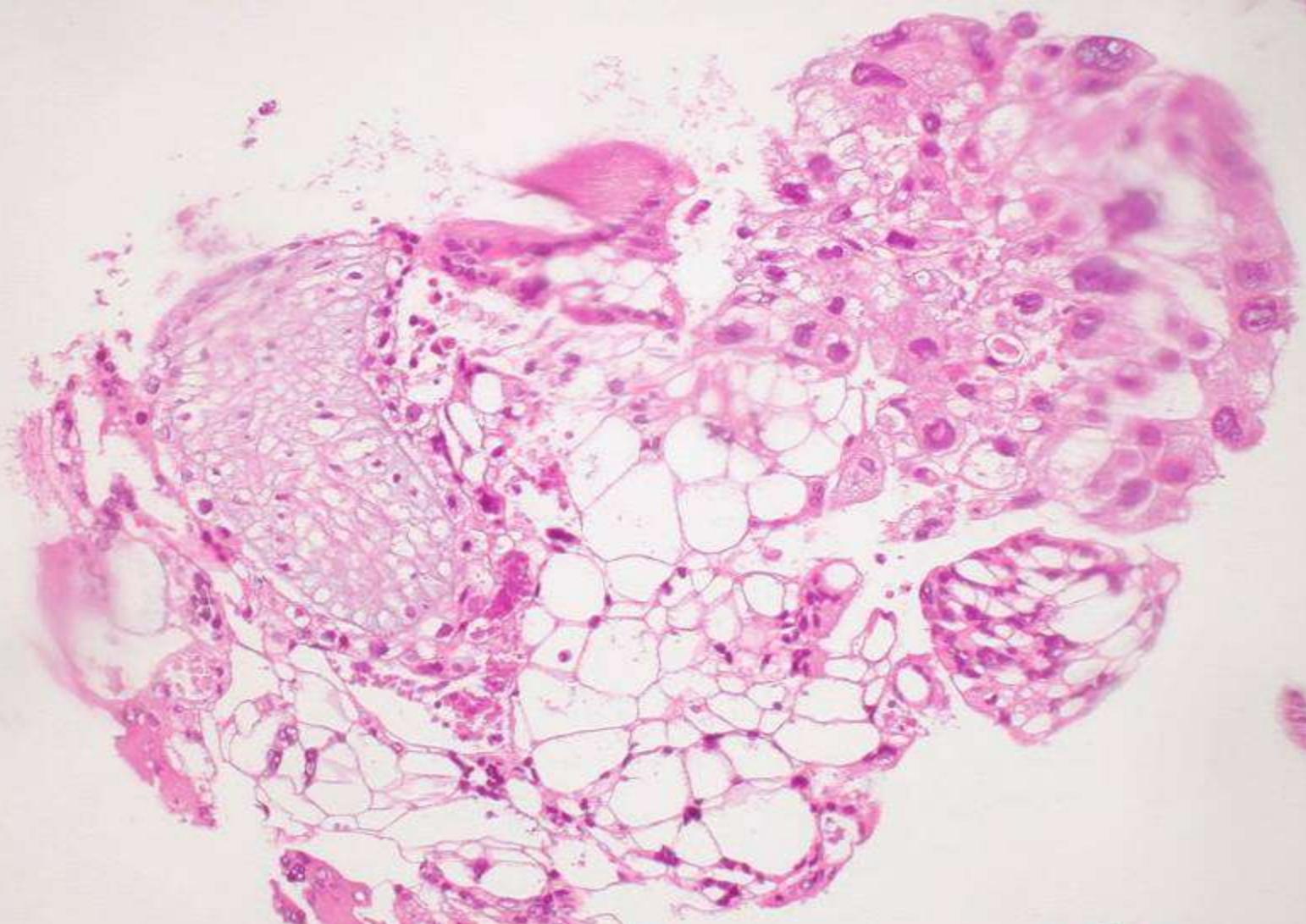




# Mola completa inicial



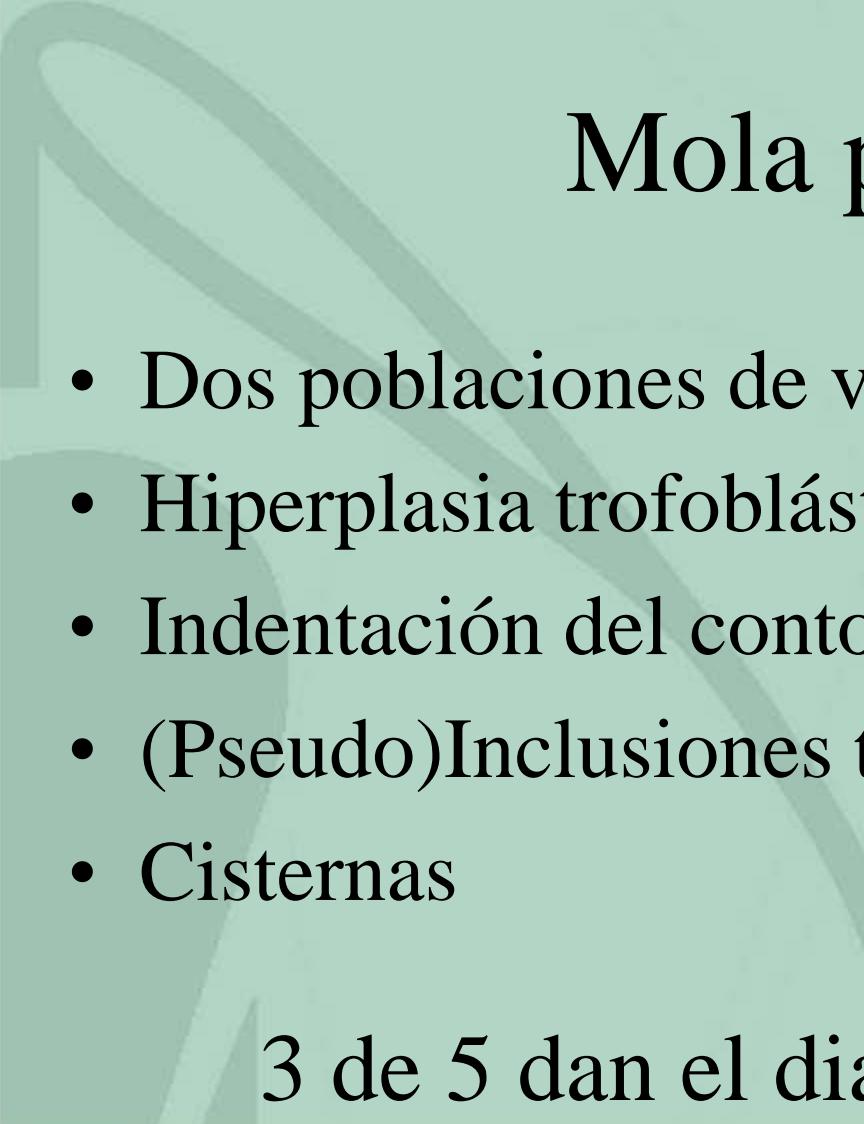




# Mola gemelar







# Mola parcial

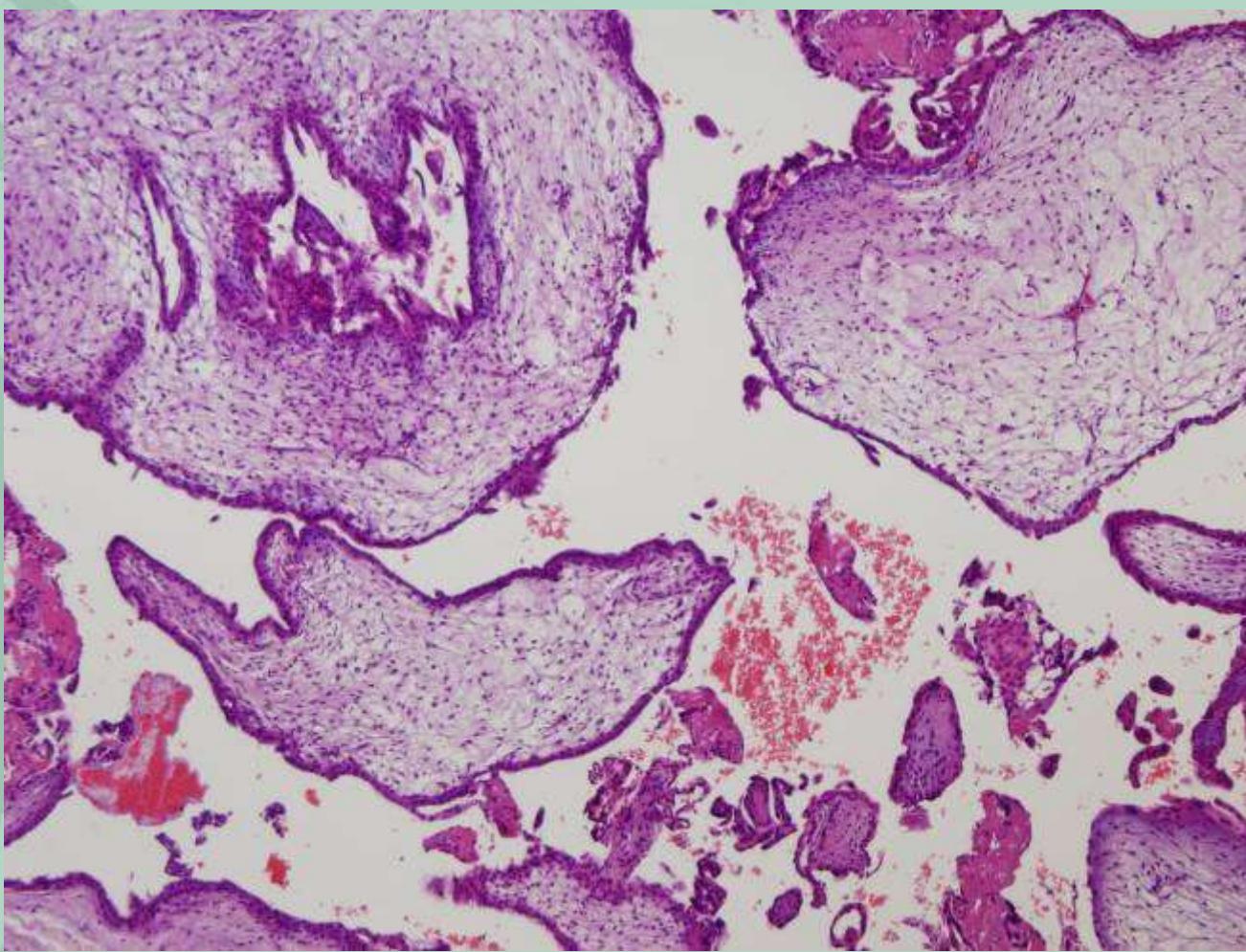
- Dos poblaciones de vellosidades diferentes
- Hiperplasia trofoblástica circunferencial
- Indentación del contorno de las vellosidades
- (Pseudo)Inclusiones trofoblásticas
- Cisternas

3 de 5 dan el diagnóstico de MHP

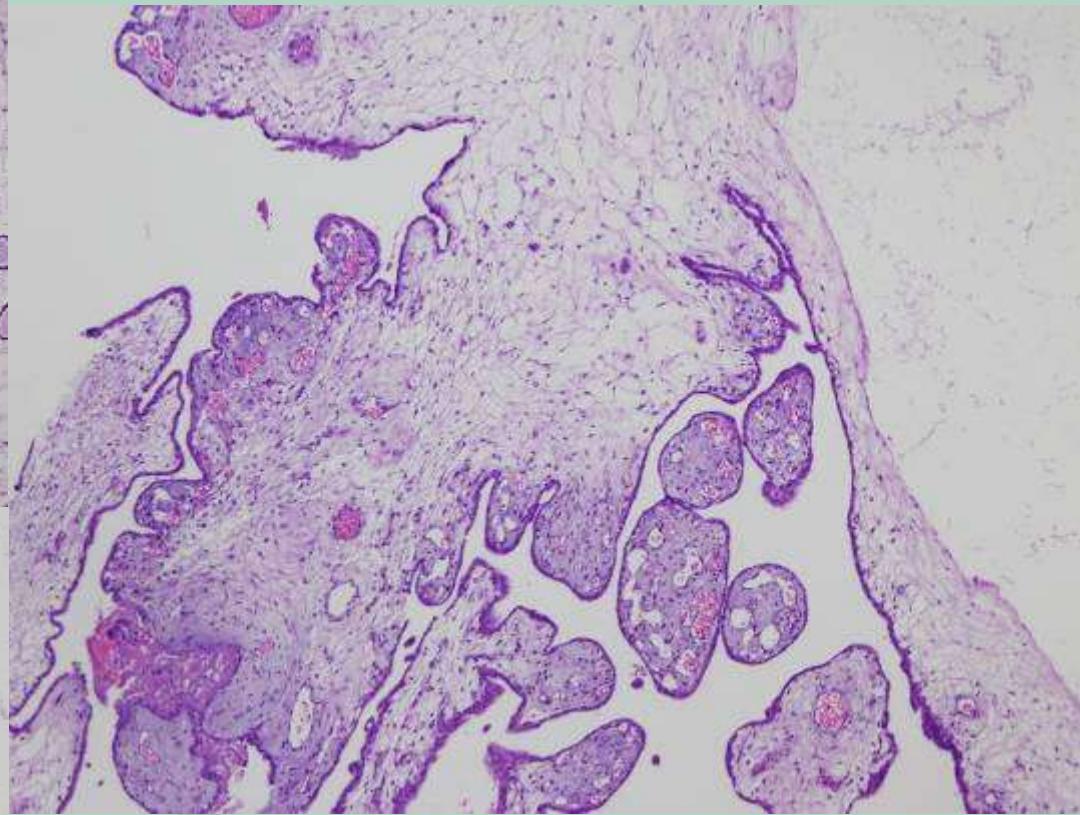
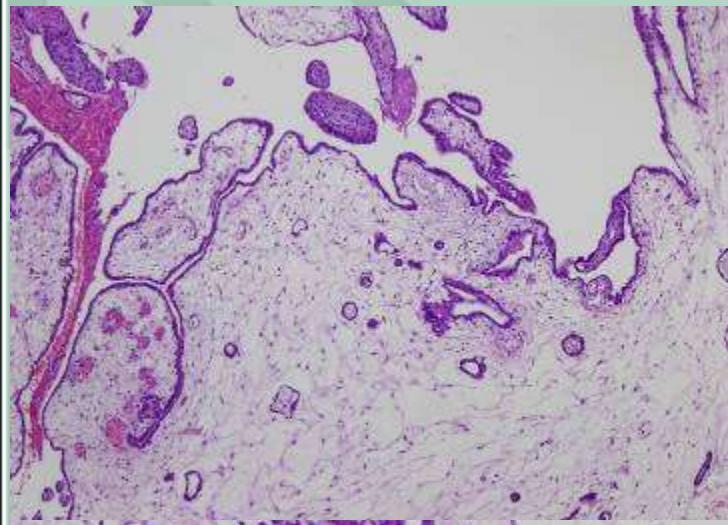


# Mola parcial





# MHP



16 semanas



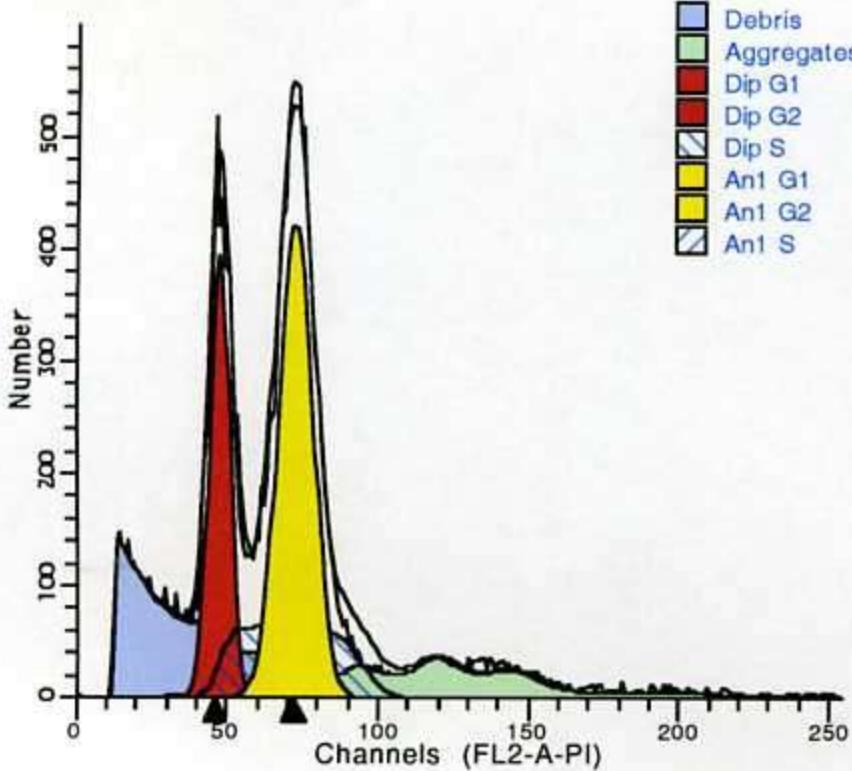
# Mola parcial inicial

	Early partial mole n=80 (%)	Late partial mole n=20 (%)
Villous edema	80 (100)	20 (100)
Cistern formation	79 (99)	20 (100)
Focal syncytiotrophoblastic hyperplasia	80 (100)	20 (100)
Villous scalloping	80 (100)	20 (100)
Trophoblastic inclusion	79 (99)	20 (100)
Nucleated red blood cells	57 (71)	15 (75)
Fetal parts	15 (19)	3 (15)
Fetal blood vessels	67 (84)	19 (95)
Extensive stromal fibrosis	2 (3)§	9 (45)§
Wandering trophoblast	32 (40)	9 (45)
Maximal size of villi (mm) <sup>a</sup>	1.2–6.2	1.5–6.5
Mean size of villi (mm) <sup>a</sup>	3.0 §§	3.4 §§

+ Dimorfismo vellositario

MHP inicial = MHP clásica

Fukunaga Virchows Arch (2000) 437:180-4



Debris  
Aggregates  
Dip G1  
Dip G2  
Dip S  
An1 G1  
An1 G2  
An1 S

File analyzed: B09\_06105-3.034  
Date analyzed: 3-Apr-2009  
Model: 2DA0n\_DSD ASF  
Analysis type: Manual analysis

Diploid: 50.74 %  
Dip G1: 52.35 % at 47.17  
Dip G2: 0.00 % at 94.35  
Dip S: 47.65 % G2/G1: 2.00  
%CV: 7.03

Aneuploid 1: 49.26 %  
An1 G1: 99.90 % at 72.41  
An1 G2: 0.00 % at 96.69  
An1 S: 0.10 % G2/G1: 1.34  
%CV: 7.97 DI: 1.53

Total Aneuploid S-Phase: 0.10 %  
Total S-Phase: 24.22 %  
Total B.A.D.: 16.39 %

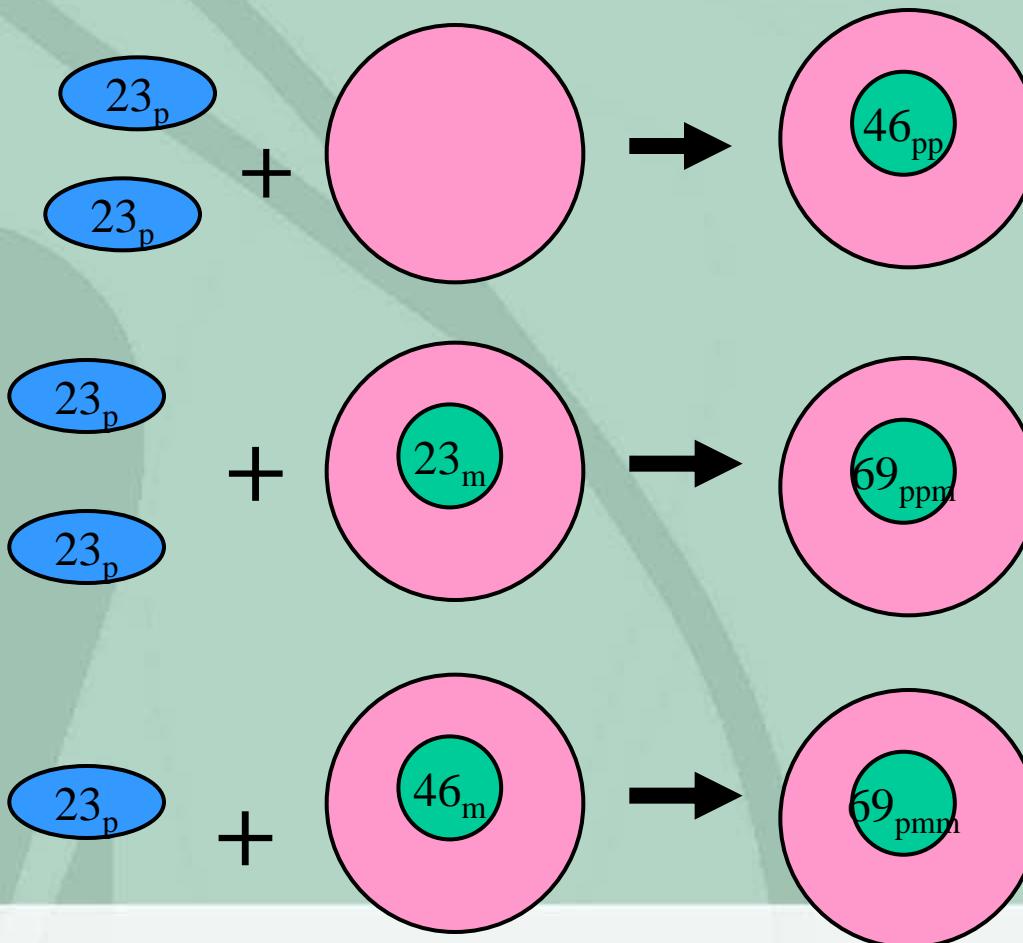
Debris: 23.75 %  
Aggregates: 12.96 %  
Modeled events: 19591  
All cycle events: 12398  
Cycle events per channel: 245  
RCS: 3.268

# Triploidia digínica





# Aneuploidias



Diploidia diandrica  
**MC**

Placenta sin embrión

Triploidia diandrica  
**MP**

Placenta predomina  
sobre embrión

Triploidia diginica  
Hipoplasia placentaria



# Origen de la triploidia y fenotipo

**Table 5**

**Parental Origin and Phenotype of Triploidy**

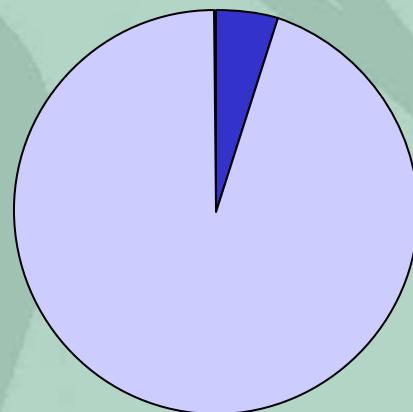
ORIGIN	TOTAL No. OF CASES	GROSS PHENOTYPIC FEATURES			DEVELOPMENTAL STAGE (wk)			
		Partial Mole	Trophoblastic Hyperplasia	Fetal Tissue	<6.0	6.0–8.5	8.5–11.5	>11.5
Maternal	27	0 (0%)	3 (11%)	16 (59%)	6	12	6	3
Paternal	58	33 (57%)	46 (79%)	13 (22%)	3	12	25	18

NOTE.—Includes cases in which results from both the DNA analysis and histological study were available.



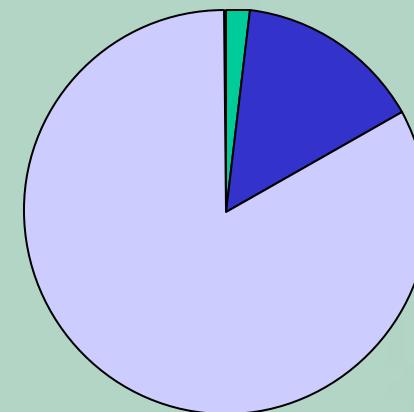
# Evolución de la mola hidatiforme

**Mola parcial**



- Coriocarcinoma
- E. persistente
- Normal

**Mola completa**



- Coriocarcinoma
- E. persistente
- Normal



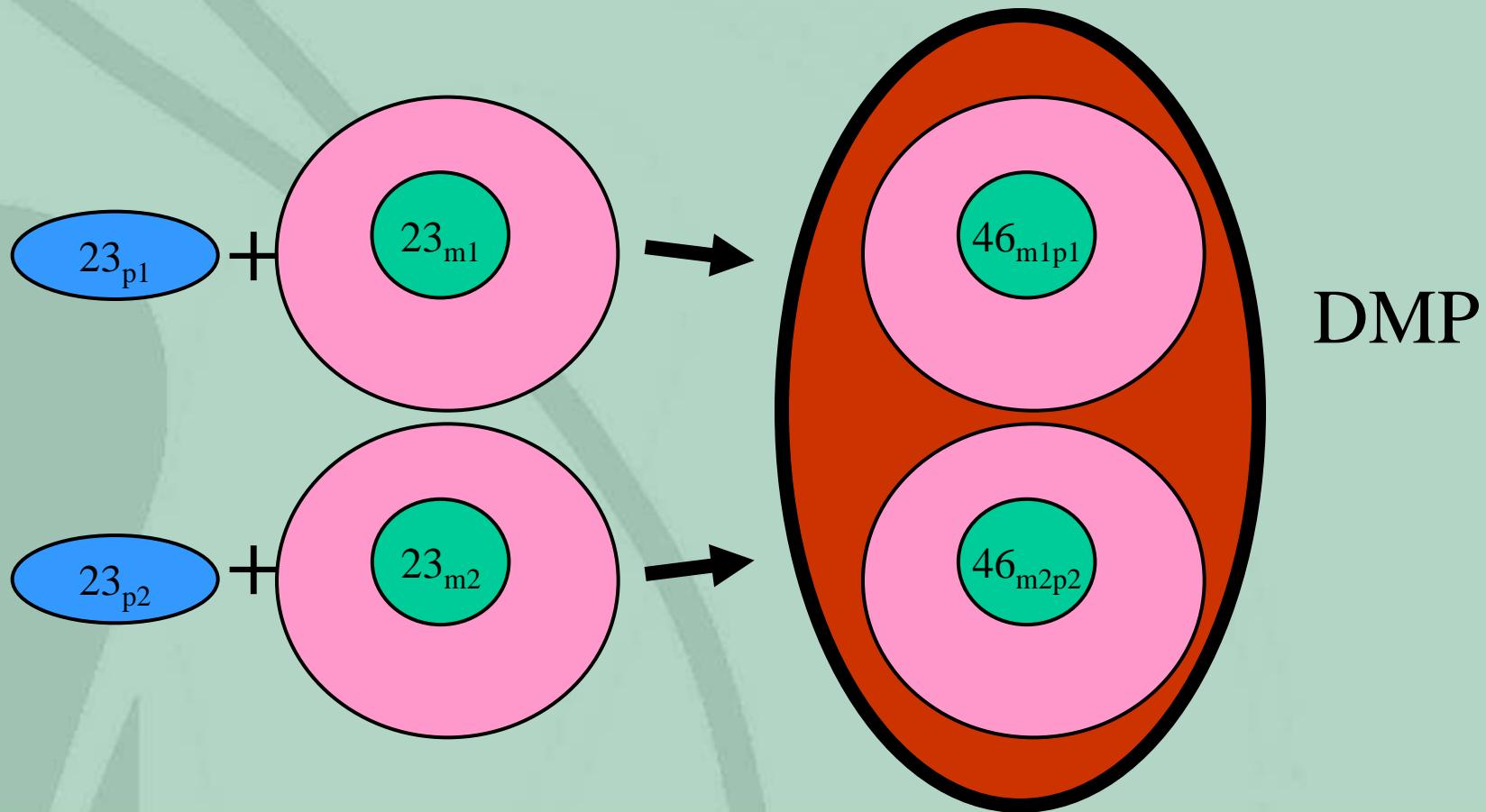
# Enfermedad persistente

- Mola persistente
- Mola invasora
- Mola metastática
- Coriocarcinoma





# Quimerismo



# Displasia mesenquimática

- Placentomegalia con anomalías de las vellosidades principales
- Dilatación quística
- Vesículas
- Proliferación fibroblástica estromal
- Anomalías en cualquier tipo de vaso
- Población de vellosidades normales



U.DIAGNOSTIC PRENATAL-ICGON-C.S.CLINIC OB/GINE



U.DIAGNOSTIC PRENATAL-ICGON-C.S.CLINIC OB/GINE

11



U.DIAGNOSTIC PRENATAL-ICGON-C.S.CLINIC OB/GINE

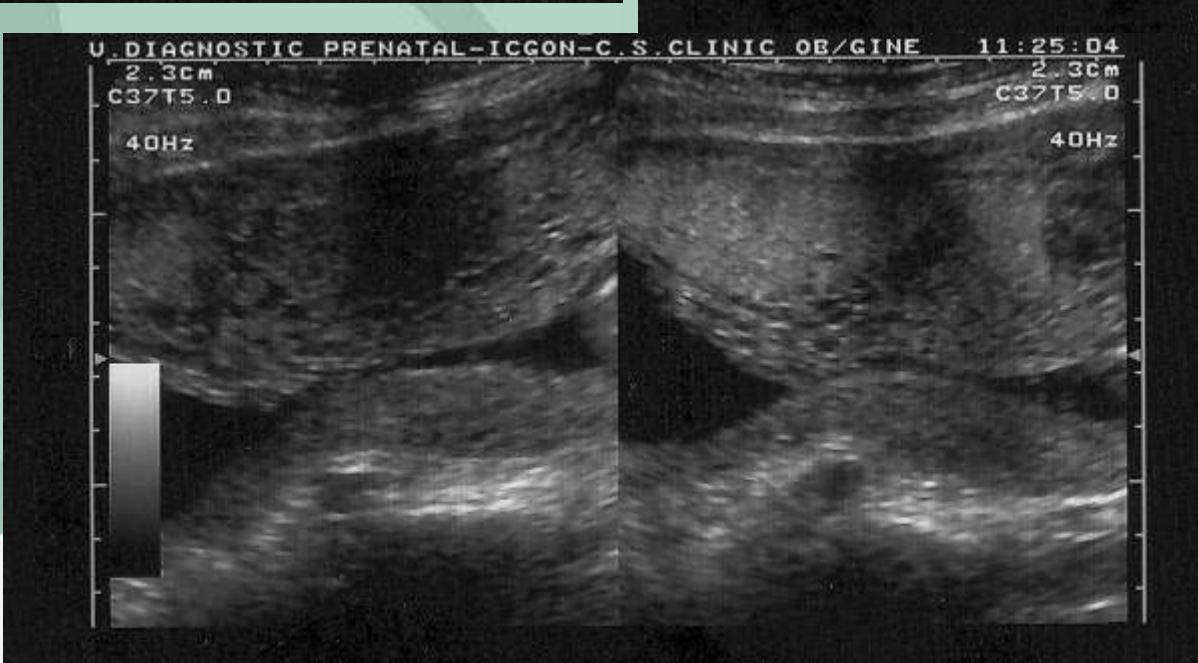
11:25:04

2.3cm  
C37T5.0

2.3cm  
C37T5.0

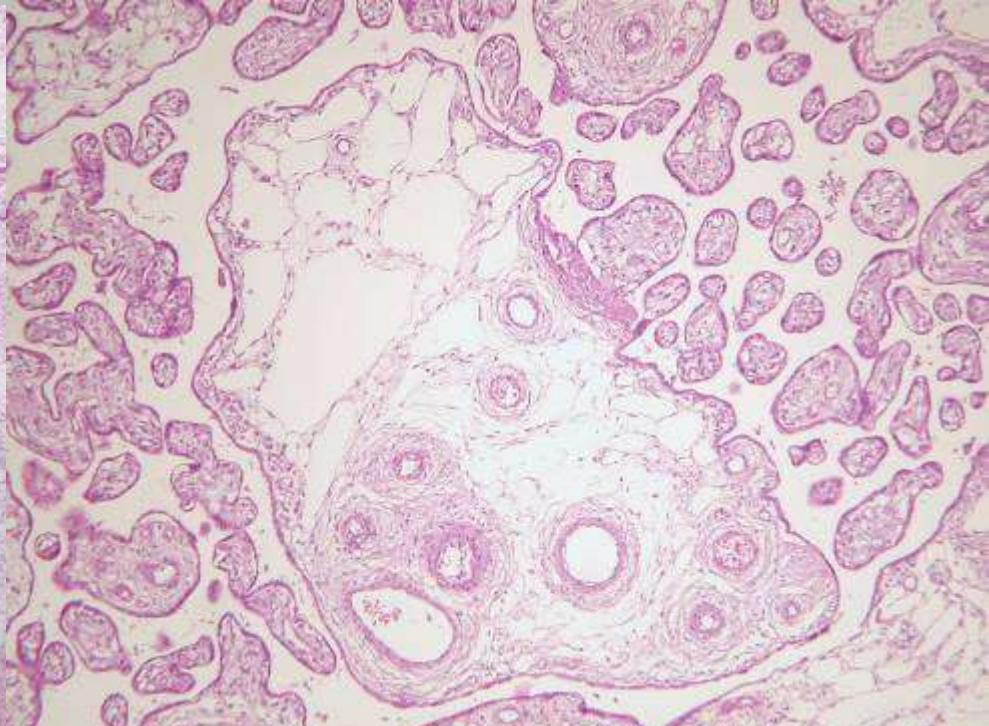
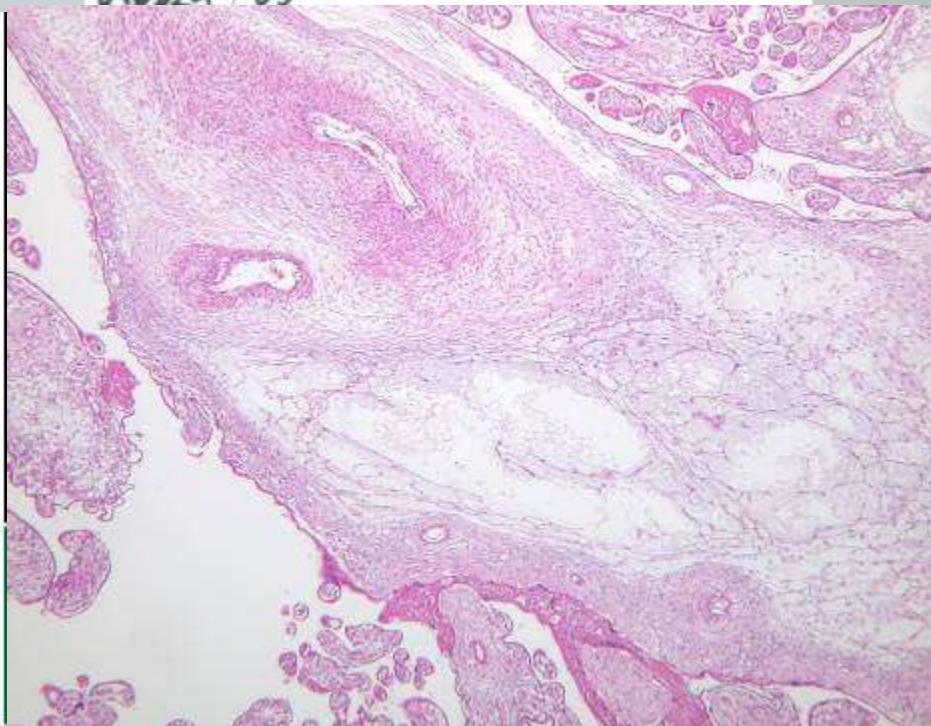
40Hz

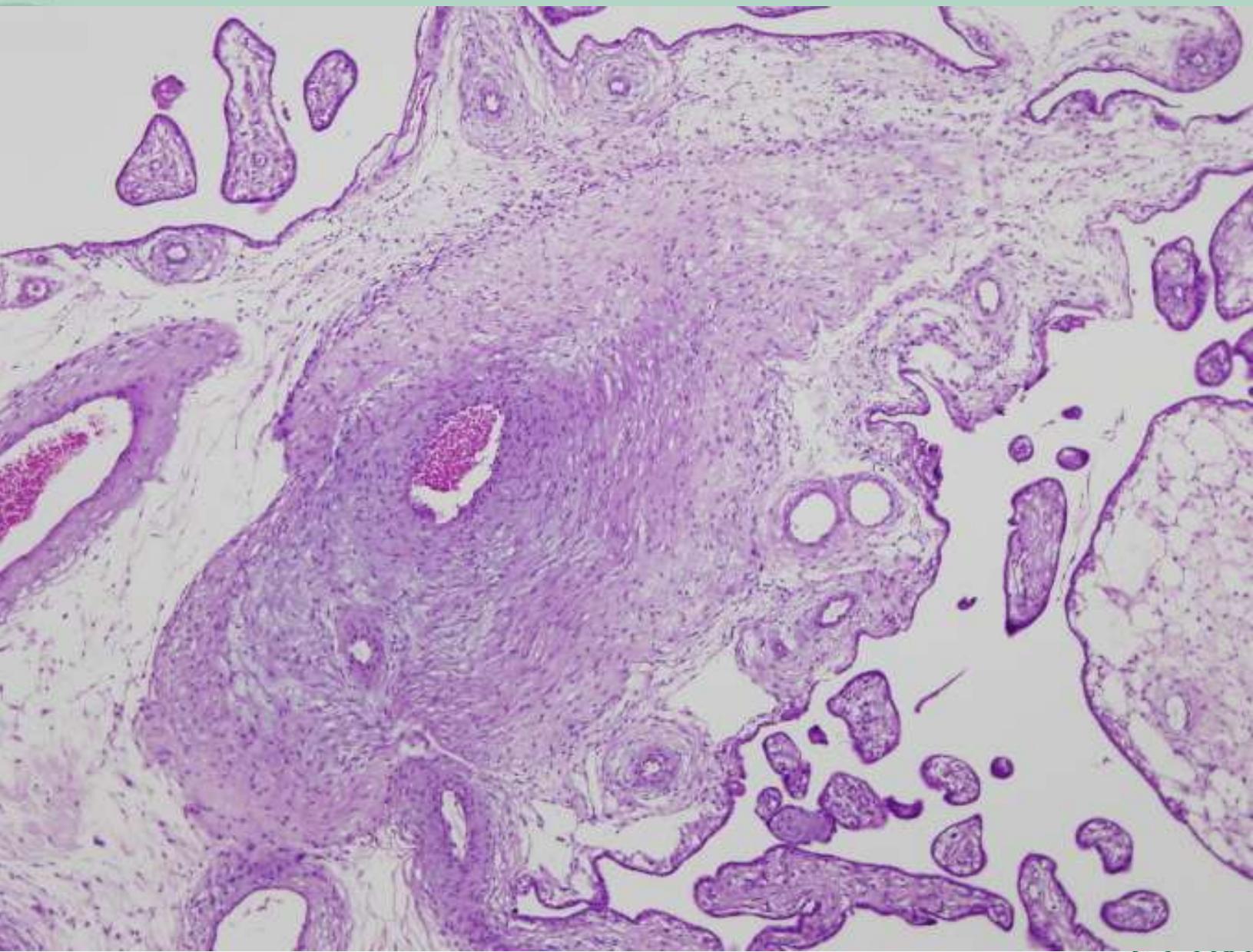
40Hz

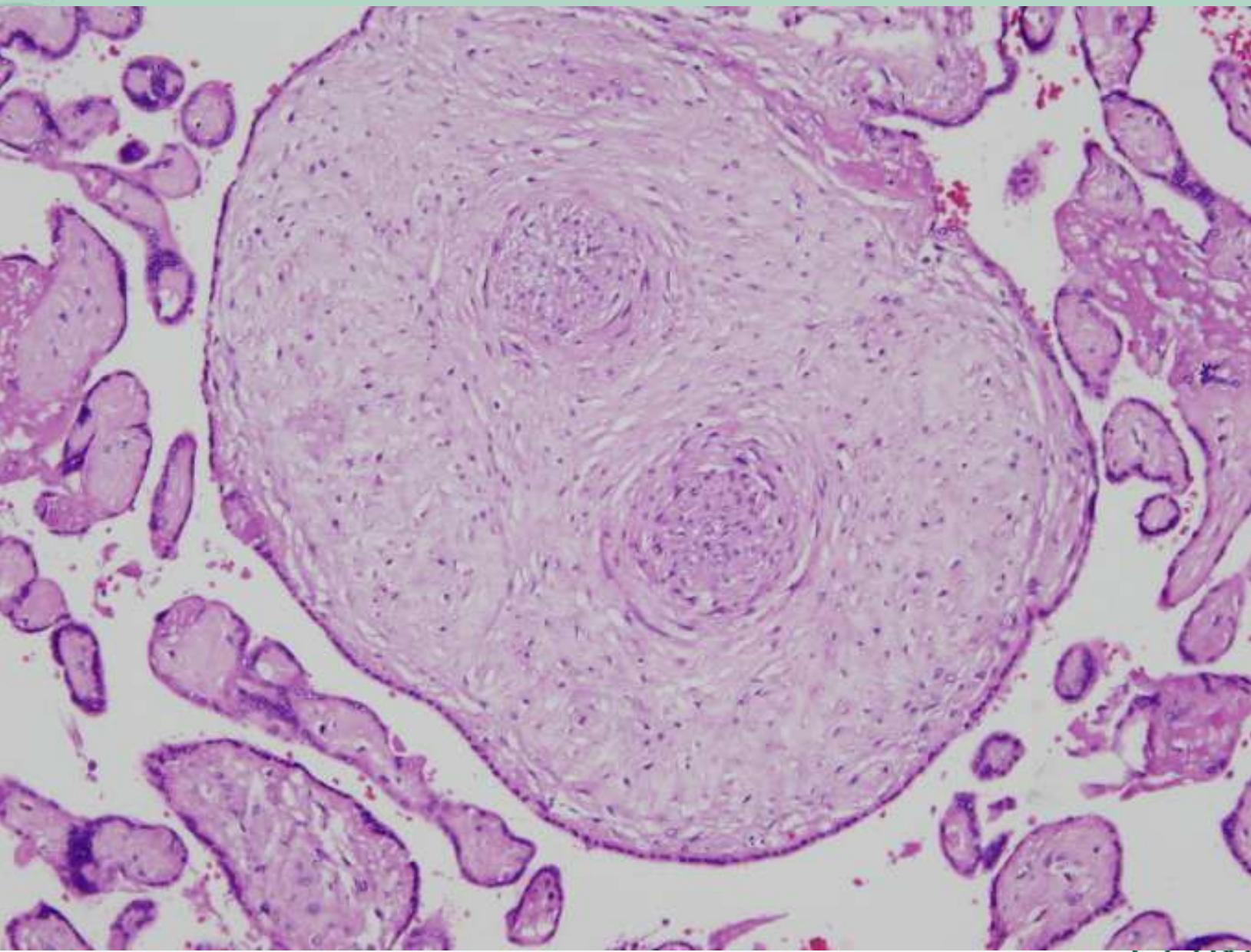


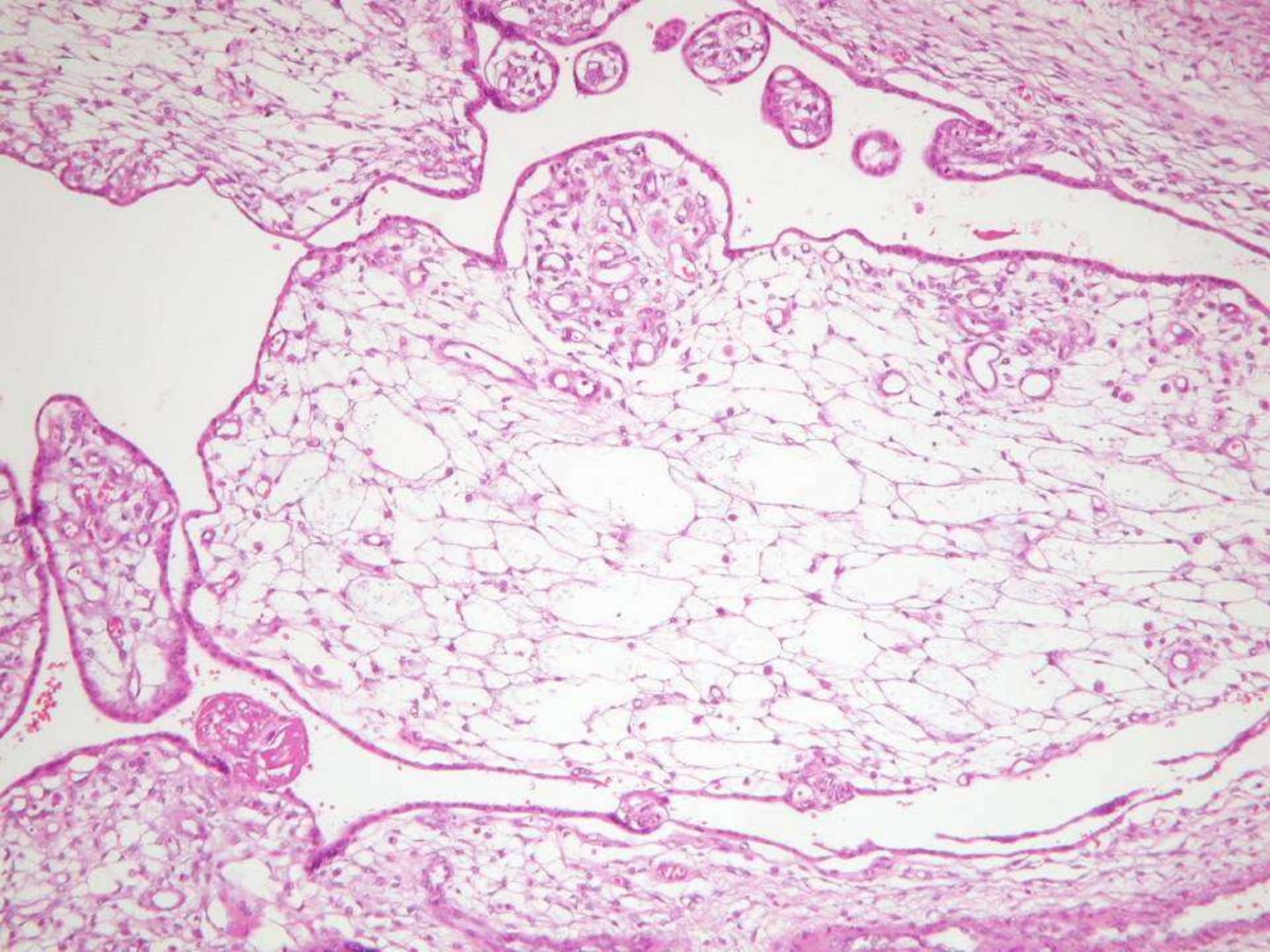


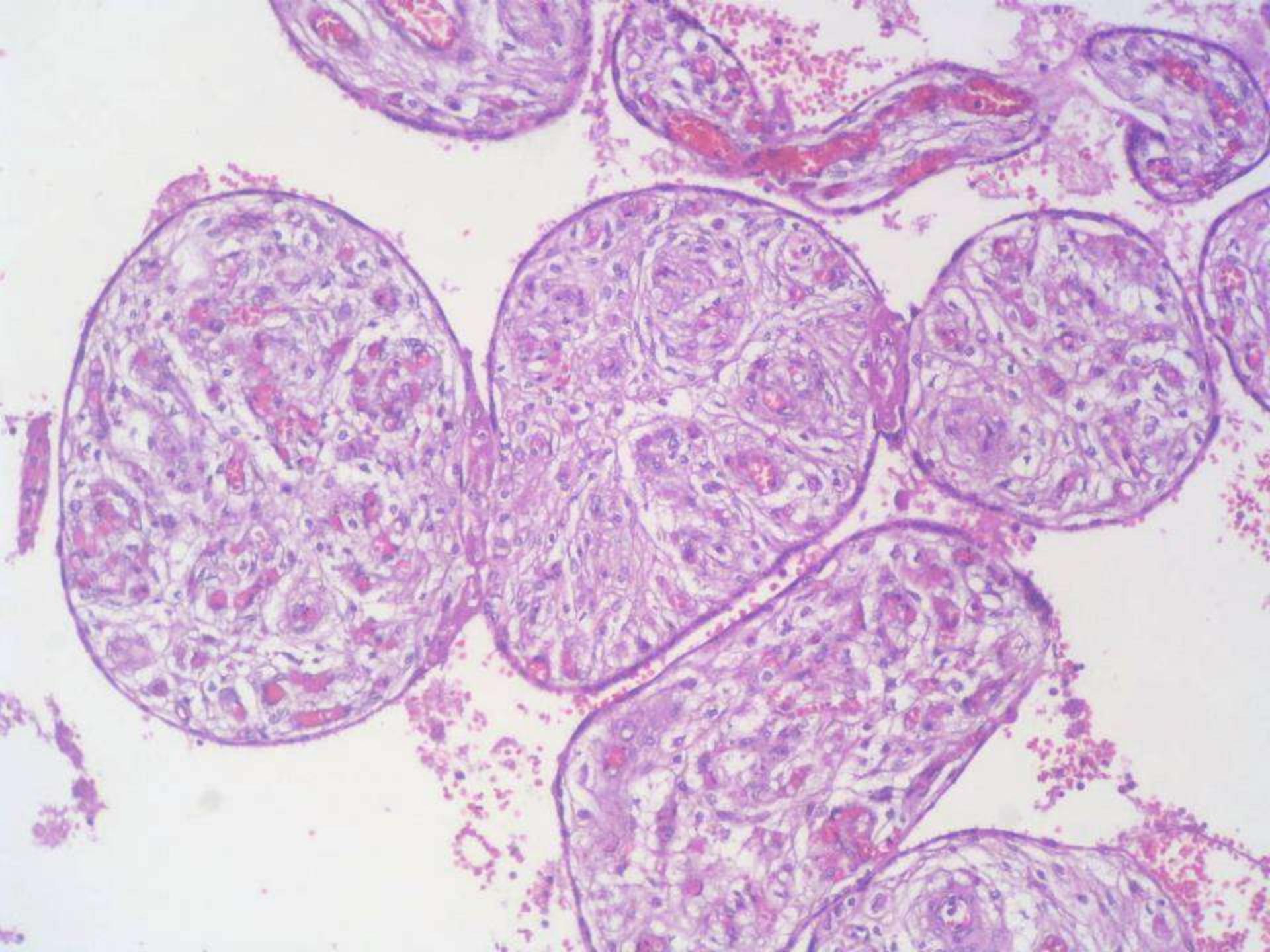
# Displasia mesenquimática

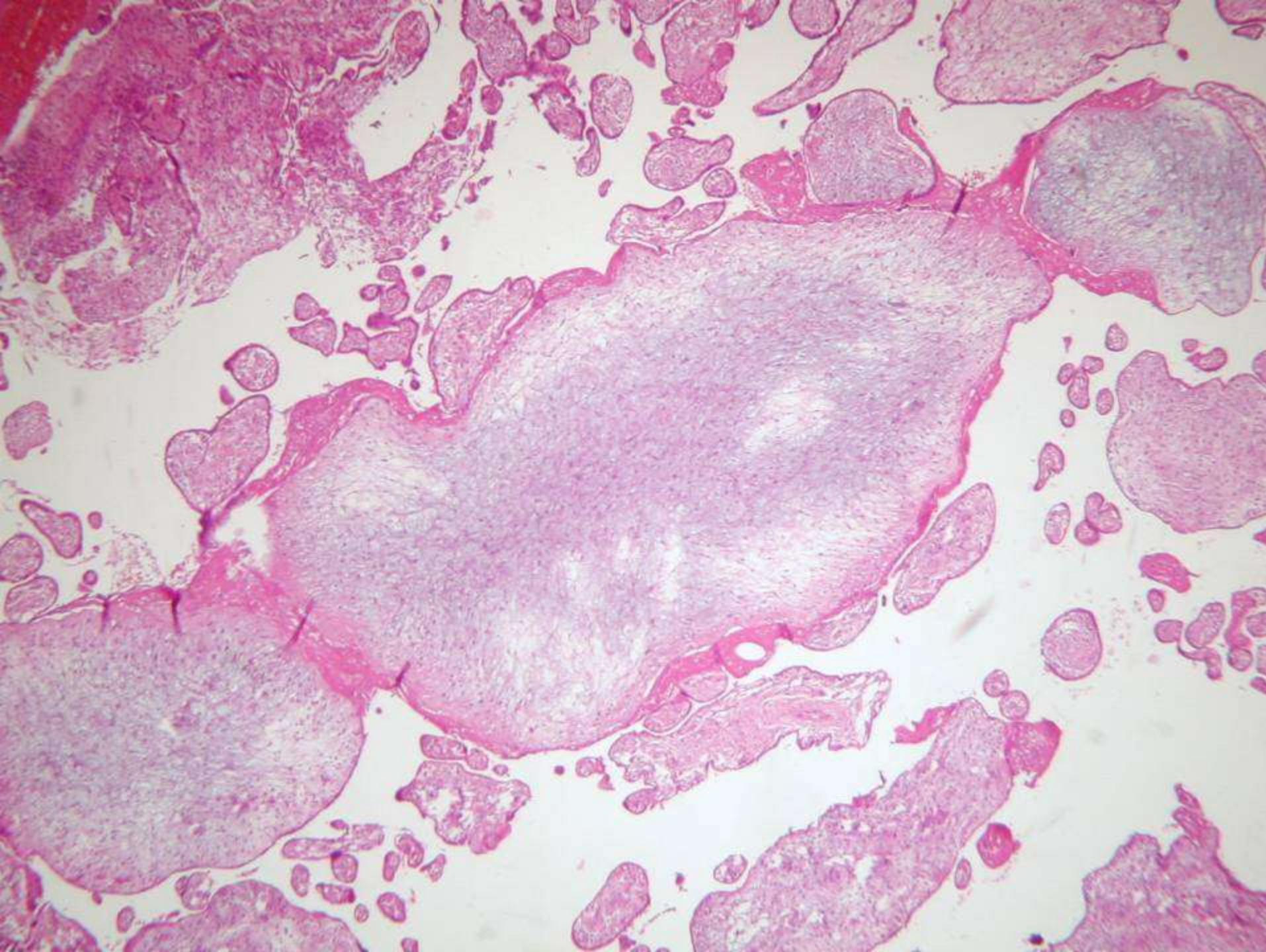










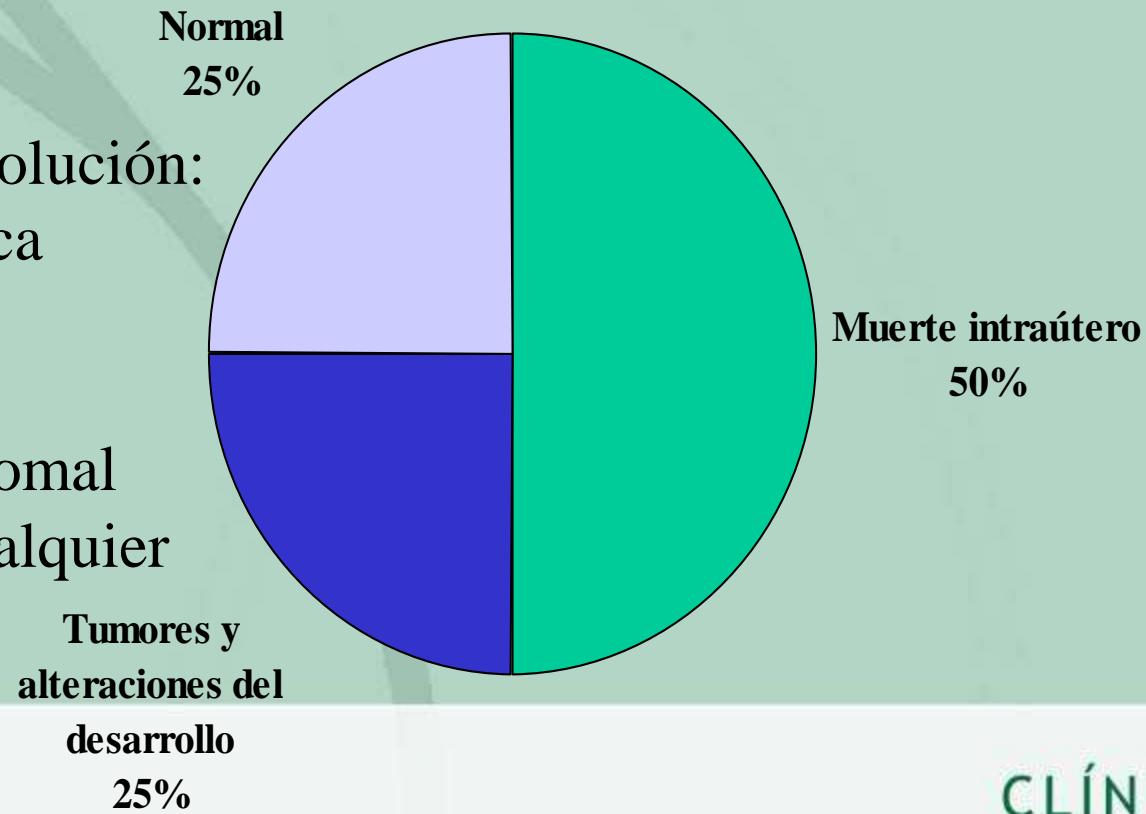


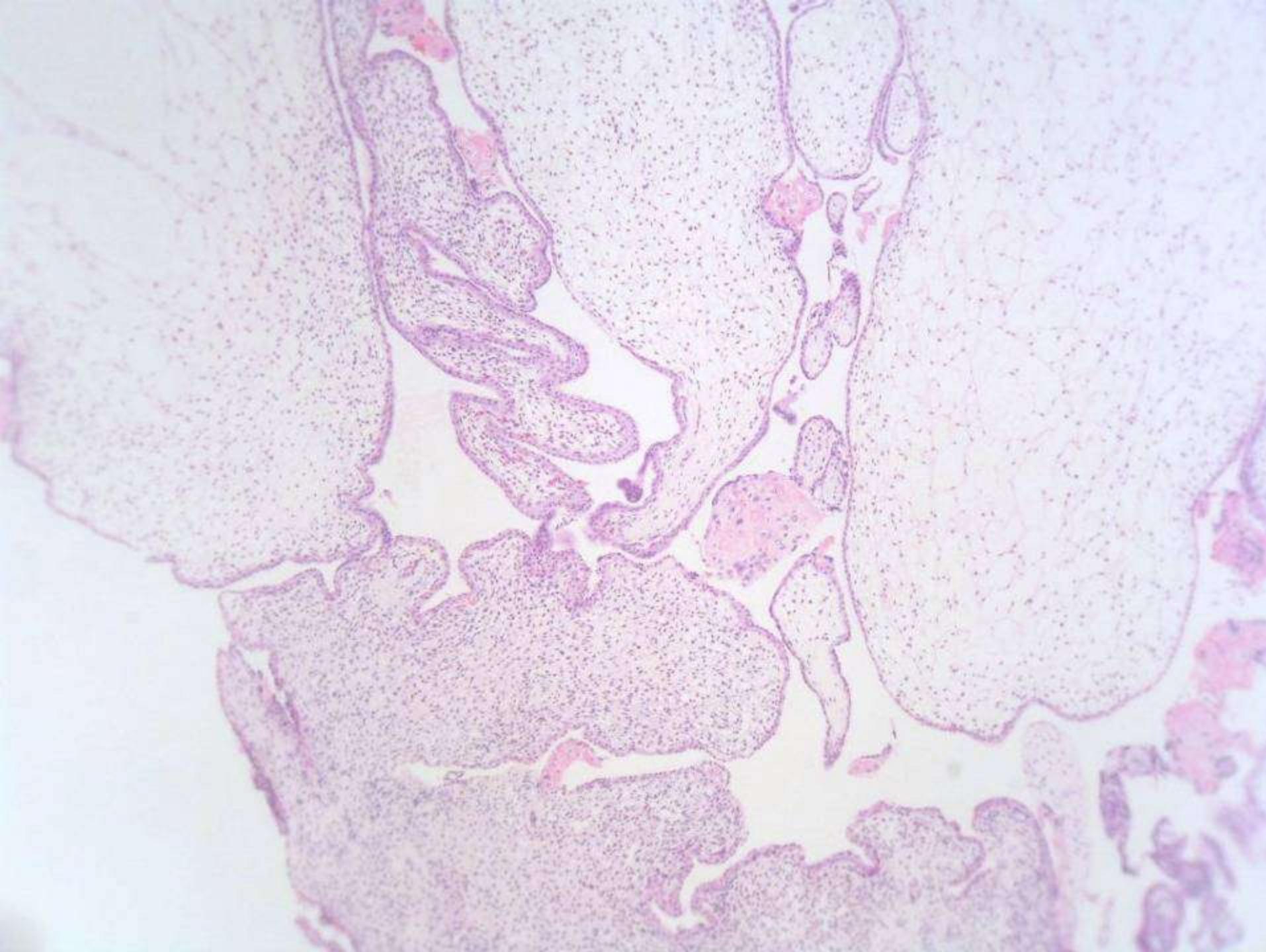


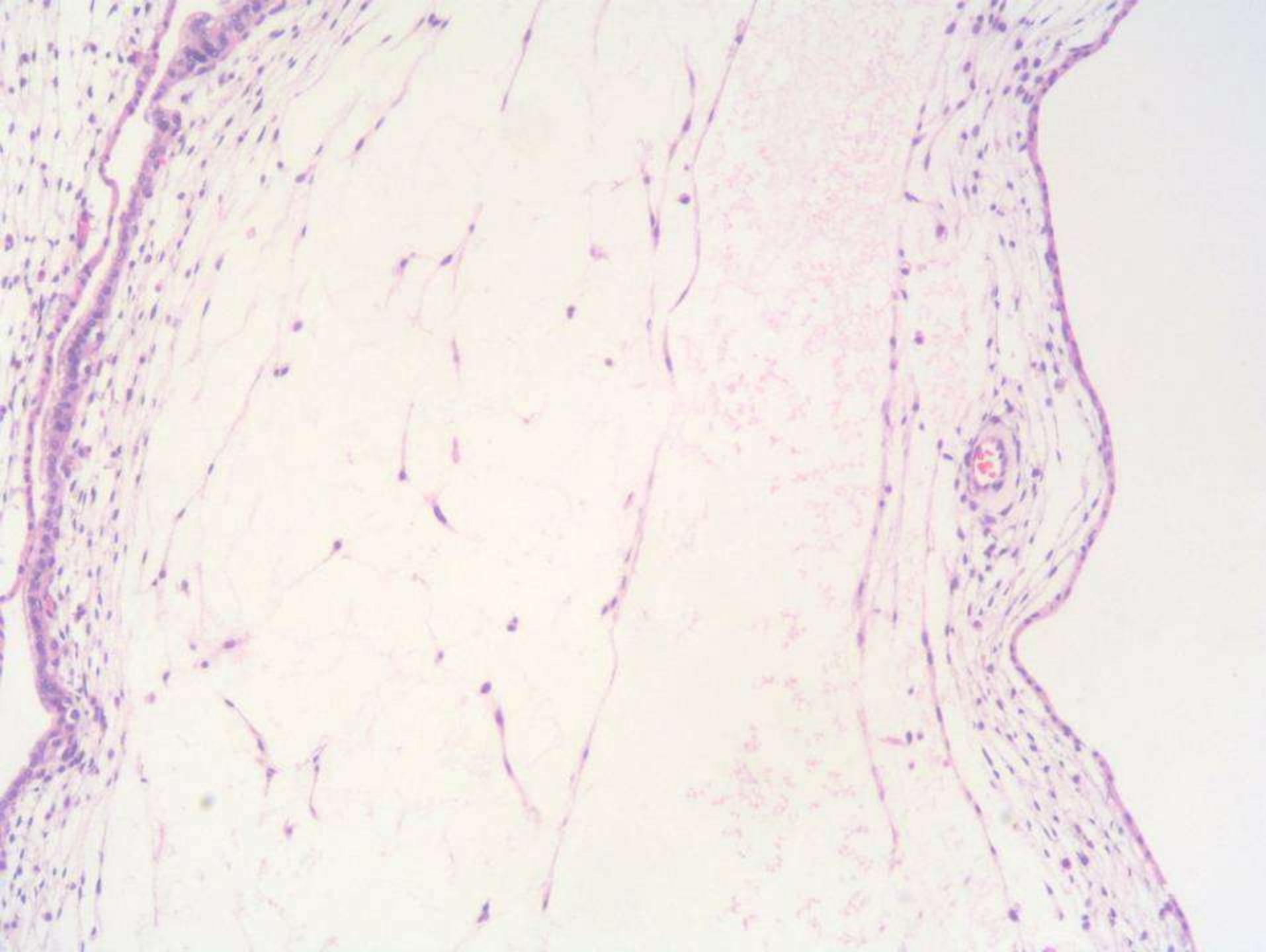
# DMP en el 1er trimestre

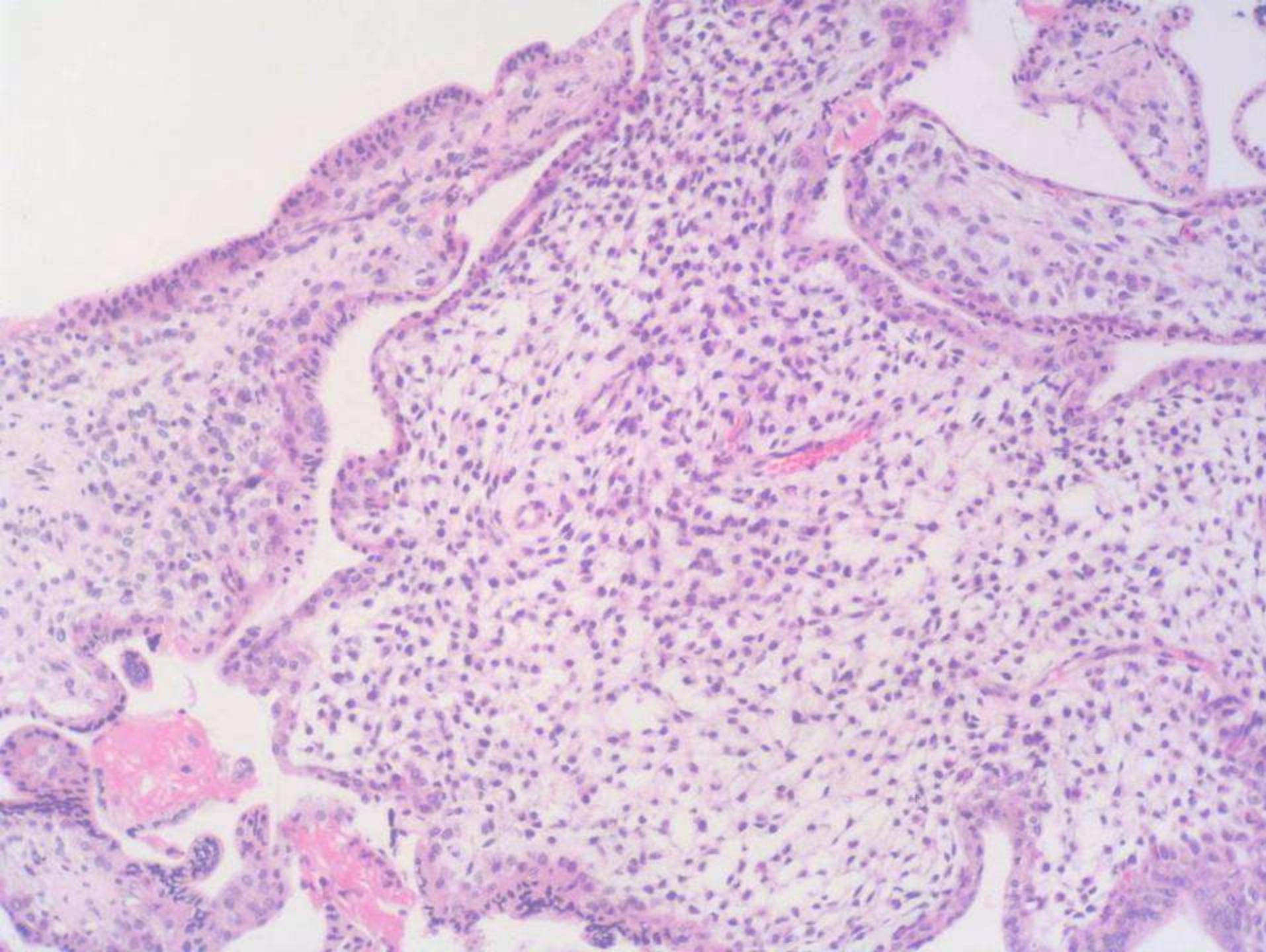
DD con MHP

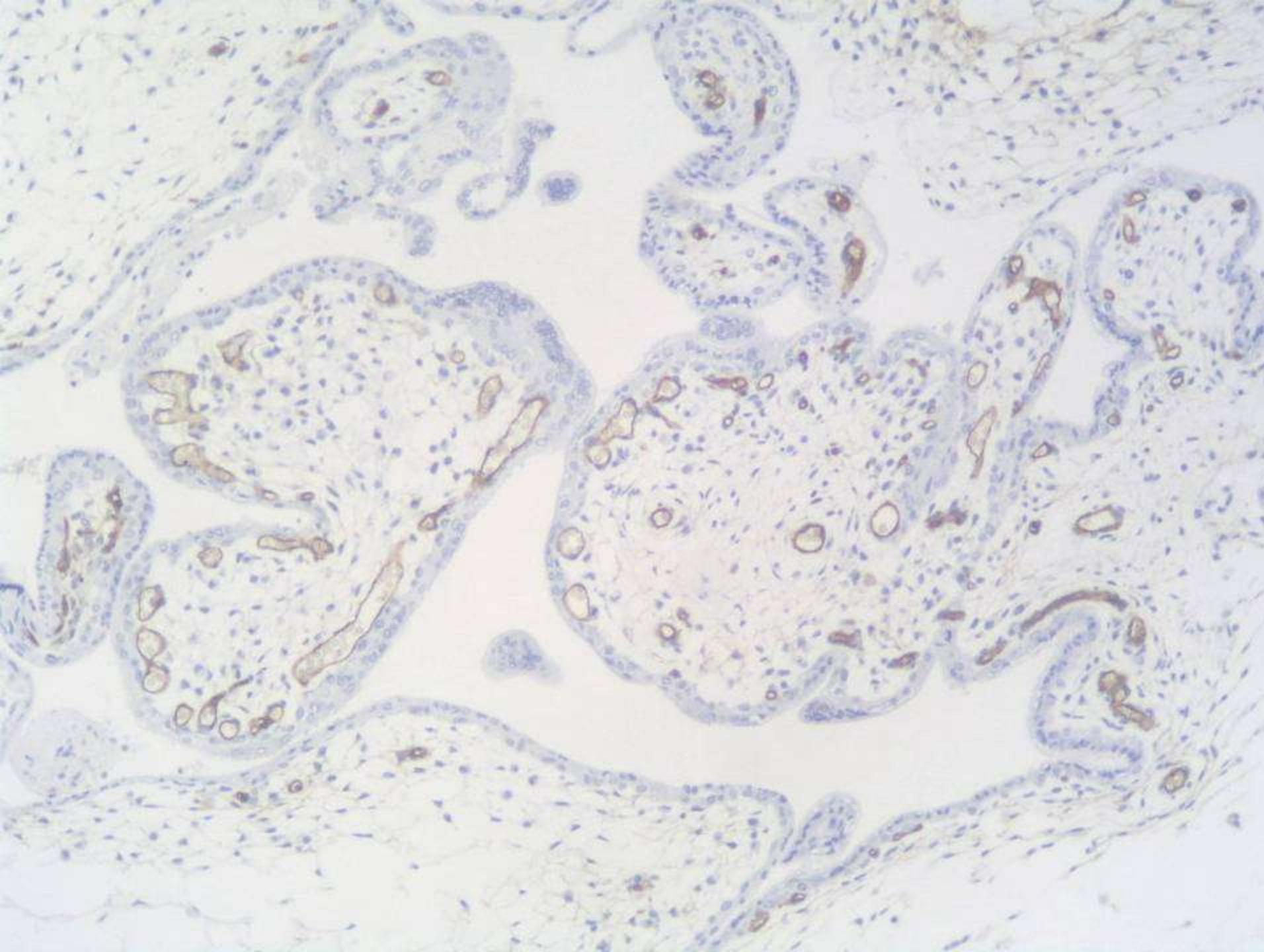
Evolución:  
Dilatación quística  
Vesículas  
Proliferación  
fibroblástica estromal  
Anomalías en cualquier  
tipo de vaso







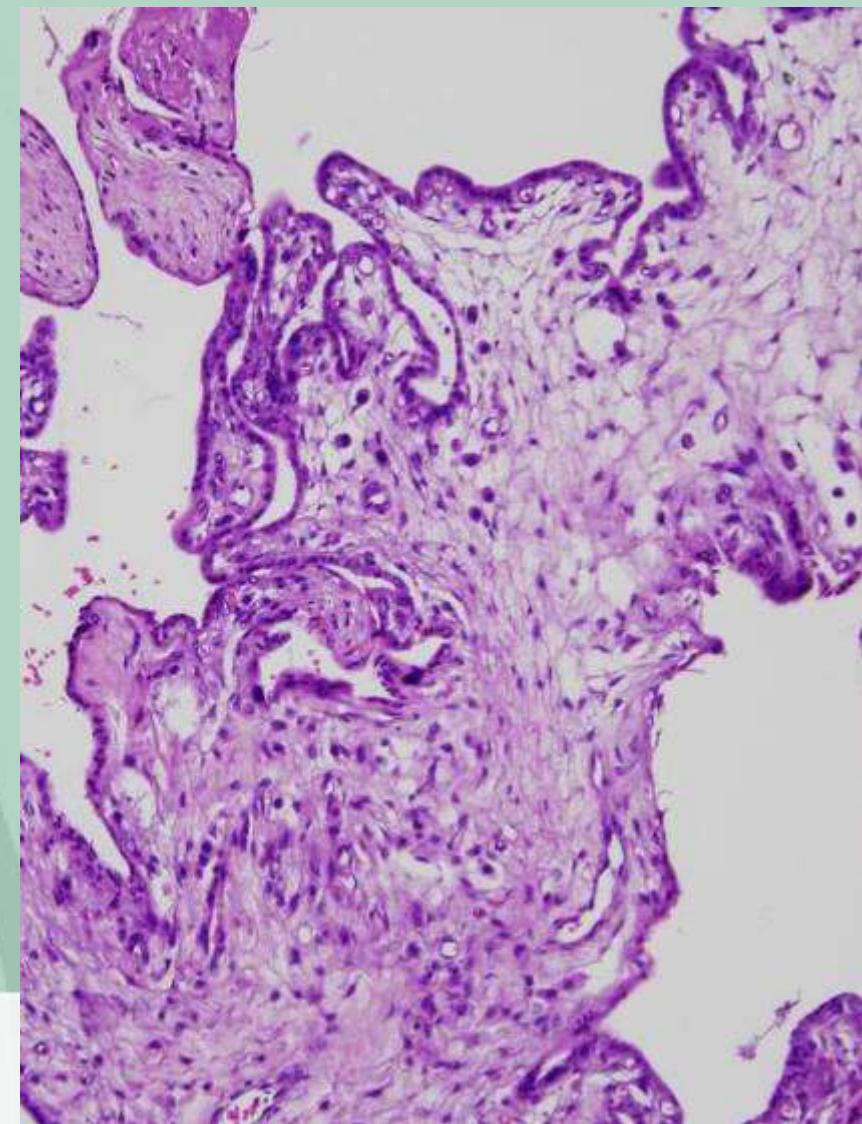
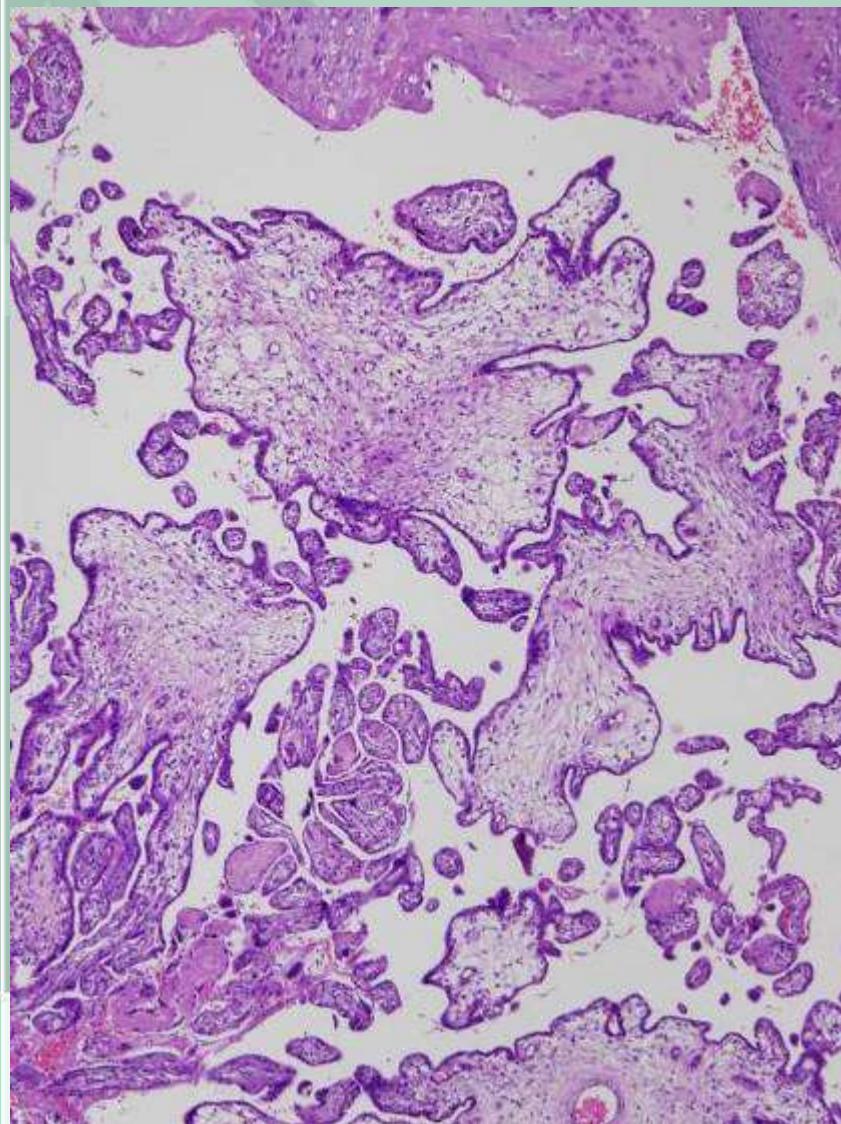




# Cromosomopatías numéricas

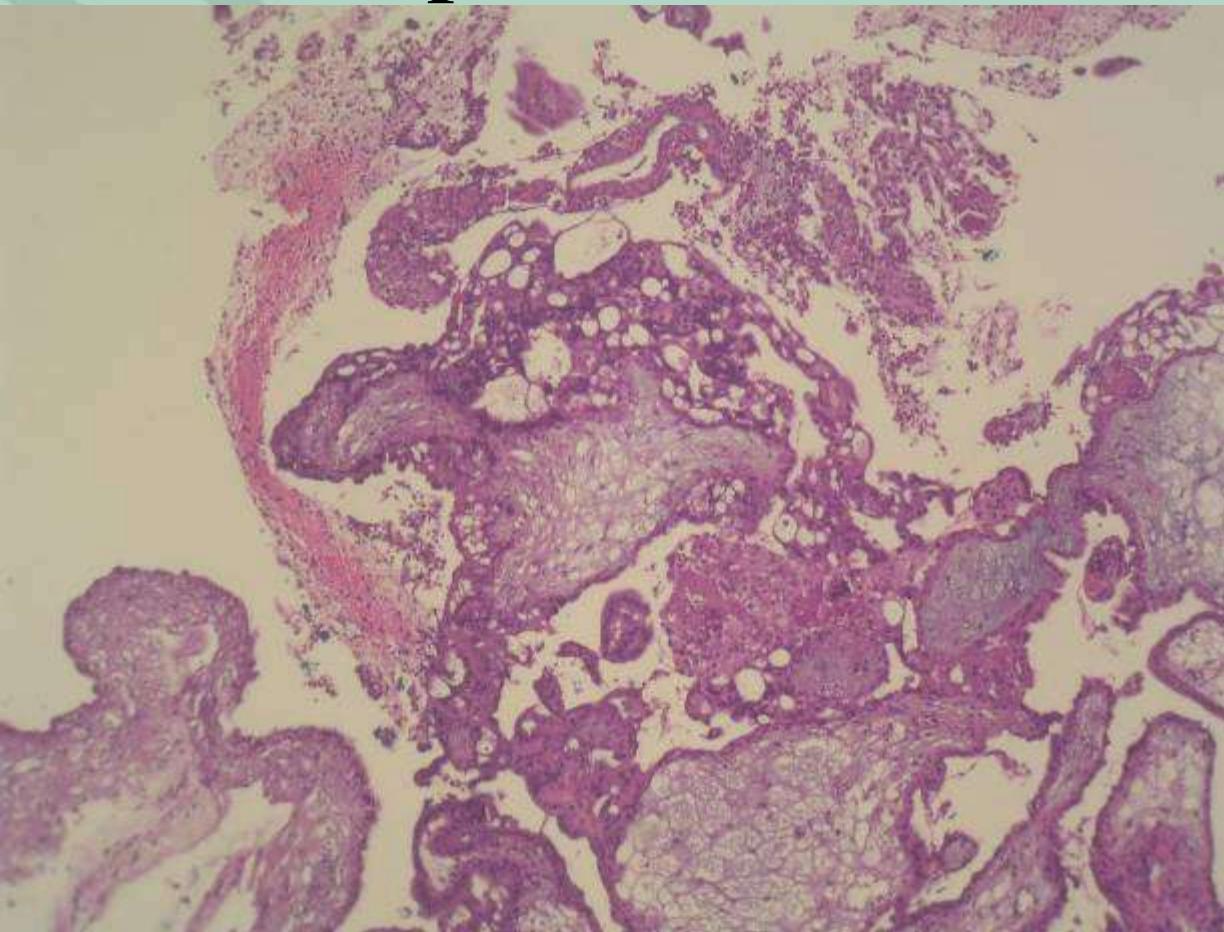
- Trisomias y monosomias no son viables
  - Excepto: 47, +21; 47, +18; 47, +13 y 45, X0
- 1<sup>a</sup> causa de pérdida gestacional temprana
- 47, +16; 47, +22; 47, +7; etc
- Arteria umbilical única
- Retraso del crecimiento
- Alteraciones morfológicas vellositarias

# Vellosidades dismórficas





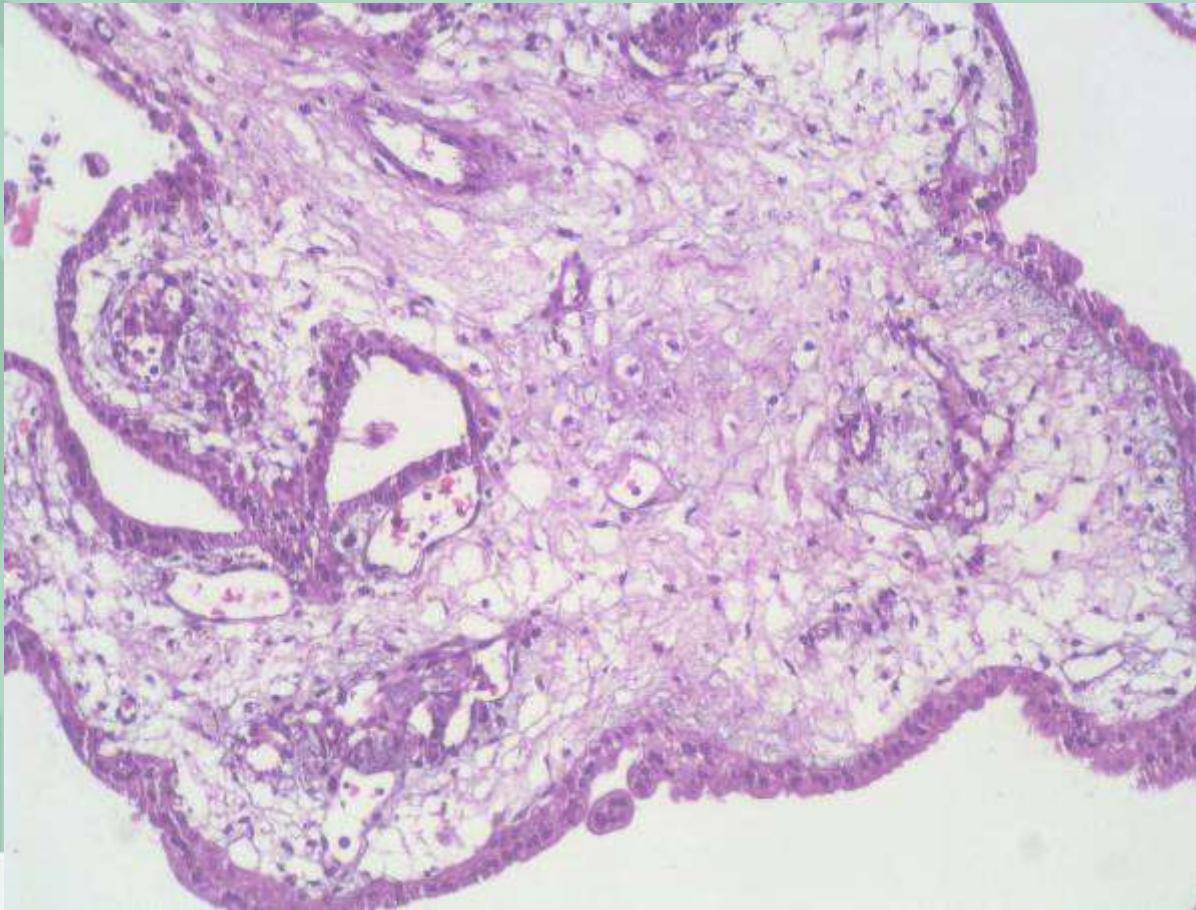
# Hiperplasia trofoblástica no específica



47, +7  
47, +15  
47, +21  
47, +22



# Vellosidades dismórficas proliferación capilar





Degeneración hidrópica

Mola parcial

Edema vellositario

Dilatación cisternal

Doble población vellositaria

Vellosidades avasculares

Proliferación capilar

Afectación de vasos de diferentes calibres

Hiperplasia trofoblástica

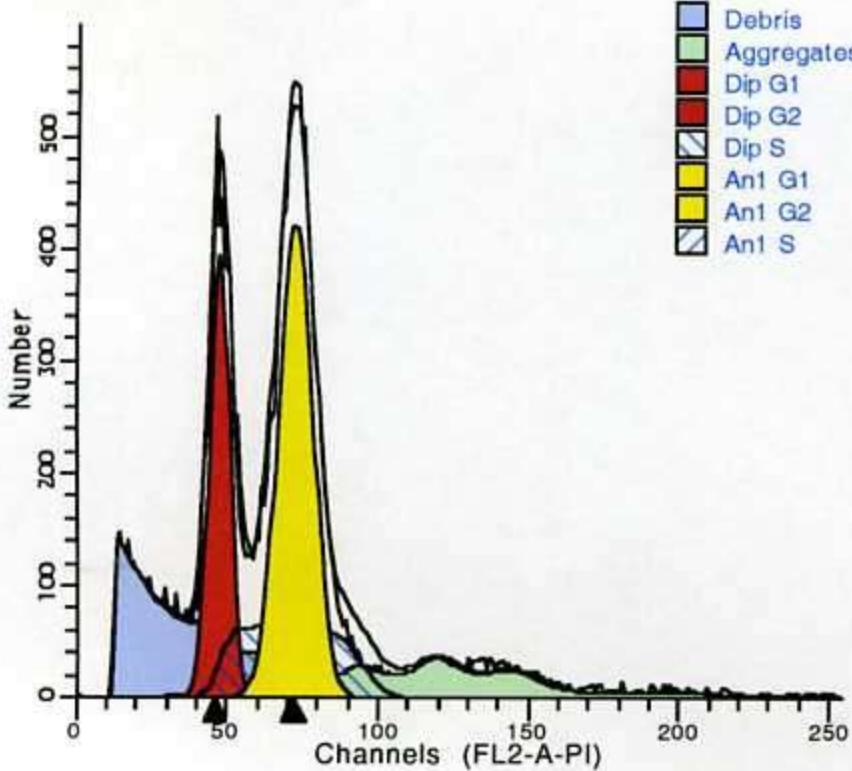
Displasia mesenquimática

Vellosidades dismórficas



# Técnicas auxiliares

- Cariotipo
- Requiere células vivas
  - Triploide: MP
  - Trisómico: VelloSIDADES dismórficas
- PCR cuantitativa fluorescente (QF-PCR)
  - Depende de los “primers”
- Citometria
- IHQ ( $p57^{kip2}$ )



Debris  
Aggregates  
Dip G1  
Dip G2  
Dip S  
An1 G1  
An1 G2  
An1 S

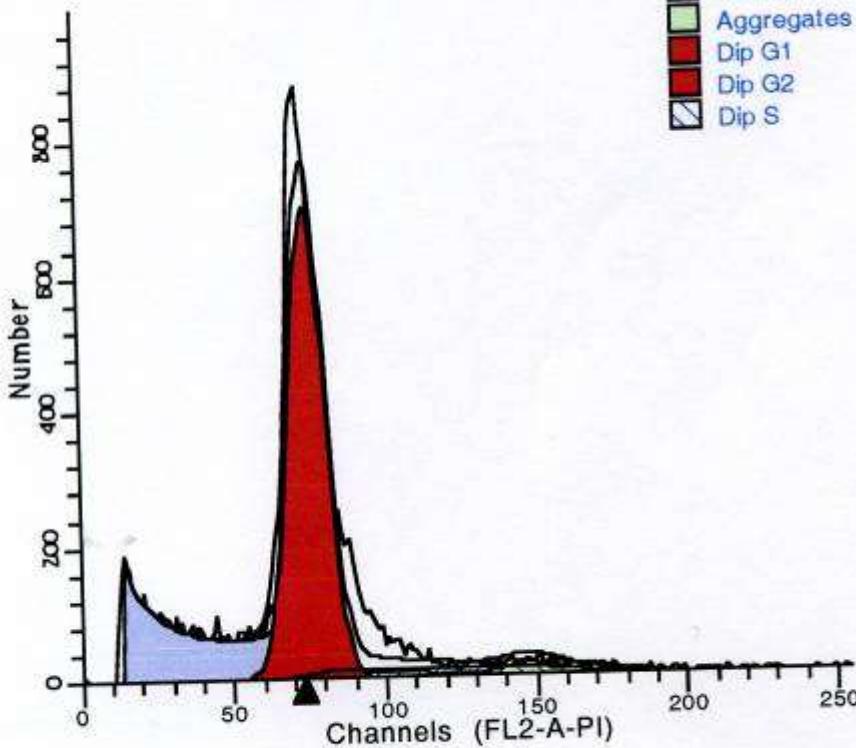
File analyzed: B09\_06105-3.034  
Date analyzed: 3-Apr-2009  
Model: 2DA0n\_DSD ASF  
Analysis type: Manual analysis

Diploid: 50.74 %  
Dip G1: 52.35 % at 47.17  
Dip G2: 0.00 % at 94.35  
Dip S: 47.65 % G2/G1: 2.00  
%CV: 7.03

Aneuploid 1: 49.26 %  
An1 G1: 99.90 % at 72.41  
An1 G2: 0.00 % at 96.69  
An1 S: 0.10 % G2/G1: 1.34  
%CV: 7.97 DI: 1.53

Total Aneuploid S-Phase: 0.10 %  
Total S-Phase: 24.22 %  
Total B.A.D.: 16.39 %

Debris: 23.75 %  
Aggregates: 12.96 %  
Modeled events: 19591  
All cycle events: 12398  
Cycle events per channel: 245  
RCS: 3.268



Debris  
Aggregates  
Dip G1  
Dip G2  
Dip S

File analyzed: B08/15902-1.036  
Date analyzed: 12-Jan-2009  
Model: 1DA0n\_DSF  
Analysis type: Manual analysis

Diploid: 100.00 %  
Dip G1: 89.74 % at 75.04  
Dip G2: 0.00 % at 163.73  
Dip S: 10.26 % G2/G1: 2.18  
%CV: 7.85

Total S-Phase: 10.26 %  
Total B.A.D.: 18.09 %

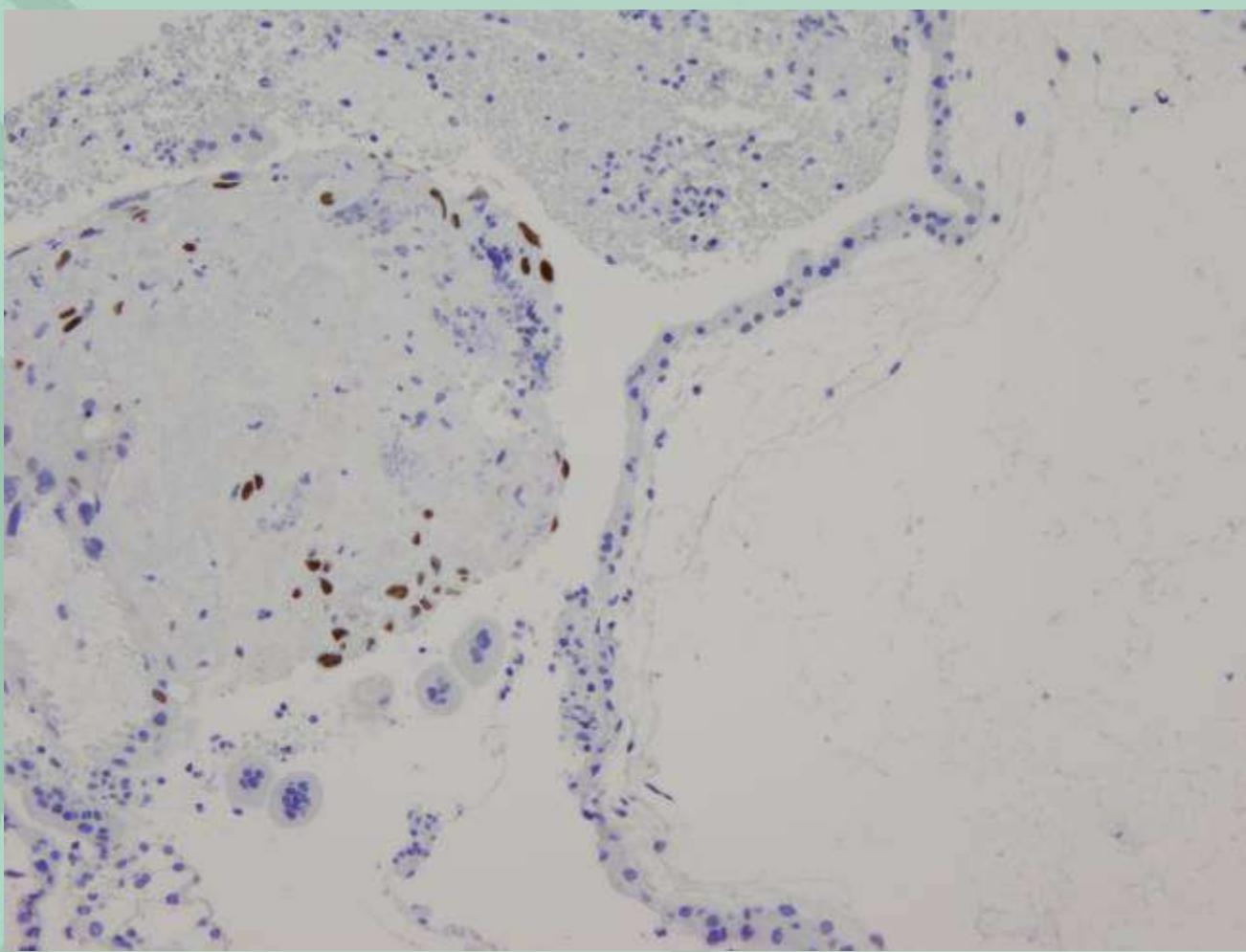
Debris: 31.52 %  
Aggregates: 5.67 %  
Modeled events: 18410  
All cycle events: 11564  
Cycle events per channel: 129  
RCS: 5.417

# IHQ p57<sup>kip2</sup>

	Citotrofoblasto	Mesenquima vellositario	Trofoblasto extravellositario	Decidua
Normal	+	+	+	+
MHP	+	+	+	+
MHC	-	-	+	+

+: >10%

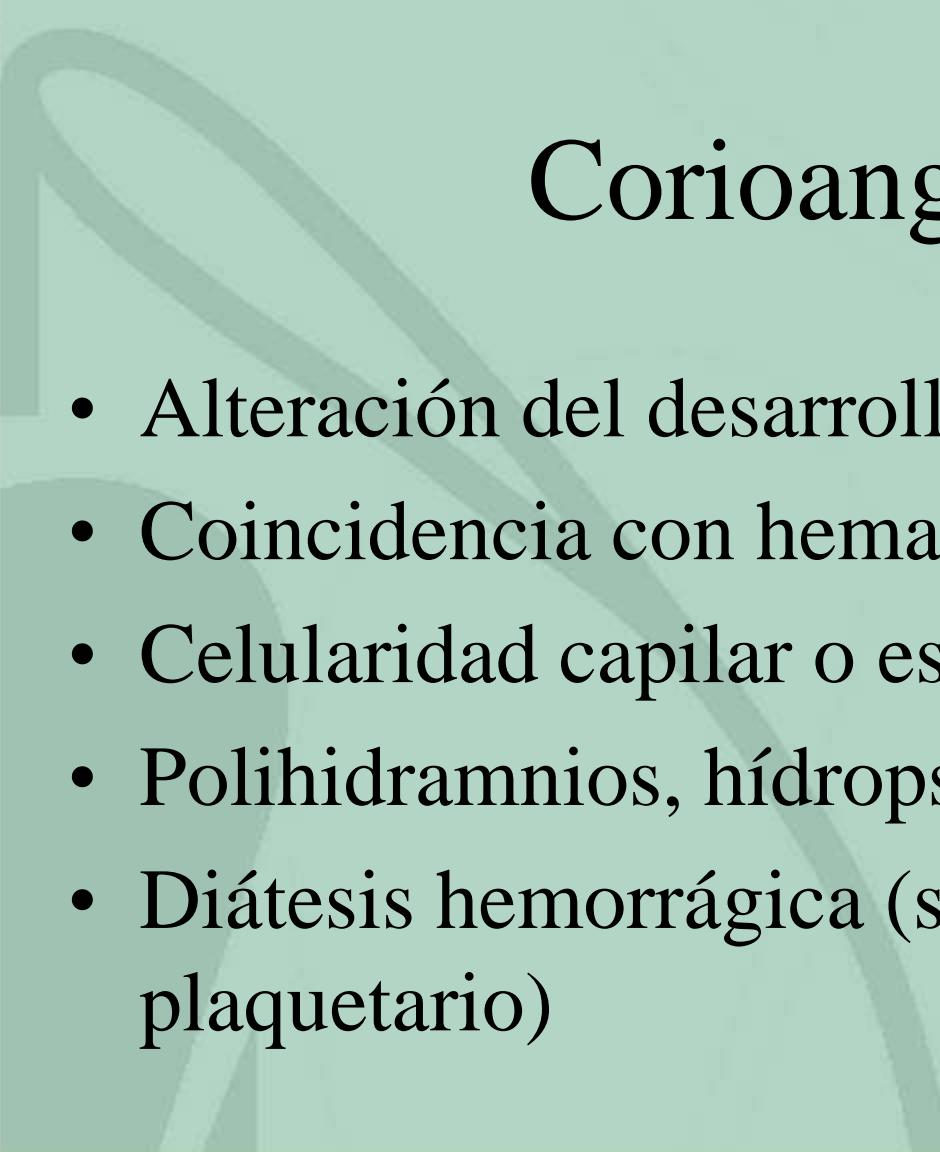
Fisher et al Human Mol Genet 2002, 11: 3267





# Tumores placentarios

- Primarios
  - Corioangioma
- Secundarios
  - Maternos
    - Melanoma, leucemia/linfoma, mama, pulmón
    - Enfermedad avanzada
    - Espacio intervellositario
  - Fetales
    - Neuroblastoma, leucemia
    - Intravellositarios



# Corioangioma

- Alteración del desarrollo
- Coincidencia con hemangiomas fetales
- Celularidad capilar o estromal
- Polihidramnios, hídrops y/o muerte fetal
- Diátesis hemorrágica (secuestro plaquetario)

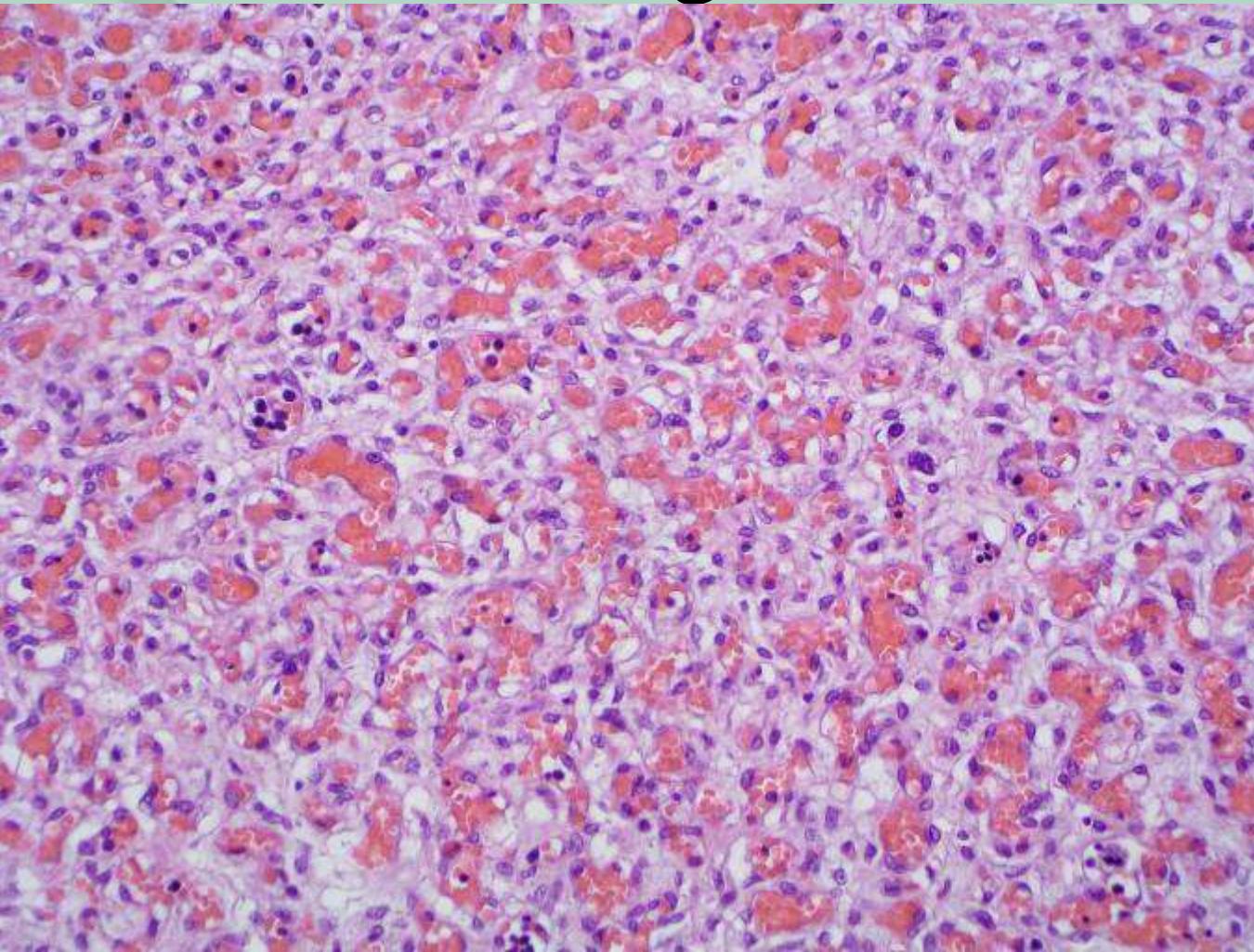
# Corioangioma



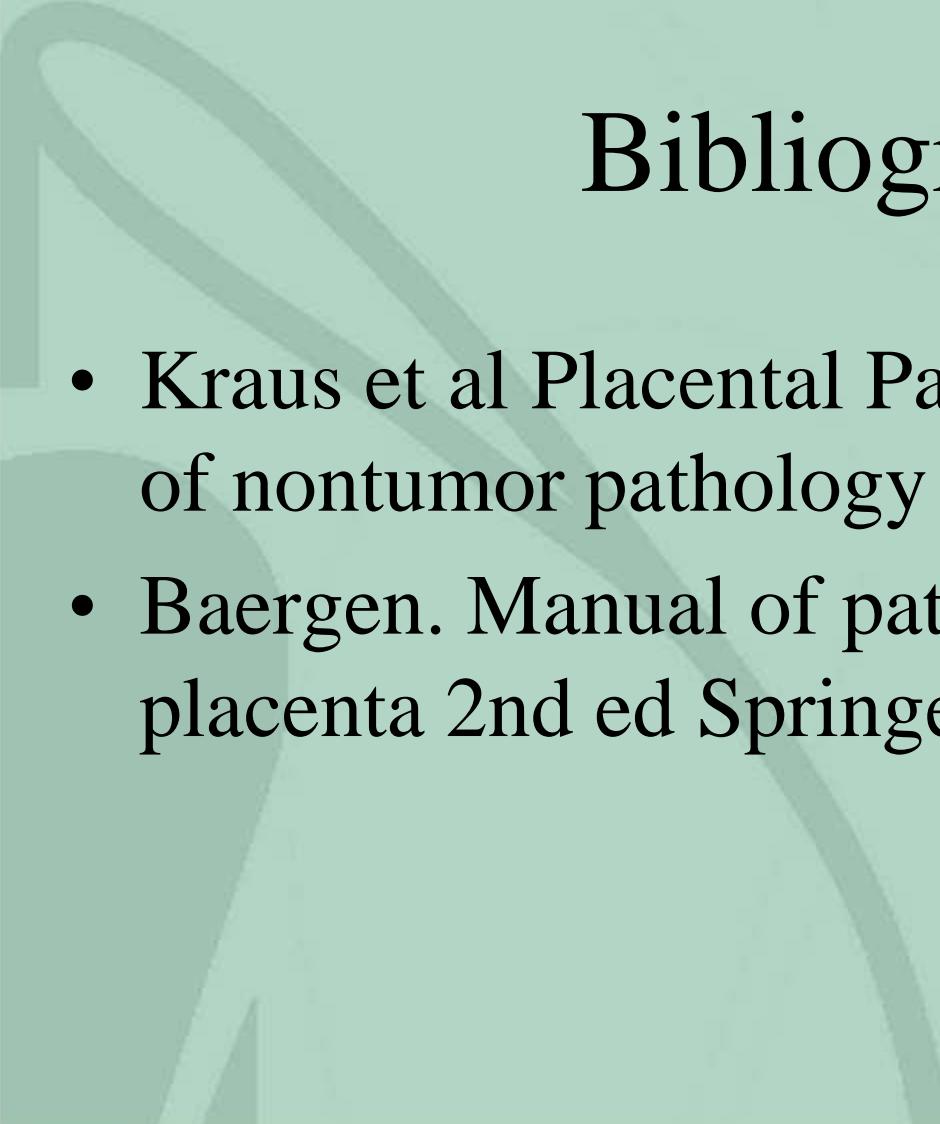
# Corioangioma



# Corioangioma

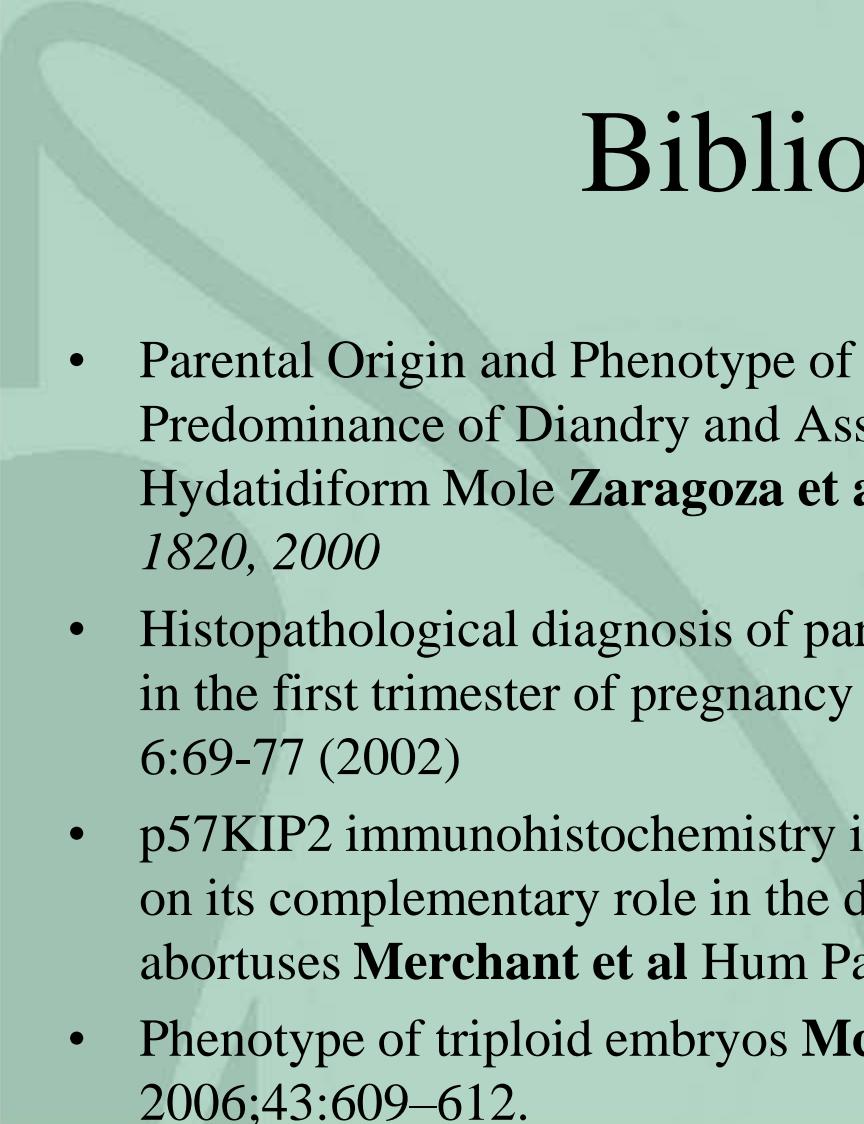






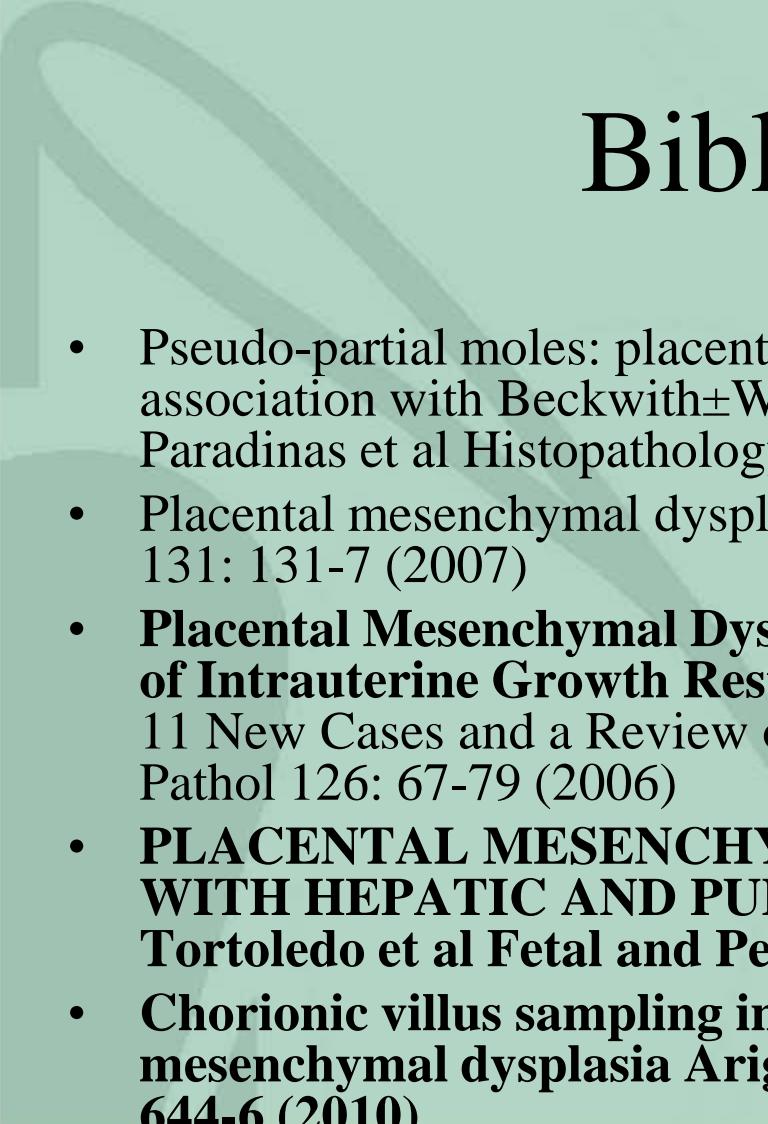
# Bibliografia

- Kraus et al Placental Pathology AFIP Atlas of nontumor pathology series (nº 3) 2004
- Baergen. Manual of pathology of the human placenta 2nd ed Springer 2011



# Bibliografia

- Parental Origin and Phenotype of Triploidy in Spontaneous Abortions: Predominance of Diandry and Association with the Partial Hydatidiform Mole **Zaragoza et al** *Am. J. Hum. Genet.* 66:1807–1820, 2000
- Histopathological diagnosis of partial and complete hydatidiform mole in the first trimester of pregnancy **Sebire et al** *Ped and Dev Pathiol* 6:69-77 (2002)
- p57KIP2 immunohistochemistry in early molar pregnancies: emphasis on its complementary role in the differential diagnosis of hydropic abortuses **Merchant et al** *Hum Pathol* 2005 36: 180-6
- Phenotype of triploid embryos **McFadden et al** *J Med Genet* 2006;43:609–612.



# Bibliografia

- Pseudo-partial moles: placental stem vessel hydrops and the association with Beckwith±Wiedemann syndrome and complete moles Paradinas et al Histopathology 39: 447-54 (2001)
- Placental mesenchymal dysplasia Parveen et al Arch Pathol Lab Med 131: 131-7 (2007)
- **Placental Mesenchymal Dysplasia Is Associated With High Rates of Intrauterine Growth Restriction and Fetal Demise A Report of 11 New Cases and a Review of the Literature** Pham et al Am J Clin Pathol 126: 67-79 (2006)
- **PLACENTAL MESENCHYMAL DYSPLASIA ASSOCIATED WITH HEPATIC AND PULMONARY HAMARTOMA** Tortoledo et al Fetal and Pediatric Pathol 29: 261-71 (2010)
- **Chorionic villus sampling in the prenatal diagnosis of placental mesenchymal dysplasia** Arigita et al Ultrasound obstet Gynecol 36: 644-6 (2010)