

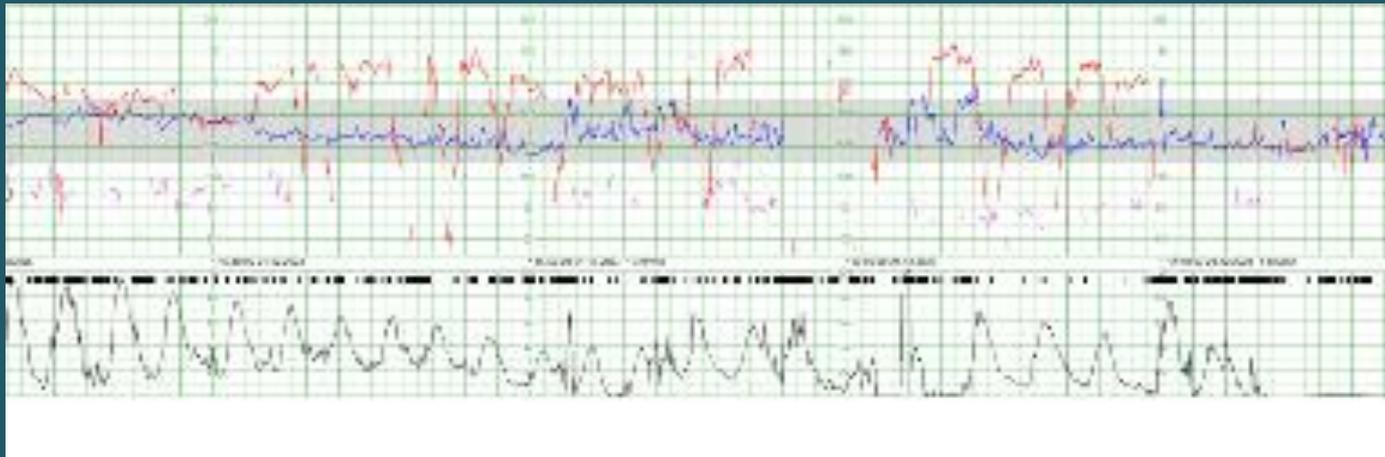
The follow up of multiple pregnancies

What should every neonatologist know?



What is your diagnosis? Case 1

- G1, spontaneous MCDA twins
- Uneventful pregnancy
- US @35w: 2x vertex, normal fluid and dopplers, p22 and p24
- Induction @36w1d
- Vaginal prostaglandins, AROM @3cm

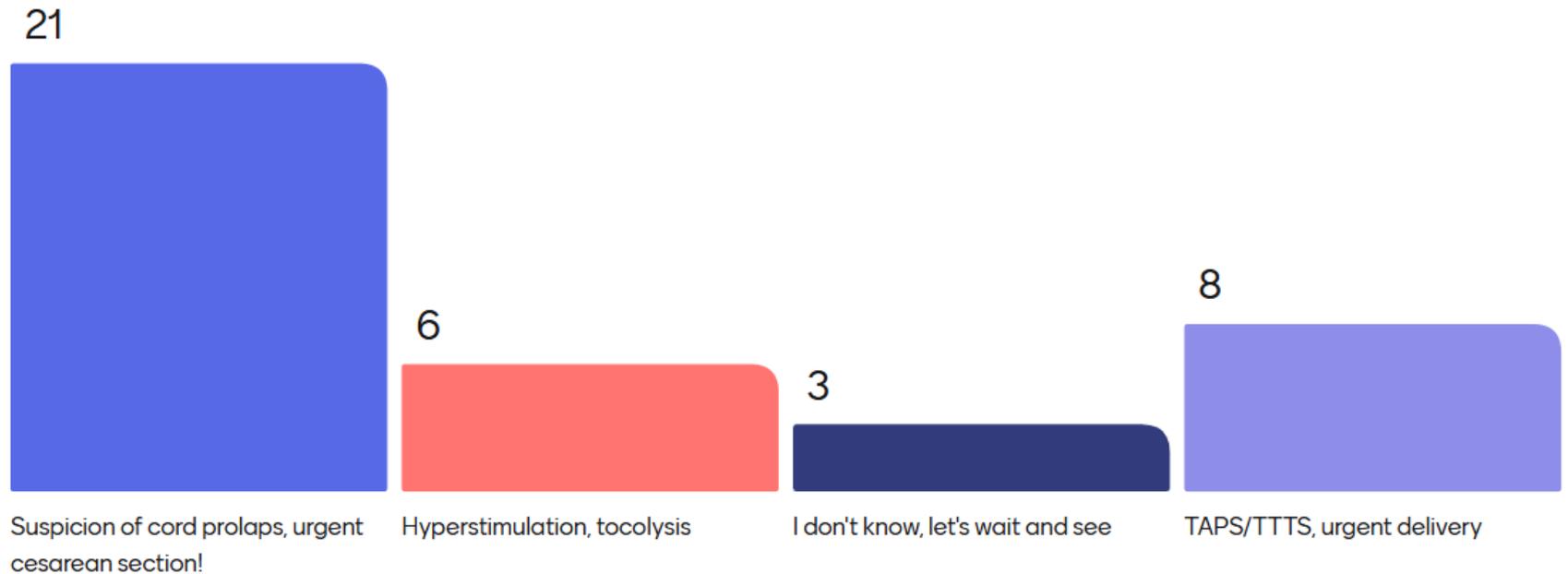


What is your diagnosis? What would you do?



What is your diagnosis? What would you do?

What is happening? What would you do?



What is your diagnosis? Case 1

- Cesarean section



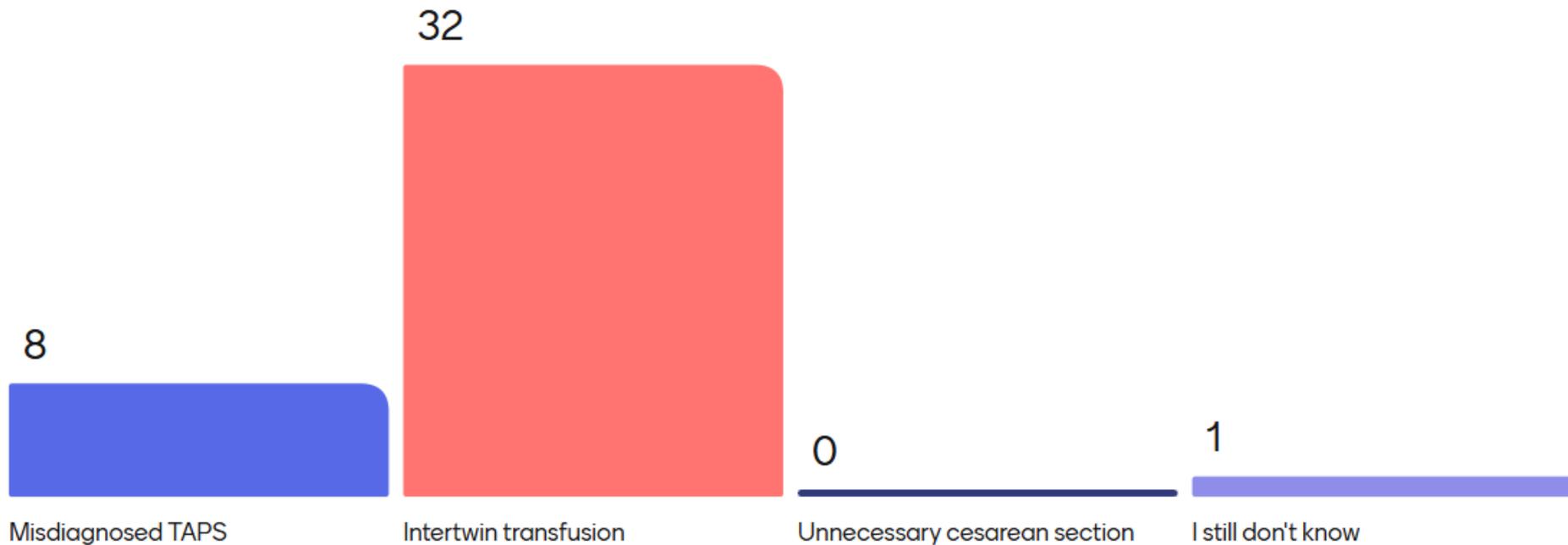
- pH 7.18
- Hb 12 gr/dL
- pH 7.22
- Hb 21 gr/dL

What is your diagnosis now?



What is your diagnosis now?

What is your diagnosis now?



Type & epidemiology of twins

Dichorionic *versus* monochorionic twins

Complications unique to MC twin pregnancies

- TTTS
- TAPS
- Acute intertwin transfusion
- Acardiac twinning

Complications not unique to MC twin pregnancies

- Discordant anomalies
- Discordant growth
- Preterm birth

MOMAs and higher order pregnancies

Type of twins

70% dizygotic



30% monozygotic



dichorionic



30% dichorionic



Poly-ovulation

incidence depends on

Family History

Ethnicity

black > asian

Maternal age

old > young

Parity

multips > primi

Mode of conception

ART >>> spontaneous

69%

monochorionic
diamniotic



1%

monochorionic
monoamniotic



≈ constant: 1 in 250 pregnancies
1 in 50 pregnancies after ART

Epidemiology of twins



Natural incidence of twins

1 in 89 births

Epidemiology of twins

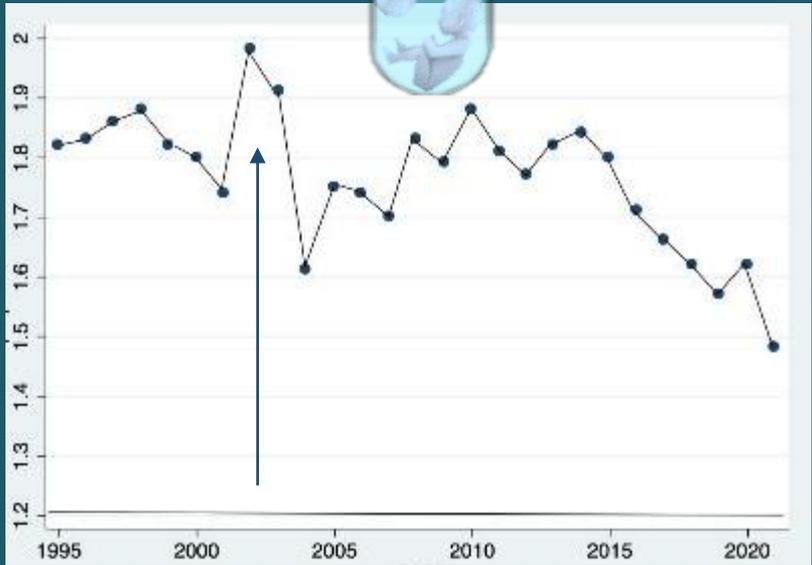


Natural incidence of twins

1 in 89 births

Flanders 2022 1 in 67 births

USA 2021 1 in 32 births



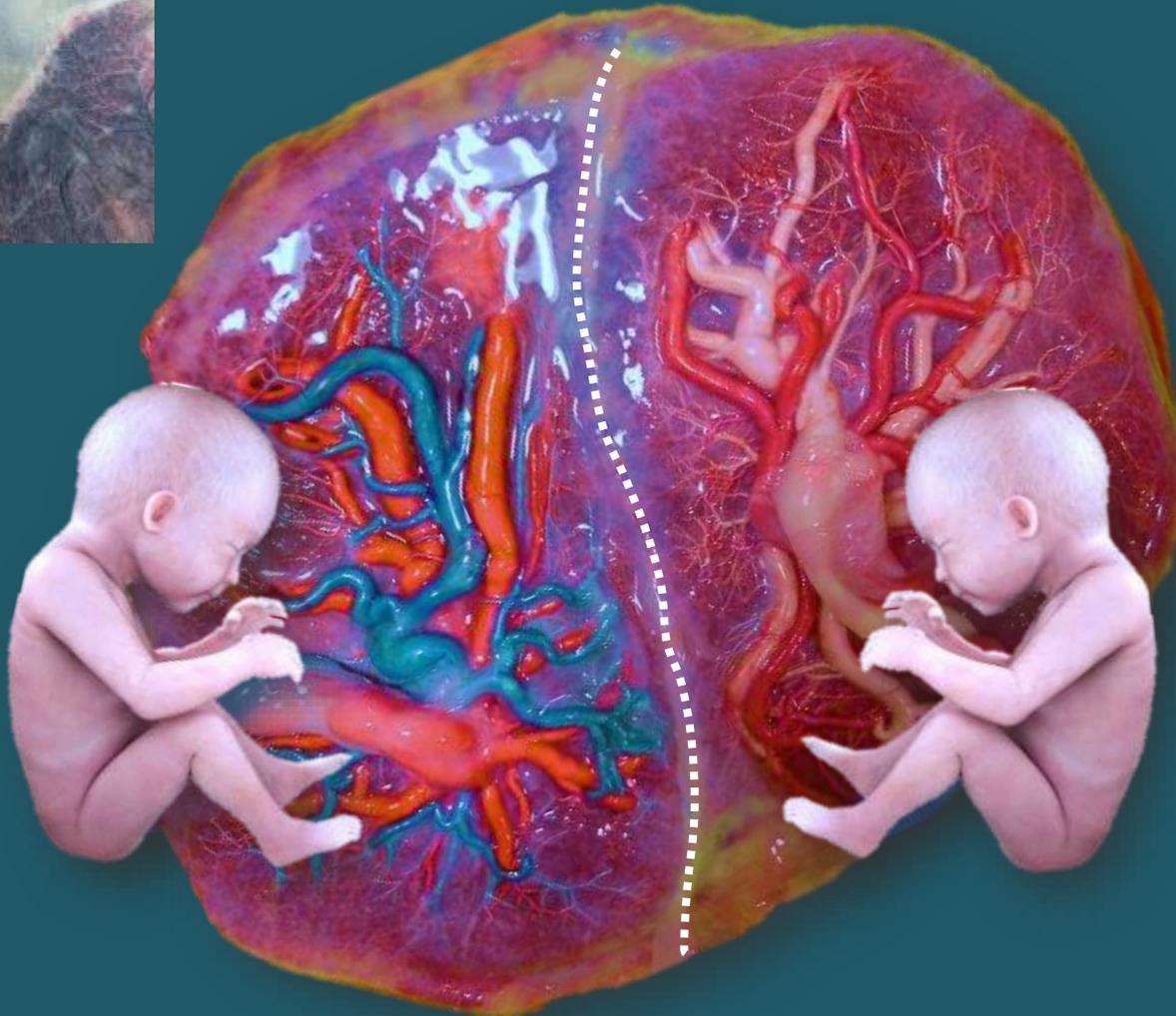
“There is NO diagnosis of twins”

There are only di- or monochorionic twins



“There is NO diagnosis of twins”

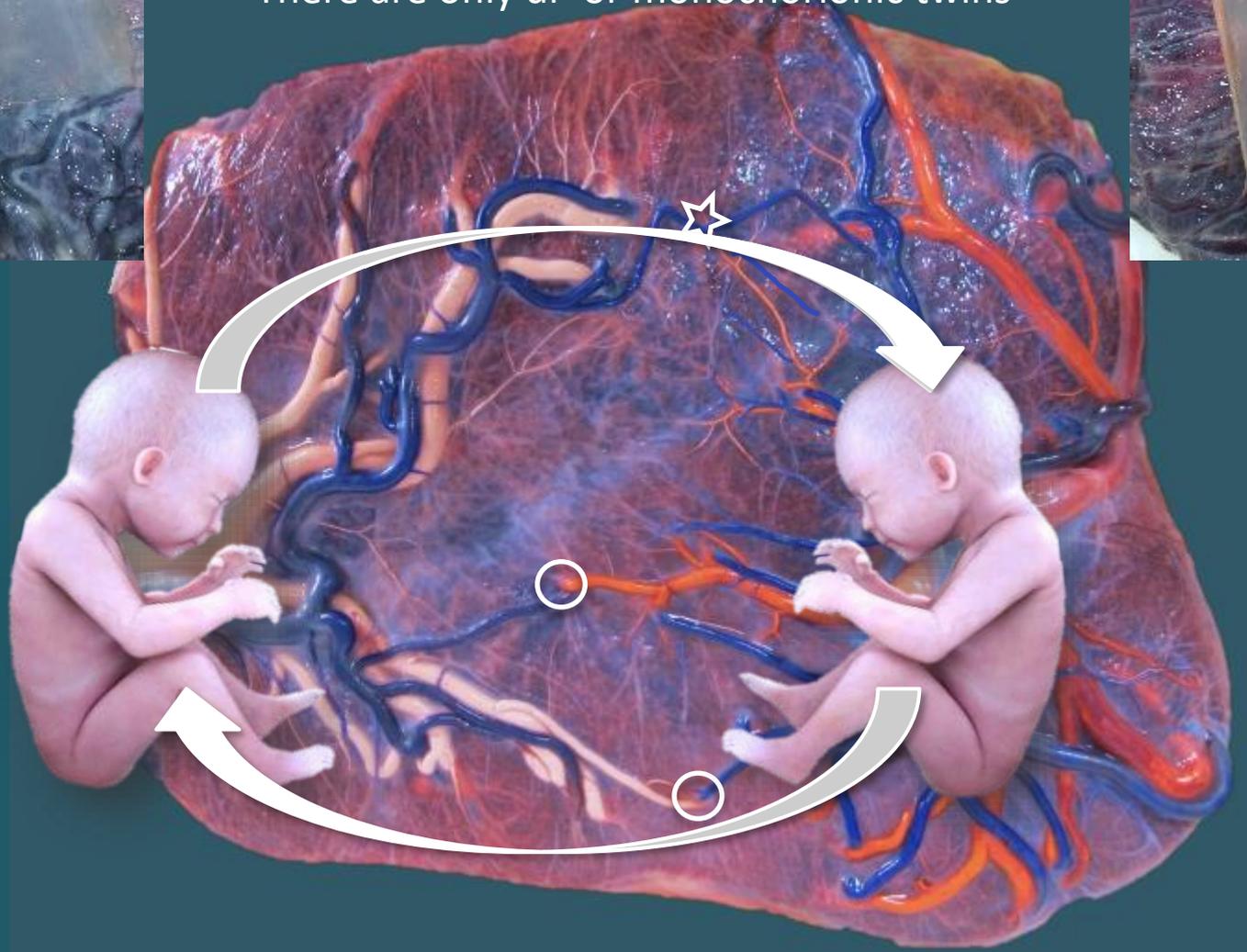
There are only di- or monochorionic twins



Dichorionic: separate circulations

“There is NO diagnosis of twins”

There are only di- or monozygotic twins

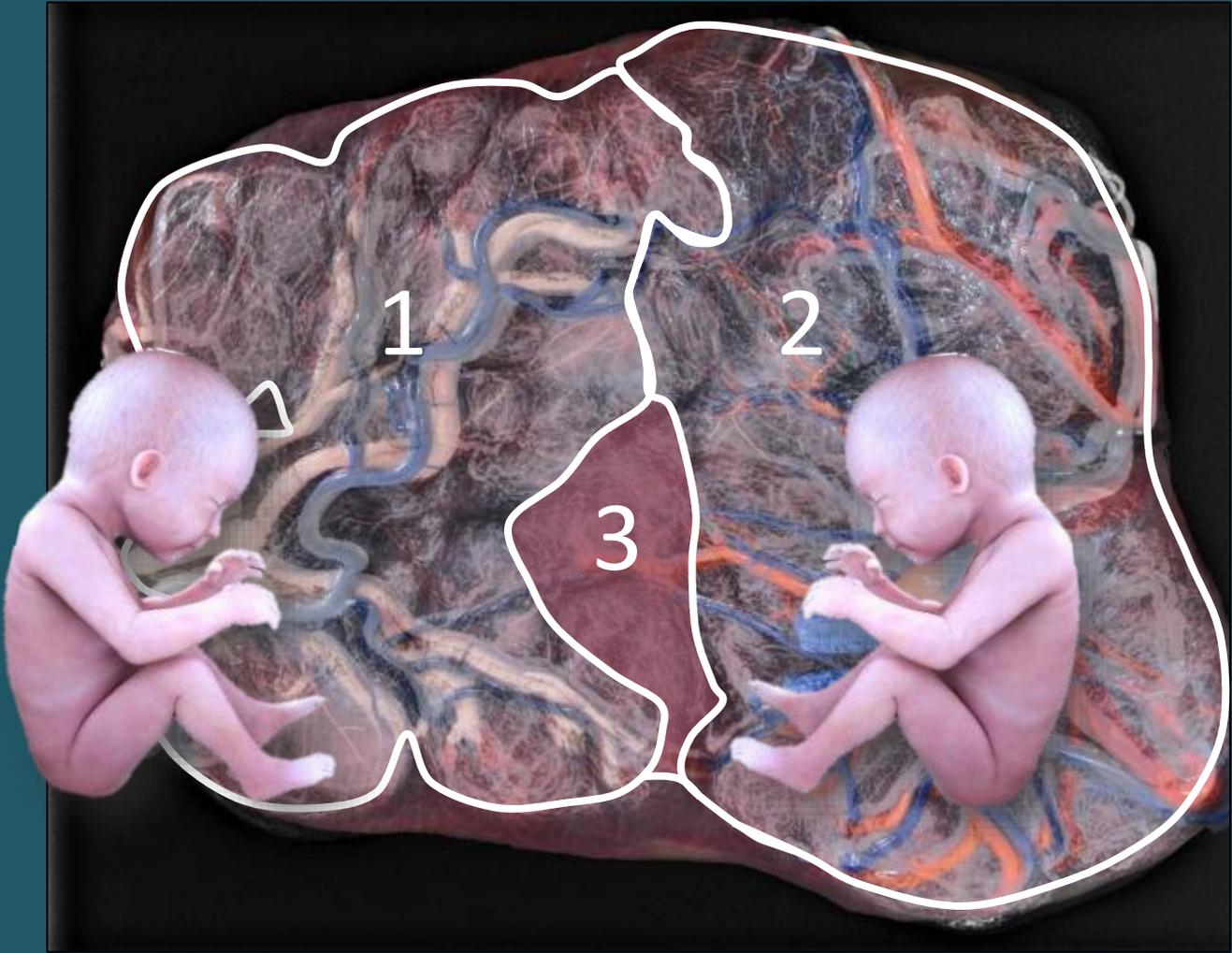


Monozygotic: shared circulations

The monochorionic placenta and its vascular anastomoses

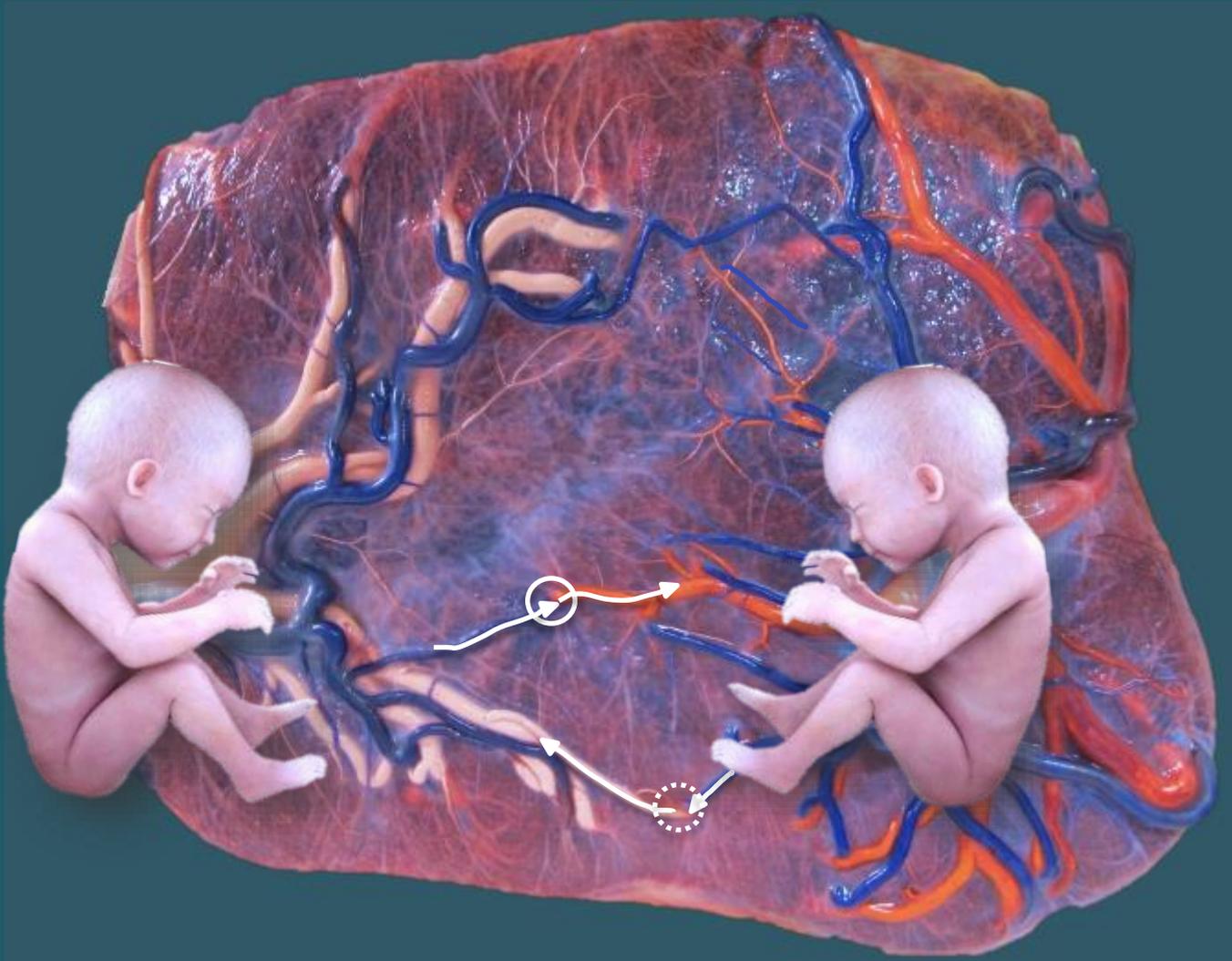


The monochorionic placenta and its vascular anastomoses



The third shared part

The monochorionic placenta and its vascular anastomoses



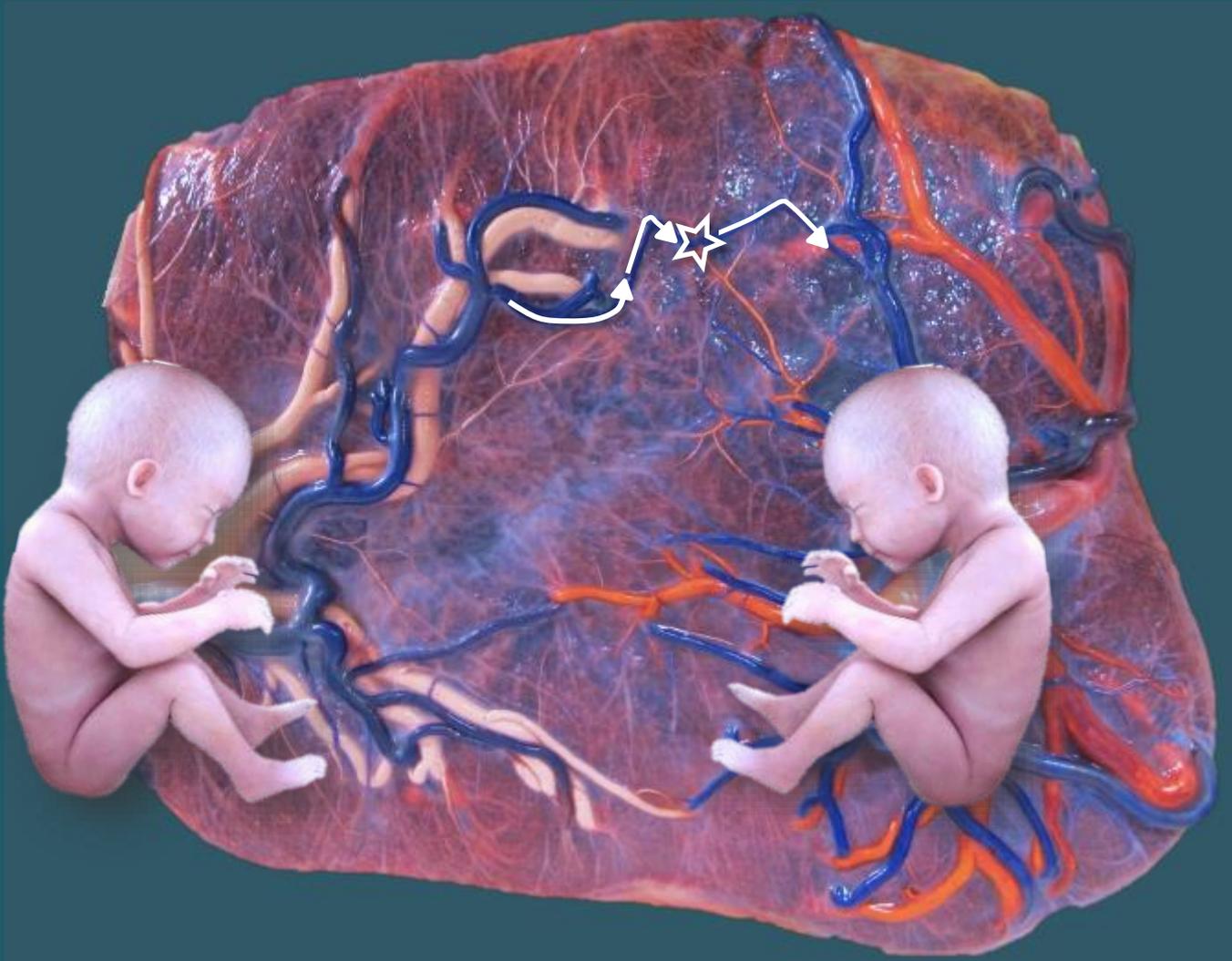
Deep unidirectional artery-to-vein (AV) anastomosis

The monochorionic placenta and its vascular anastomoses



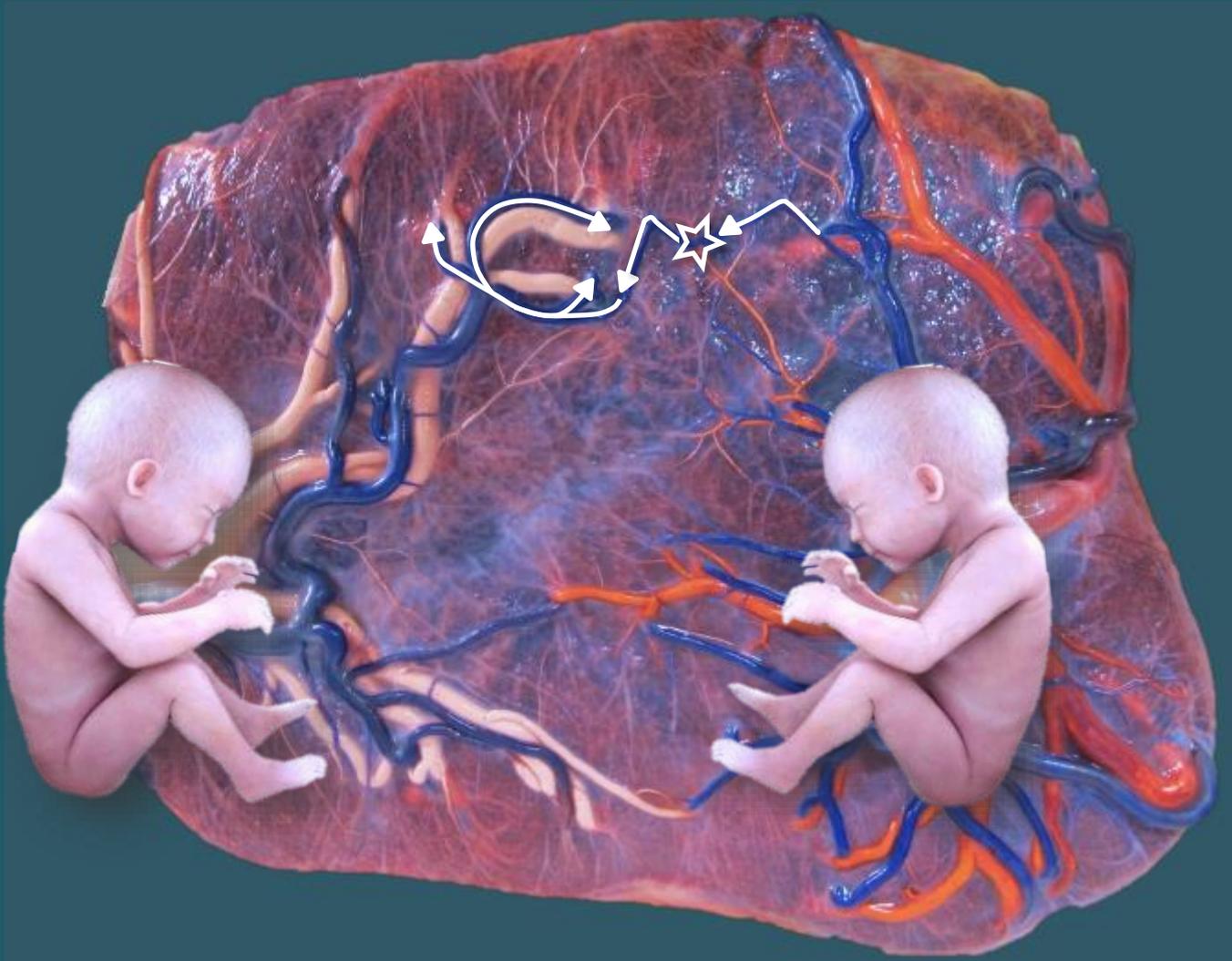
Superficial and bidirectional artery-to-artery (AA) anastomosis

The monochorionic placenta and its vascular anastomoses



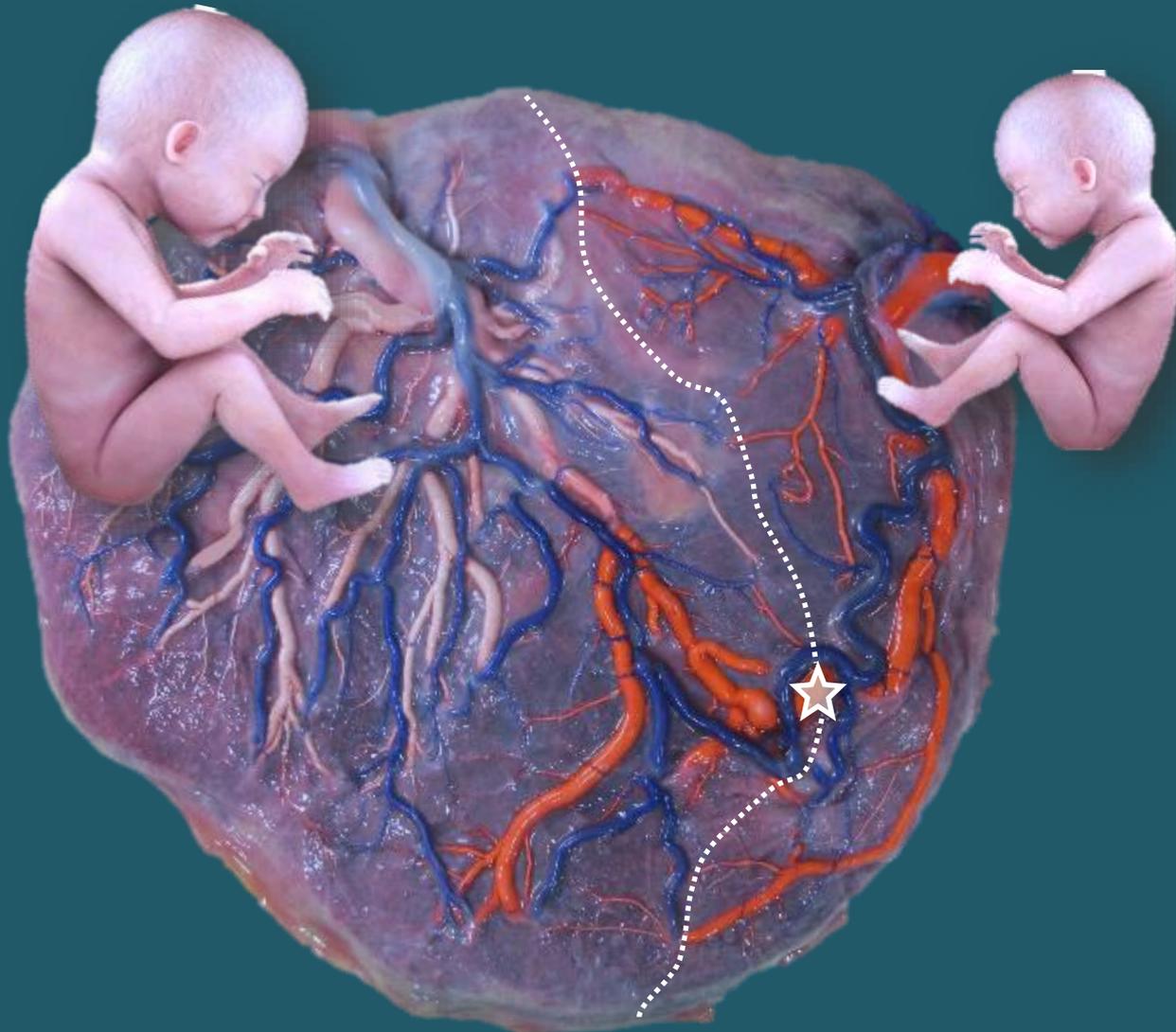
AA is a flexible AV anastomosis

The monochorionic placenta and its vascular anastomoses



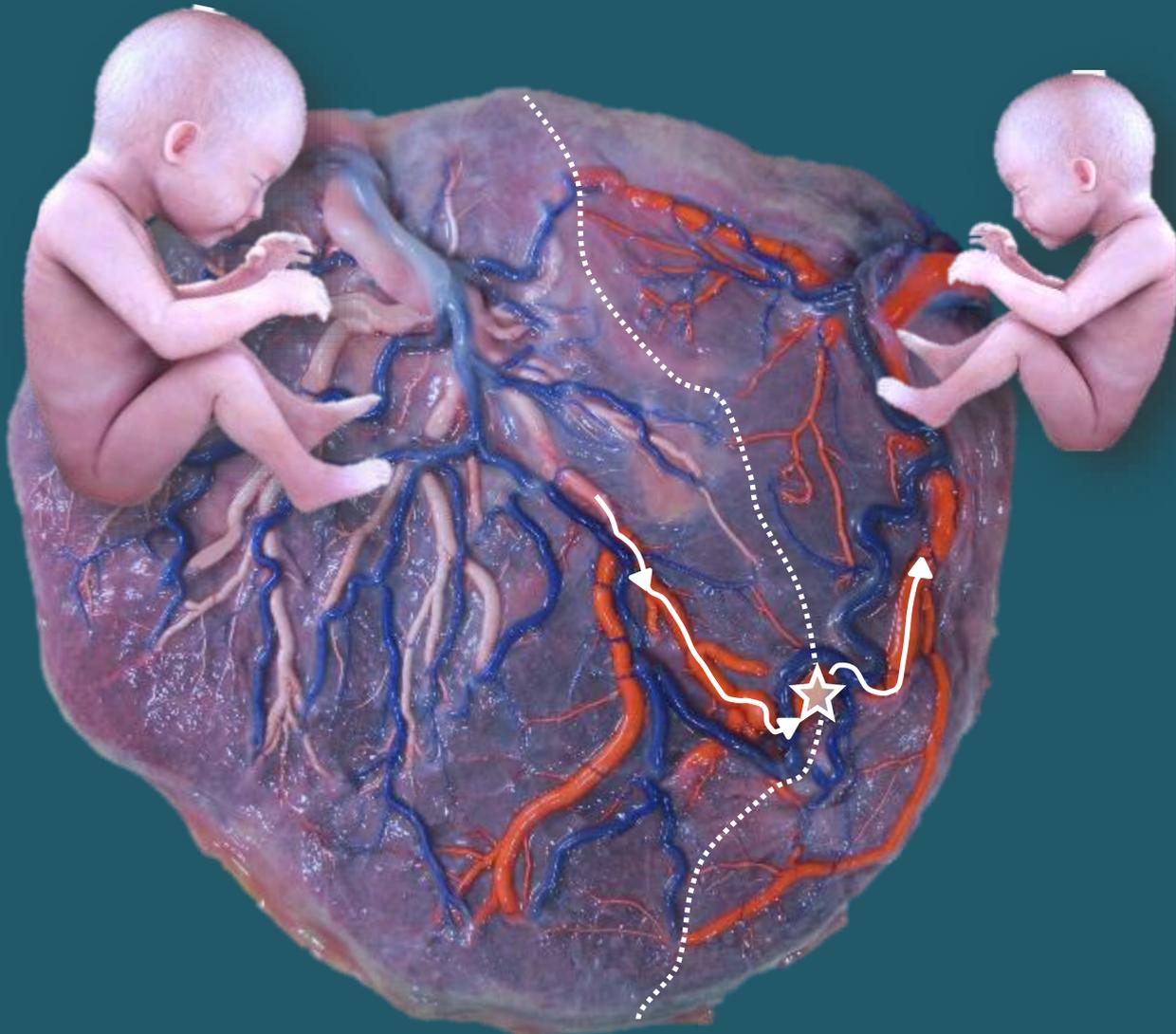
AA is a flexible AV anastomosis

The monochorionic placenta and its vascular anastomoses



Superficial and bidirectional vein-to-vein (VV) anastomosis

The monochorionic placenta and its vascular anastomoses



VV makes the placental sharing flexible

“There is NO diagnosis of twins”

Mono- vs dichorionic twins



Outcome	Monochorionic	Dichorionic
Fetal loss < 24 wks	8%	2%
Perinatal death	2.5%	1%
Birth < 32 wks	14%	7%

Survival	88%	96%
Developmental delay	7-10%	4%
Cerebral palsy	2%	0,5%

“There is NO diagnosis of twins”



Complications unique to MC twins



Complications not unique to MC twins

The follow up of twin pregnancies: how do we do it?



1st trimester



- Chorionicity and amnionicity determination
- Dating
- Labelling
- Detailed anatomy scan
- Screening and diagnosis of chromosomal anomalies

1st trimester: Chorionicity and amnionicity determination

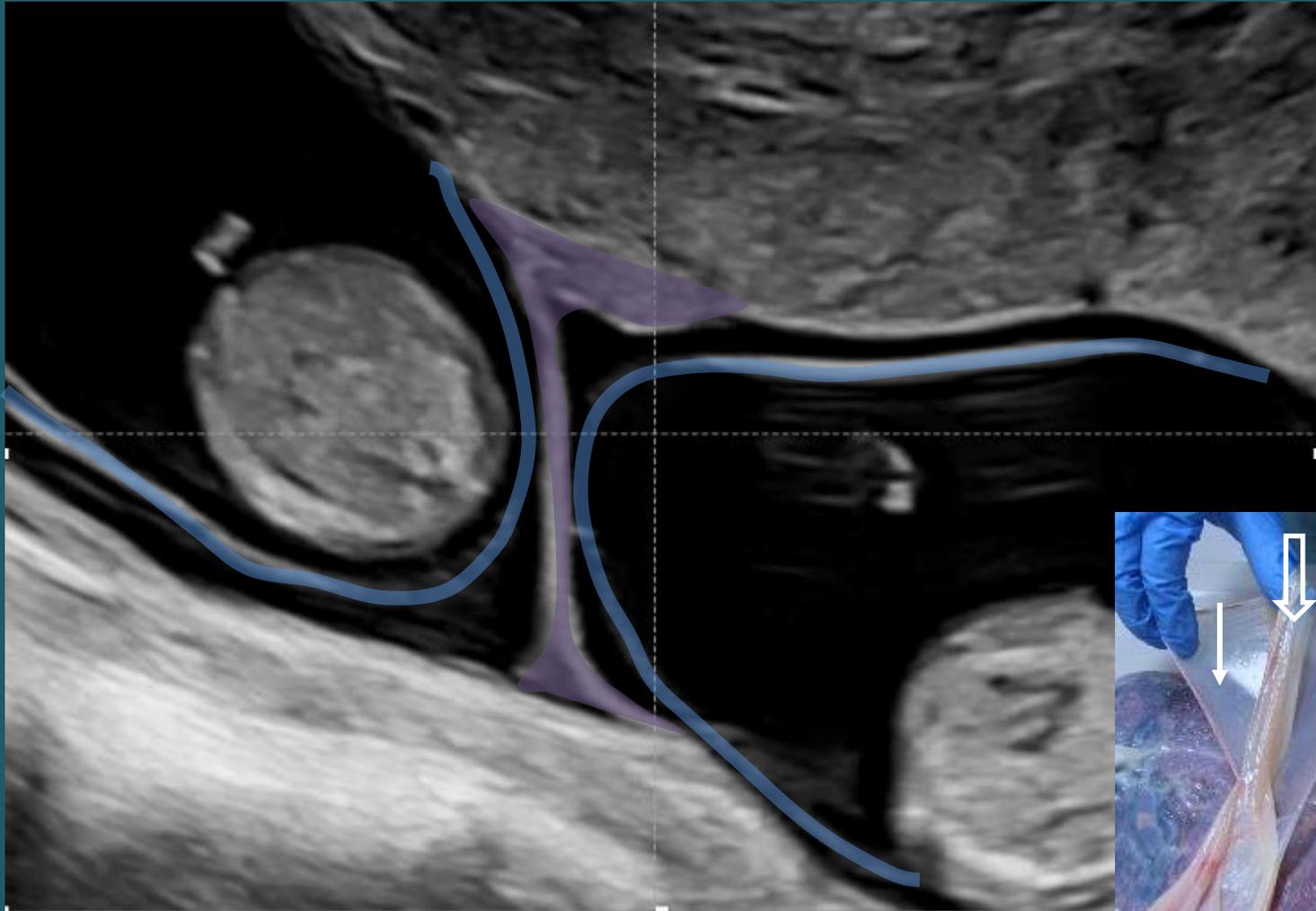
Look at what separates the twins



DC twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

Look at what separates the twins



DC twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

Look at what separates the twins



DC twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

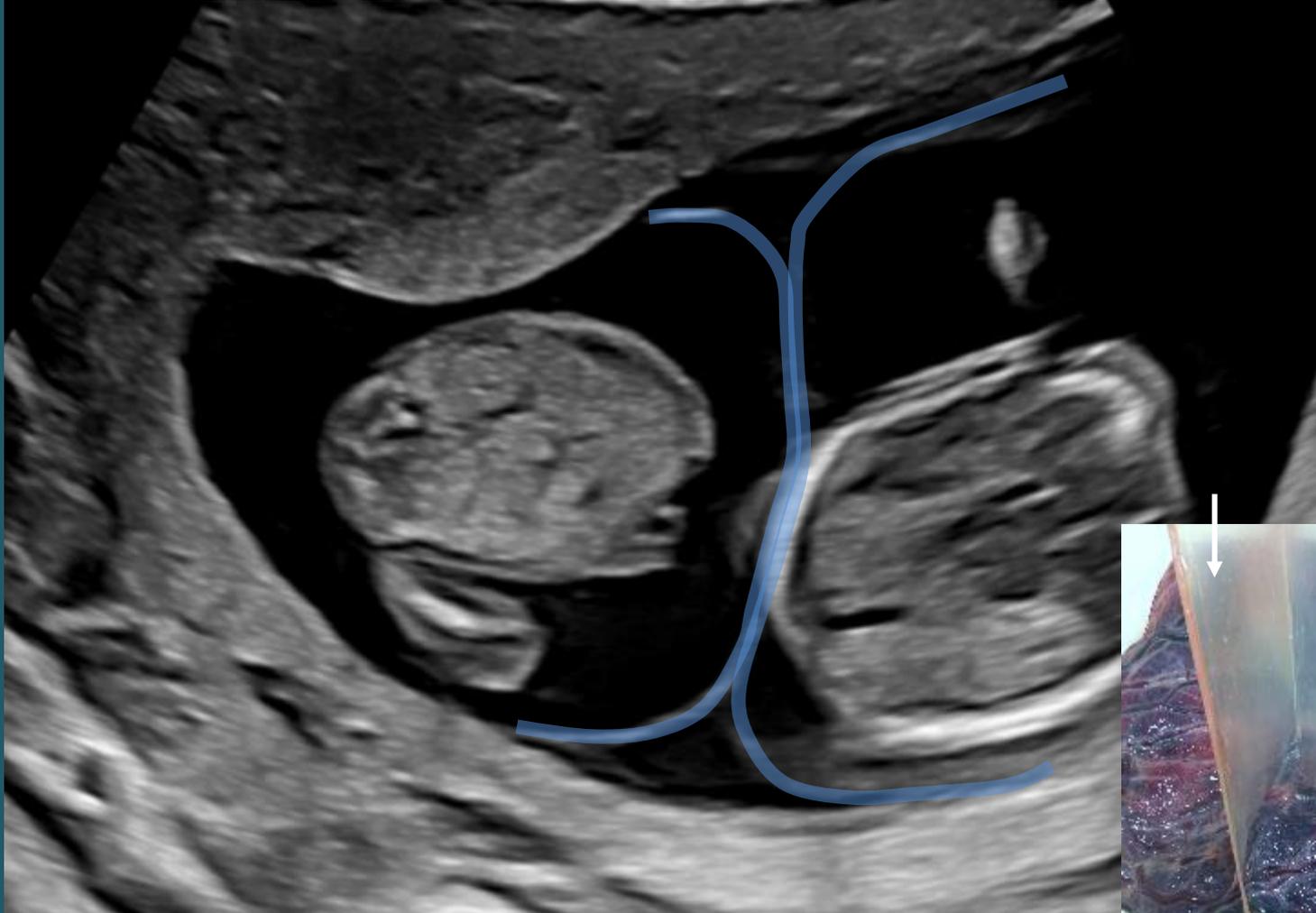
Look at what separates the twins



MCDA twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

Look at what separates the twins



MCDA twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

Look at what separates the twins



MCDA twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

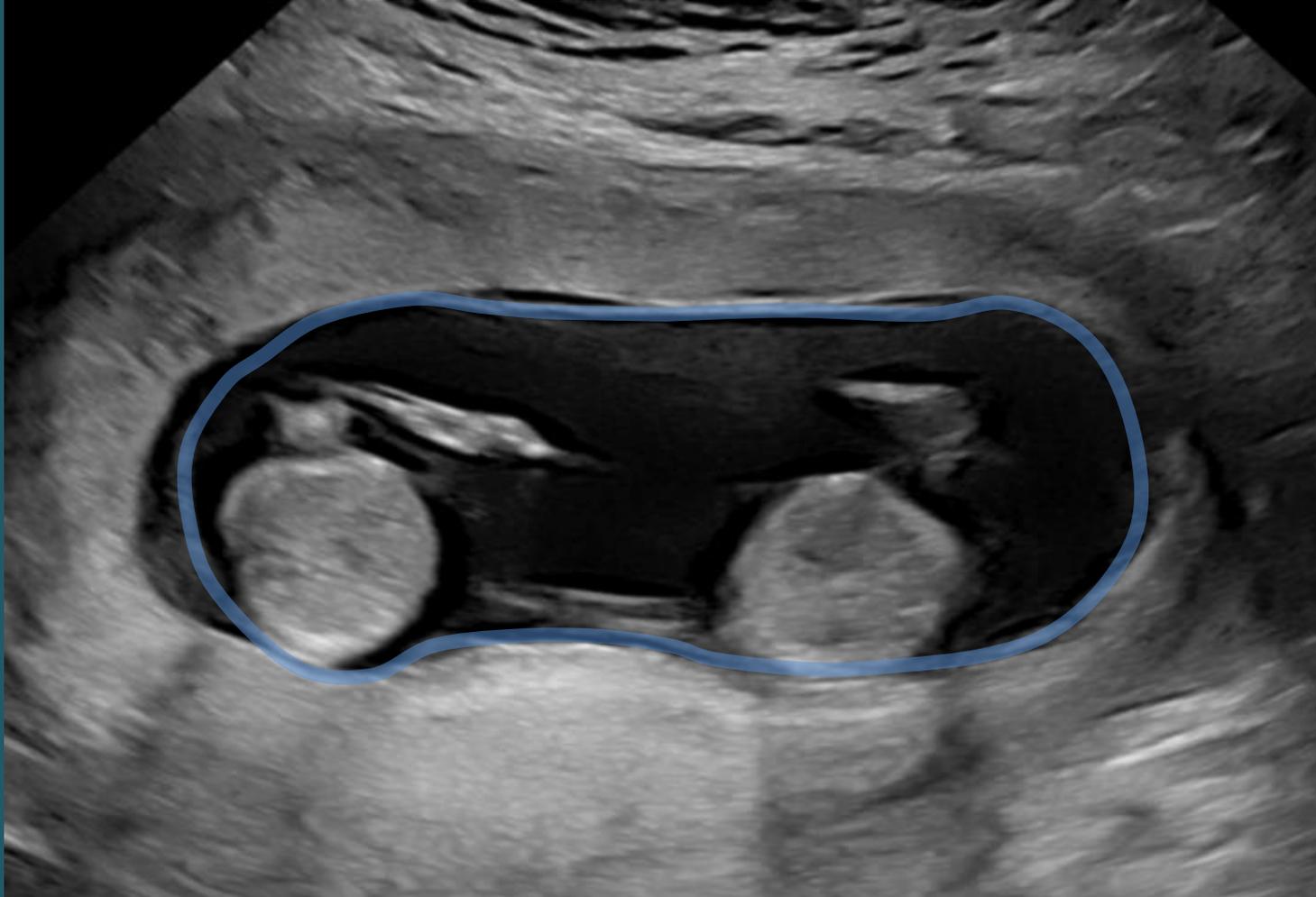
Look at what separates the twins



MCMA twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

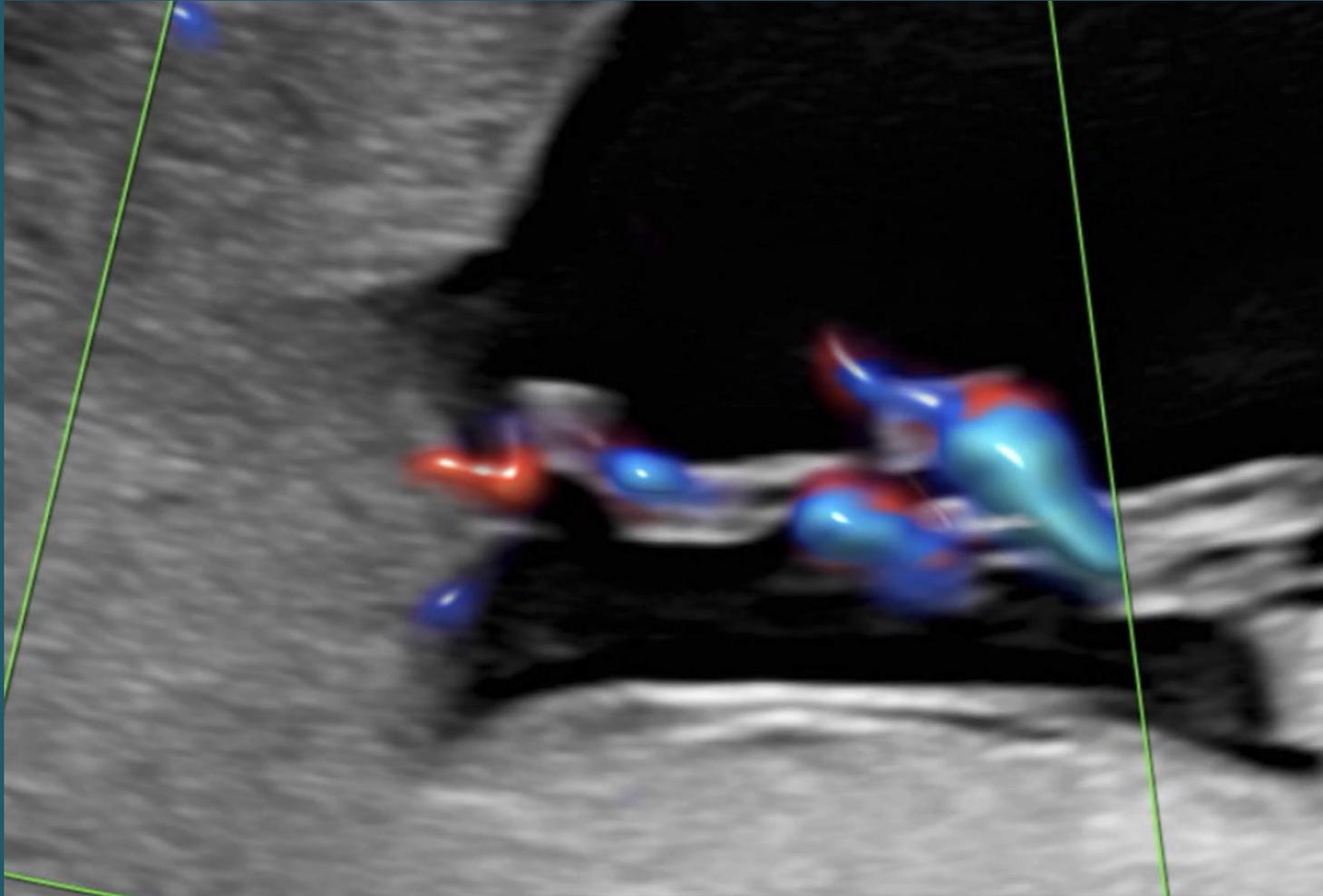
Look at what separates the twins



MCMA twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

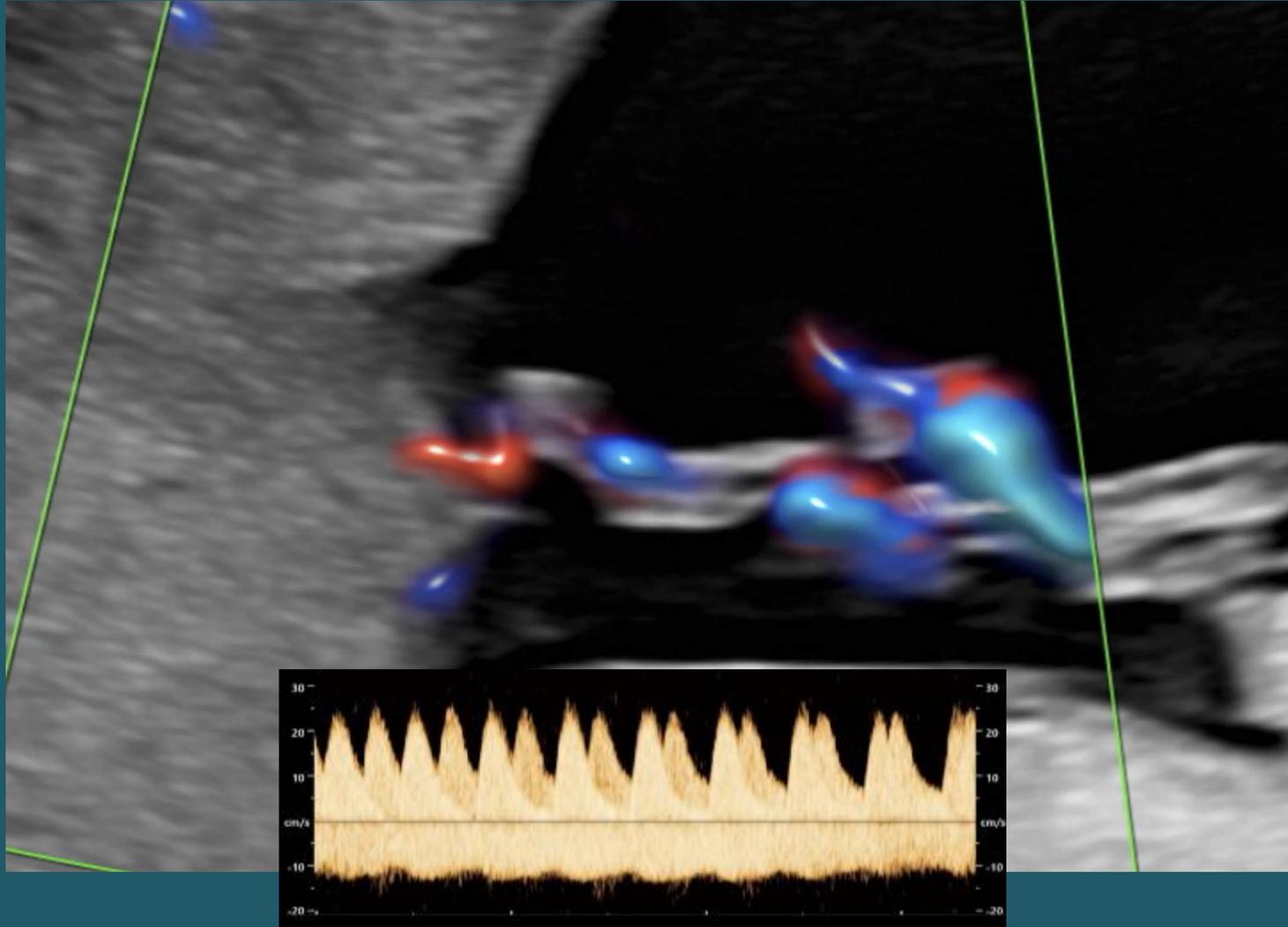
Look at what separates the twins



MCMA twin at 12 weeks

1st trimester: Chorionicity and amnionicity determination

Look at what separates the twins



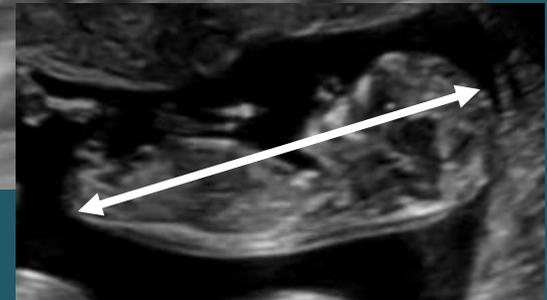
MCMA twin at 12 weeks

1st trimester: Dating



1st trimester: Dating

Use the CRL of the larger twin

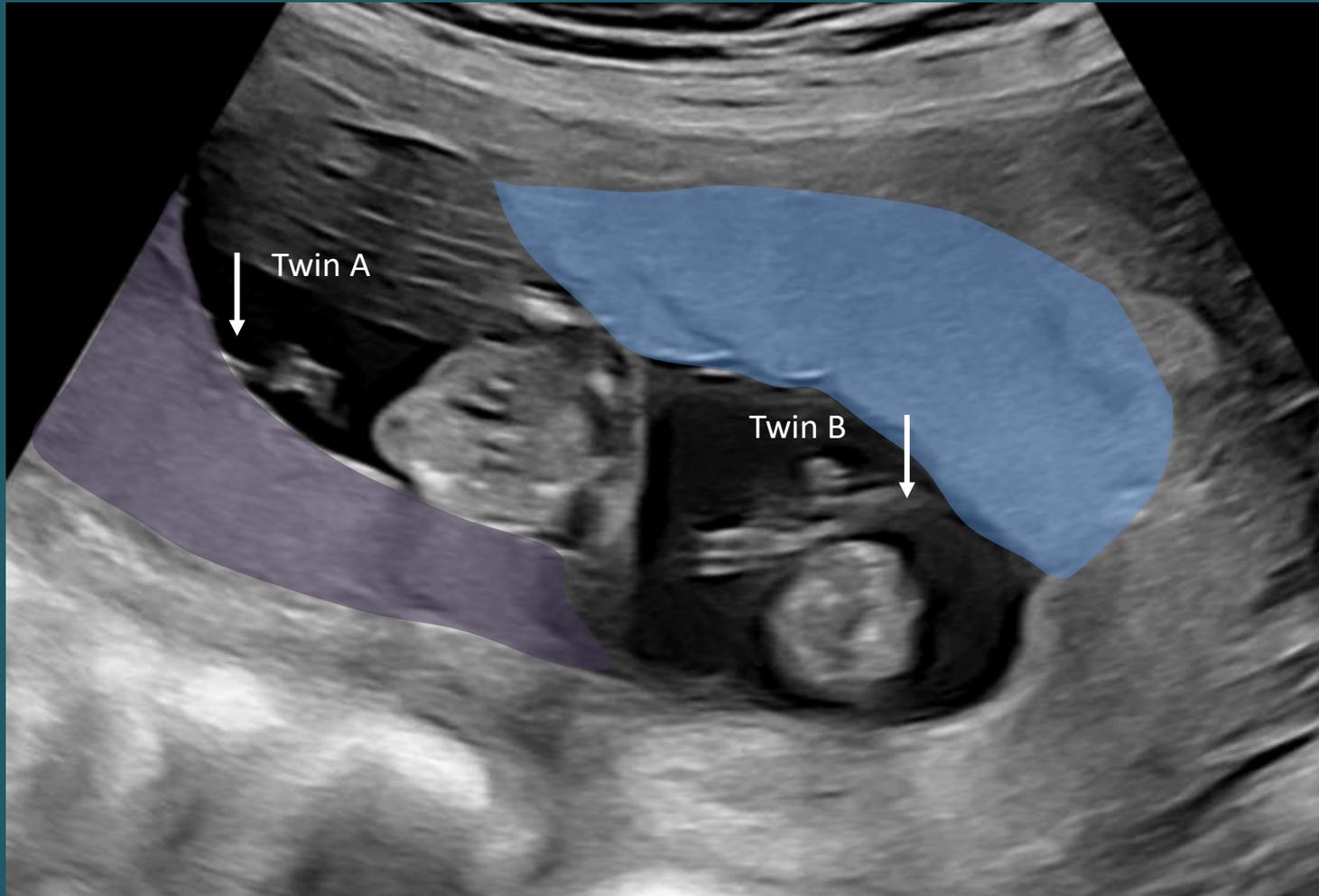


1st trimester: Labelling

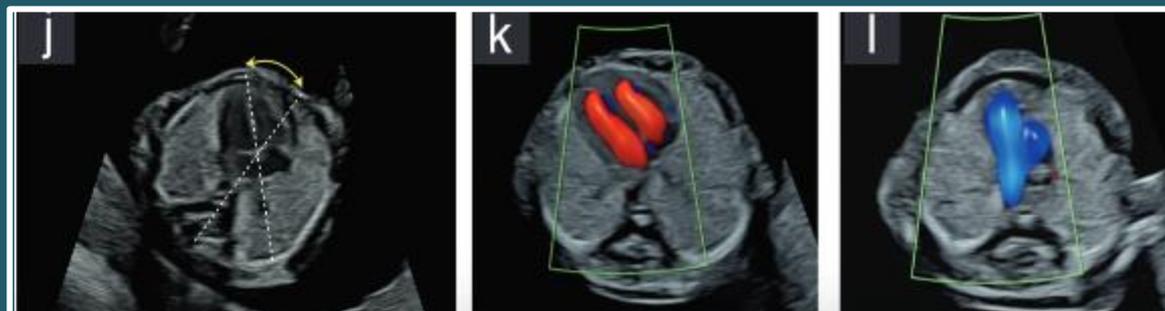
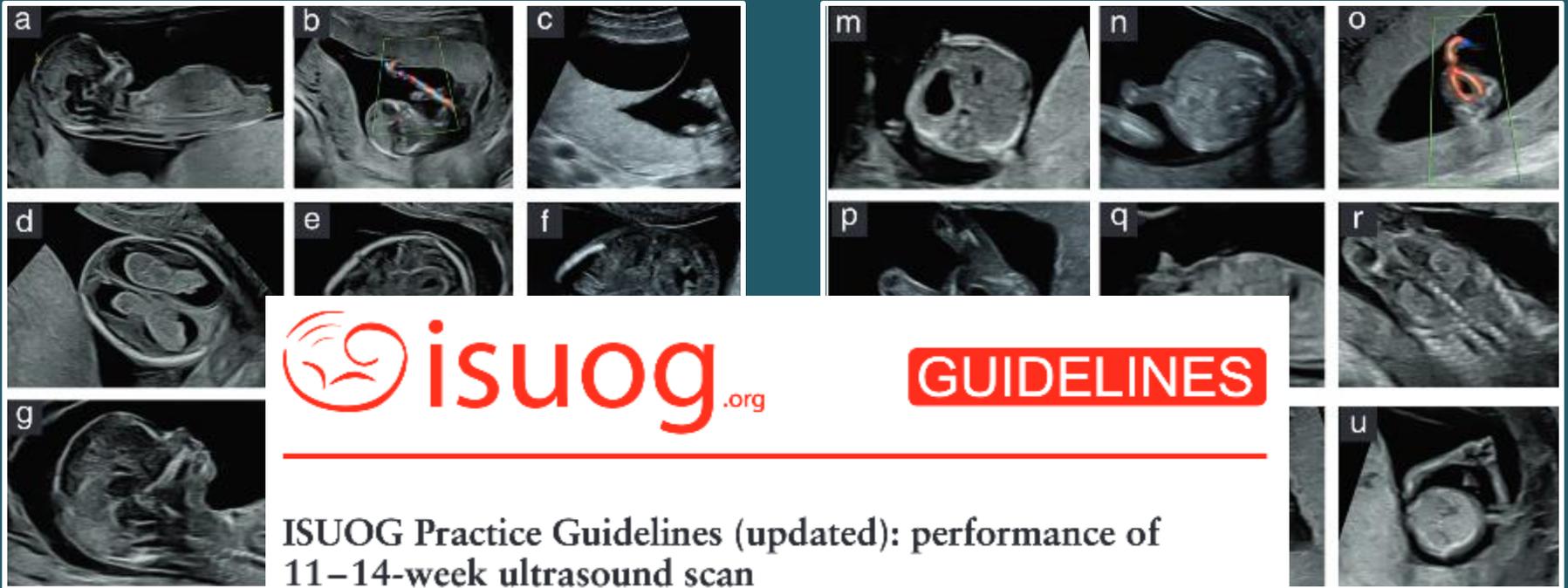


1st trimester: Dating

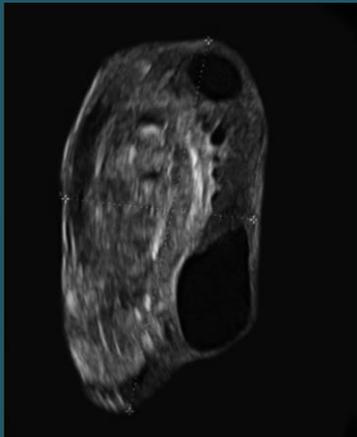
Use the position, placenta and cord insertion



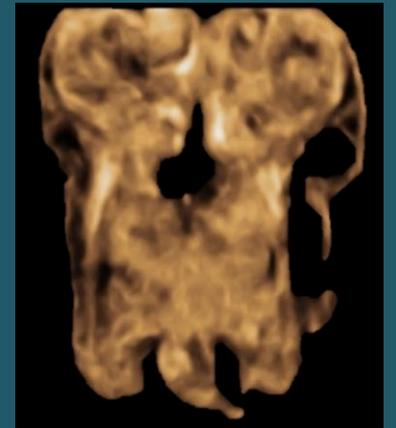
1st trimester: Detailed anatomy scan



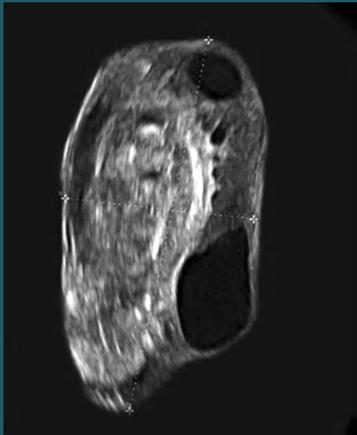
1st trimester: Detailed anatomy scan



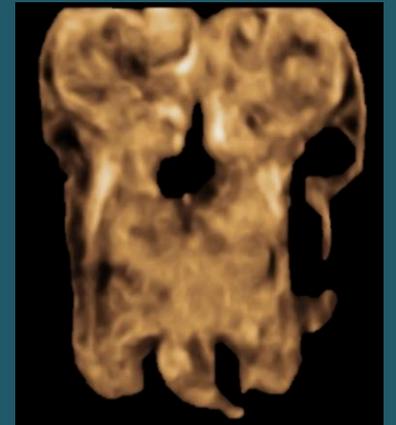
- Singletons 1 in 40
- DC twins 1 in 20 pairs
- MCDA twins 1 in 10 pairs
- MA twins 1 in 5 pairs



1st trimester: Detailed anatomy scan



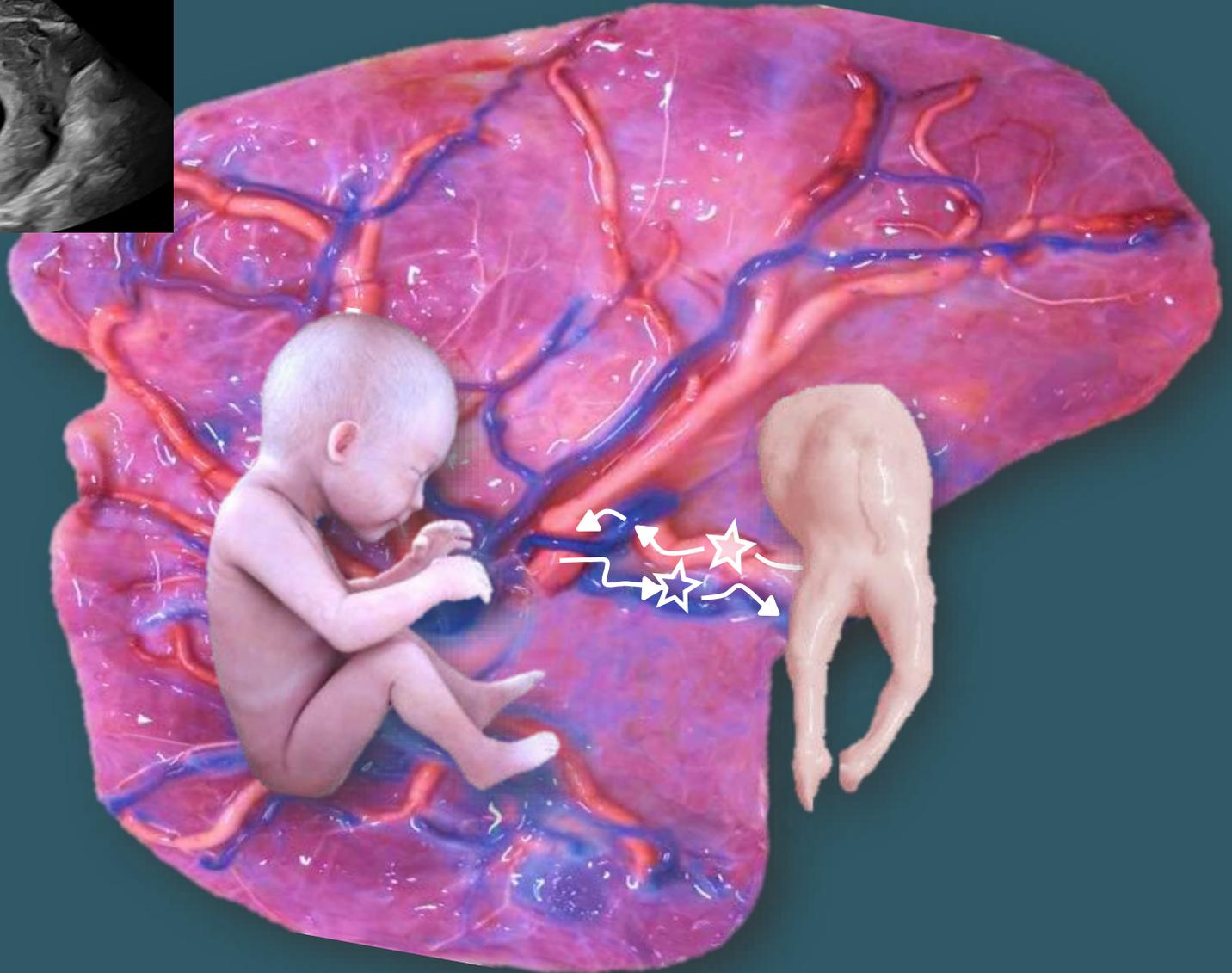
Acardiac twinning



Conjoint twinning

First trimester: 50% of anomalies detected in MC twins and 25% in DC twins

Complication unique to MC twins: Acardiac twinning



1st trimester: Screening for chromosomal anomalies in DC twins

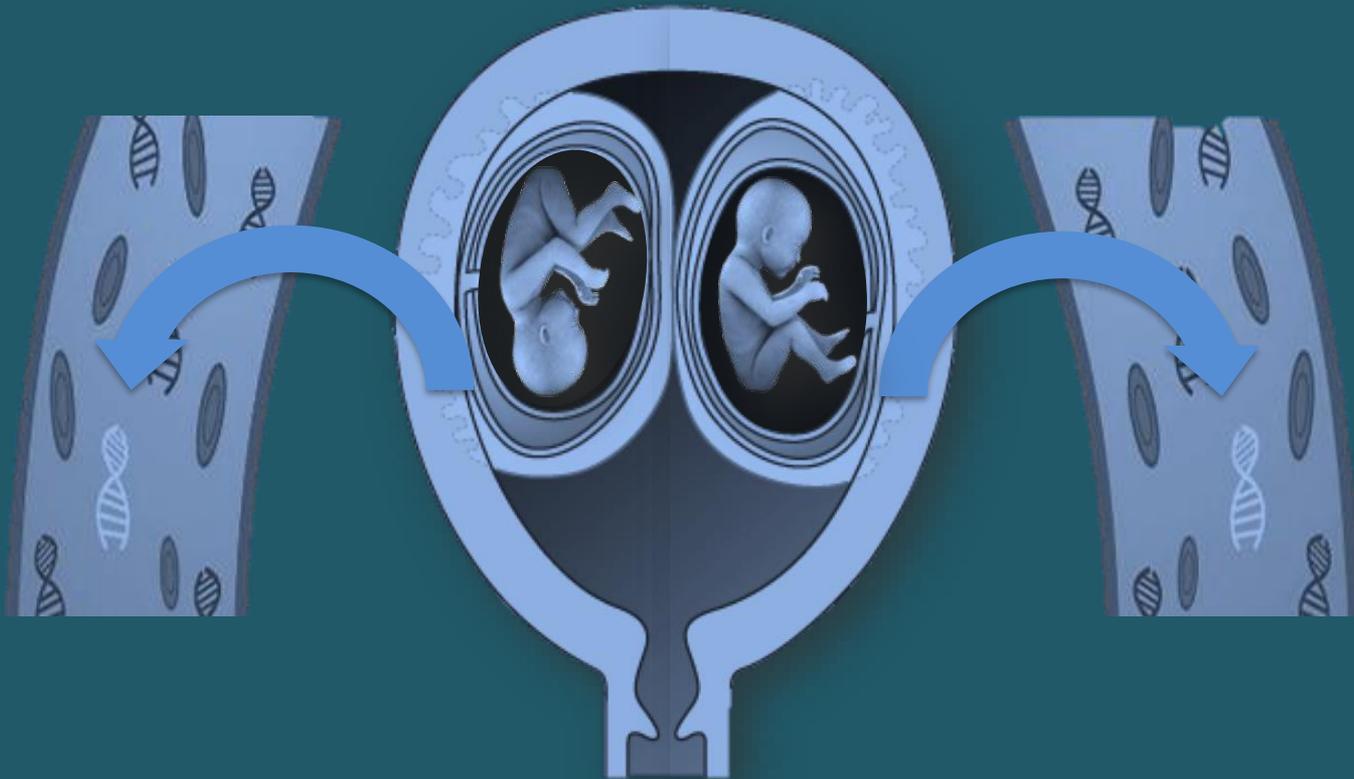
Cell-free fetal DNA testing or non-invasive prenatal testing (NIPT)



T21 DR of 99% at a 0.02% FPR

1st trimester: Screening for chromosomal anomalies in MC twins

Cell-free fetal DNA testing or non-invasive prenatal testing (NIPT)



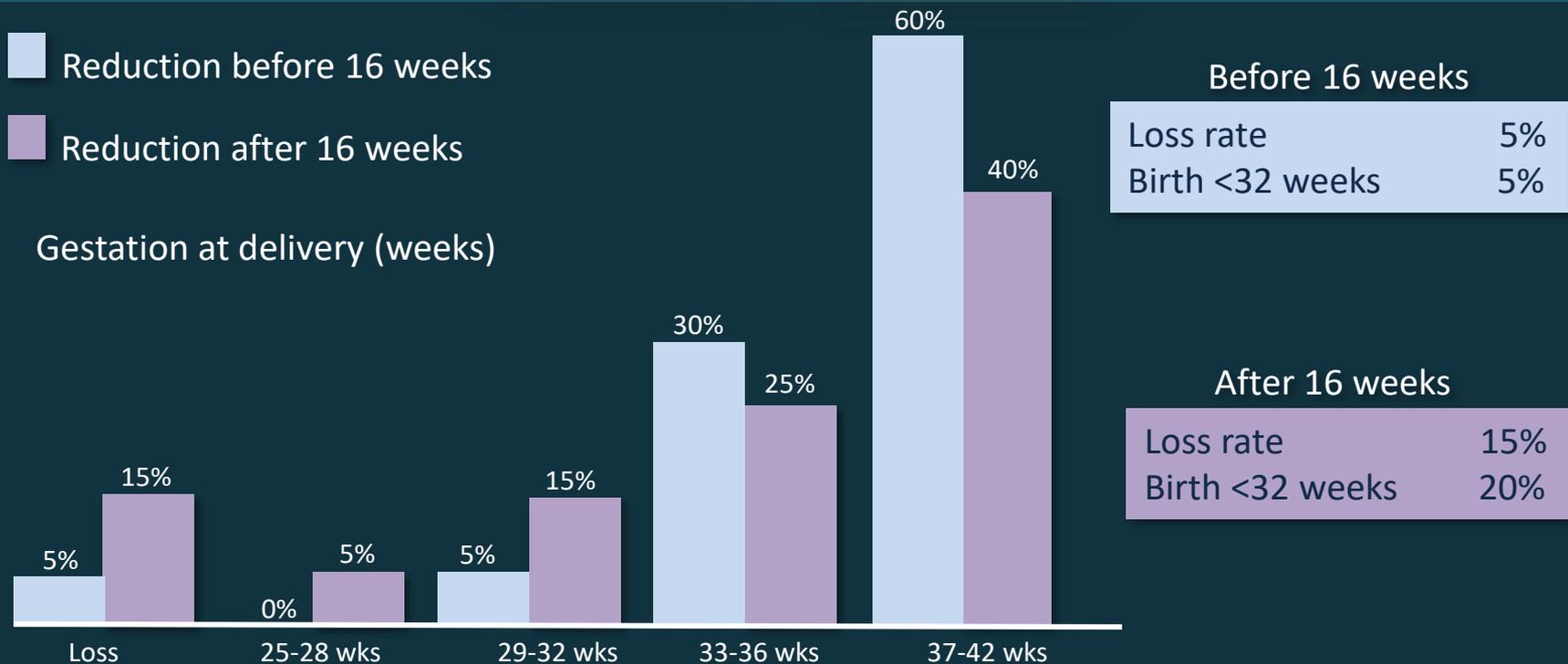
Accuracy in MC twins should be that of singletons but very few data

1st trimester: Diagnosis of chromosomal anomalies

Dual CVS in DC twins allows for a 1st trimester reduction

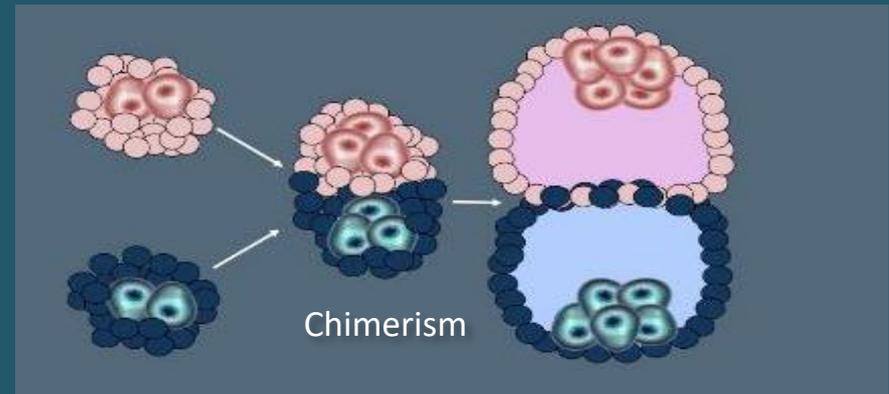
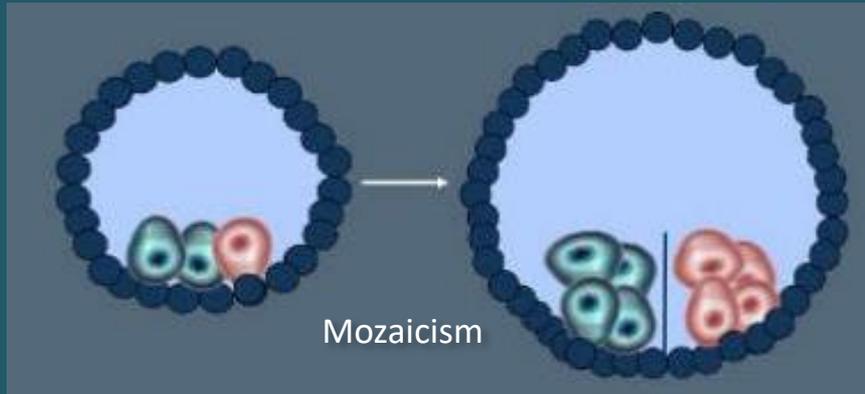


Low procedure-related loss



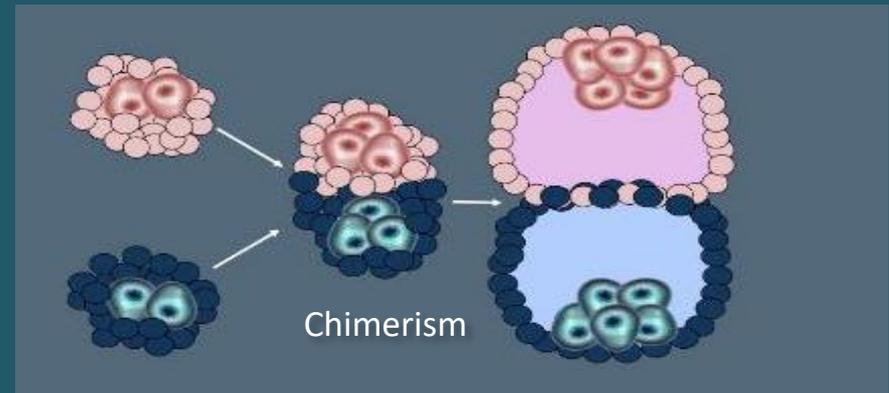
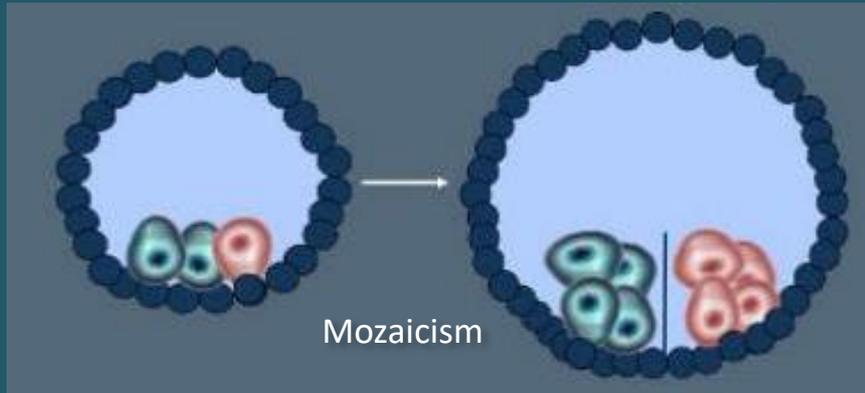
1st trimester: Diagnosis of chromosomal anomalies

Rarely, MC twins may be discordant for chromosomal anomalies



1st trimester: Diagnosis of chromosomal anomalies

Rarely, MC twins may be discordant for chromosomal anomalies



Dual amniocentesis with sampling of both sacs

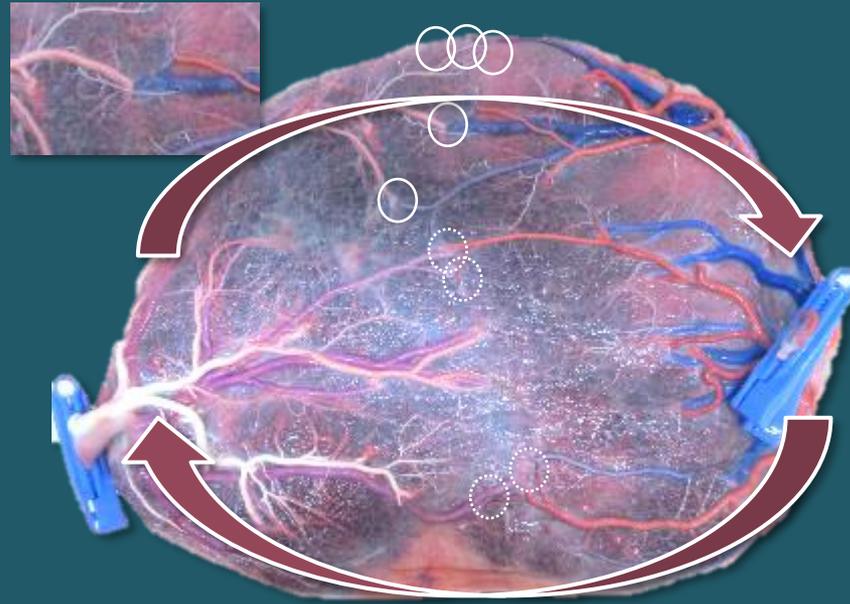


2nd – 3rd trimester

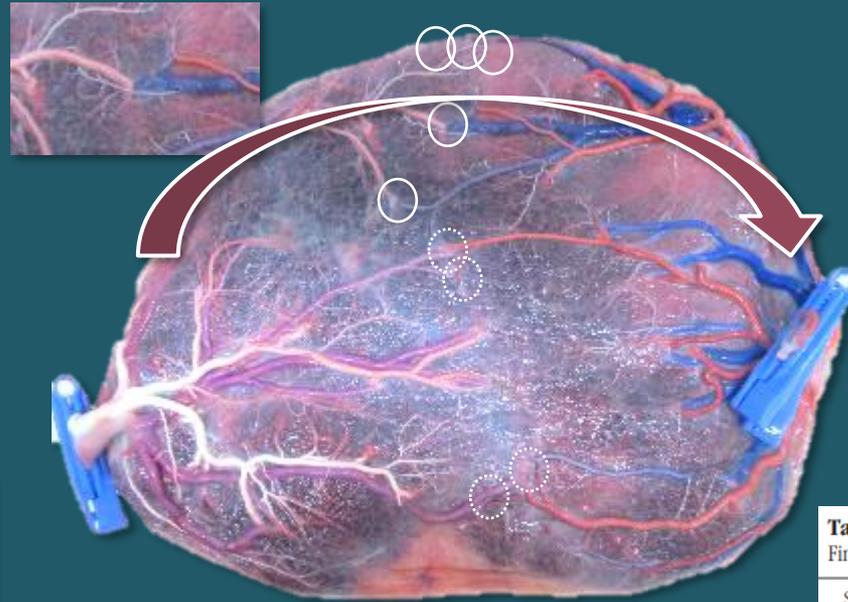


- Screening and treatment of TTTS
- Screening and treatment of TAPS
- Structural anomalies and selective reduction
- Poor growth and its management
- Cervical length screening and prevention of preterm birth

2nd – 3rd trimester: Unique for MCDA twins: Screening for TTTS



2nd – 3rd trimester: Unique for MCDA twins: Screening for TTTS



Progressive

10-15% of MCDA twins
16-26 weeks

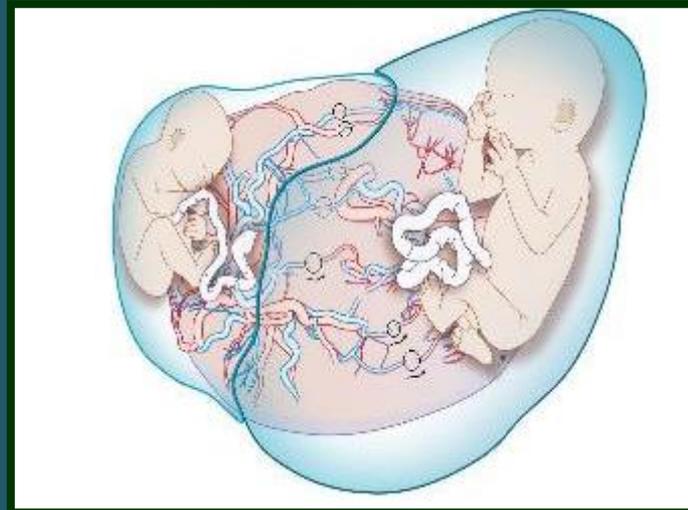


Table 1 Staging of TTTS Based on Sonographic and Doppler Findings

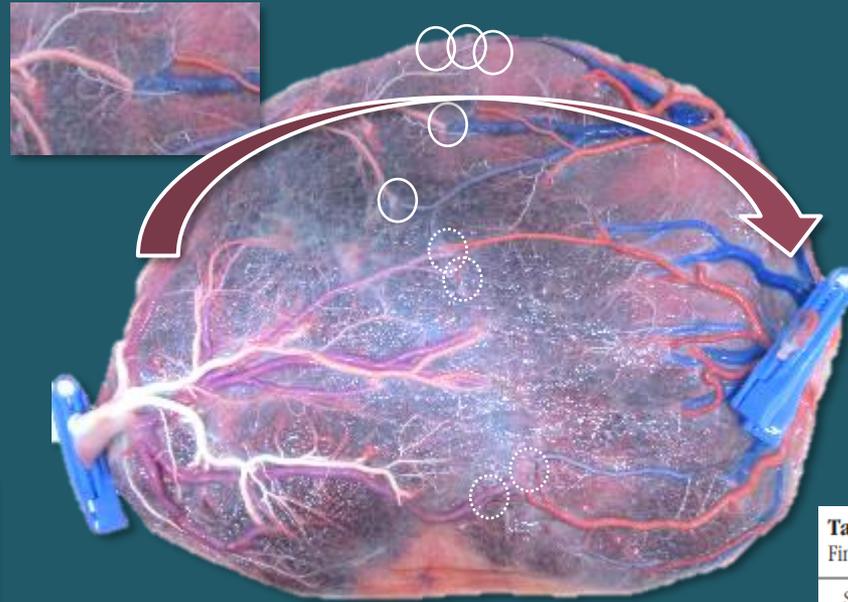
Stage	Poly/ oligohydramnios*	Absent bladder in donor	CADs†	Hydrops	Demise
I	+	-	-	-	-
II	+	+	-	-	-
III	+	+	+	-	-
IV	+	+	+	+	-
V	+	+	+	+	+

*Polyhydramnios, MVP of >8 cm; oligohydramnios, MVP of <2 cm.

†CADs, defined as the presence of at least one of the following: (1) UA AEDW/REDV, (2) RFDV, or (3) pulsatile umbilical venous flow (PUVF).

Atypical presentations
possible!

2nd – 3rd trimester: Unique for MCDA twins: Screening for TTTS



10-15% of MCDA twins
16-26 weeks

Progressive

Table 1 Staging of TTTS Based on Sonographic and Doppler Findings

Stage	Poly/ oligohydramnios*	Absent bladder in donor	CADs†	Hydrops	Demise
I	+	-	-	-	-
II	+	+	-	-	-
III	+	+	+	-	-
IV	+	+	+	+	-
V	+	+	+	+	+

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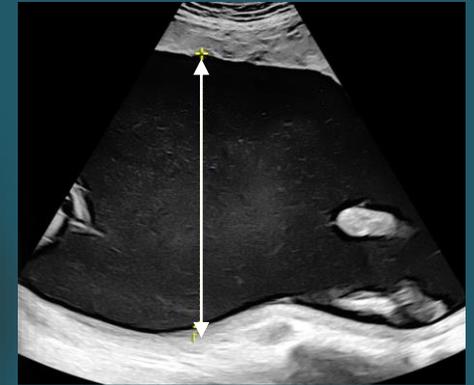
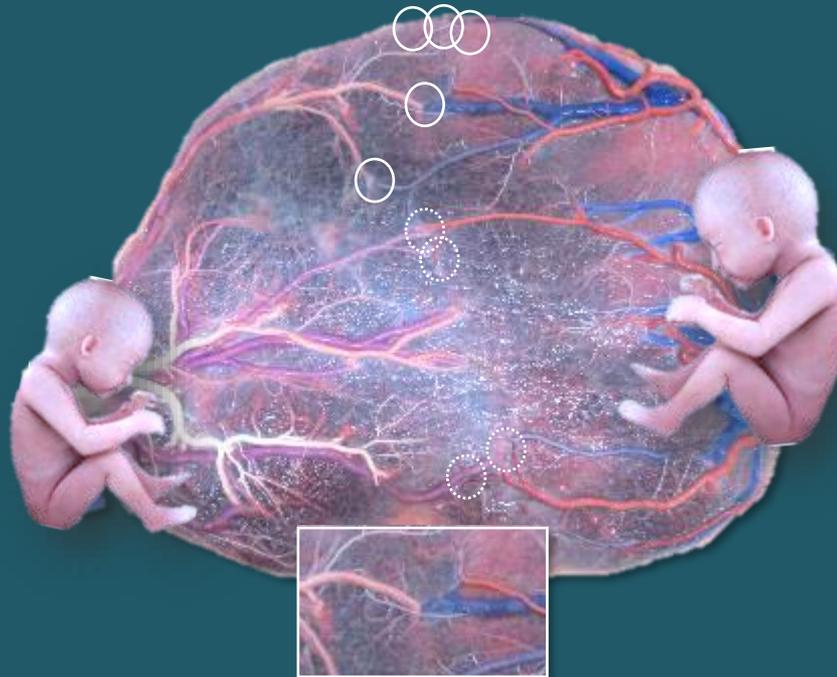
Atypical presentations
possible!

TTTS is the main cause of death or handicap in MC twins

2nd – 3rd trimester: Unique for MCDA twins: Screening for TTTS



$\leq 2\text{cm}$



$\geq 6\text{ cm}$	$< 16\text{ weeks}$
$\geq 8\text{ cm}$	$16\text{ to }20\text{ weeks}$
$\geq 10\text{ cm}$	$> 20\text{ weeks}$



2nd – 3rd trimester: Unique for MCDA twins: Screening for TTTS

Look for the intertwin septum and bladders

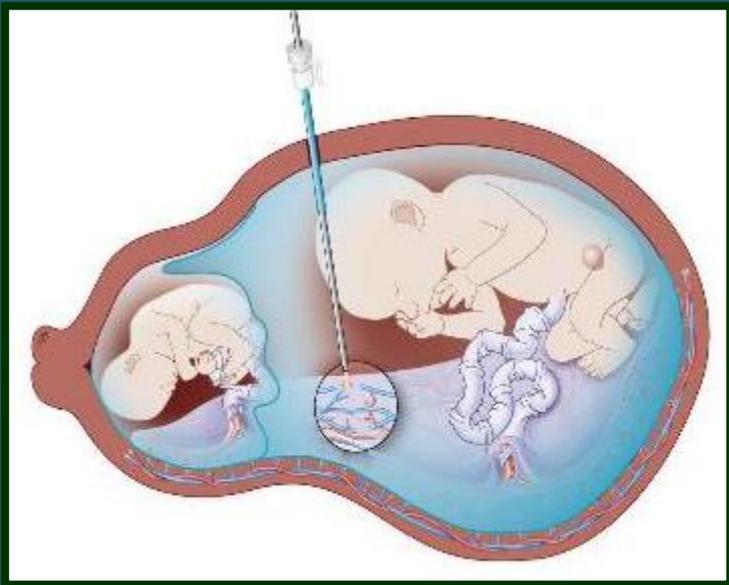
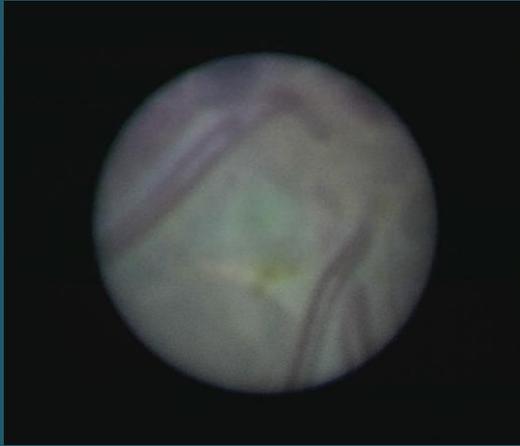


2nd – 3rd trimester: Unique for MCDA twins: Screening for TTTS

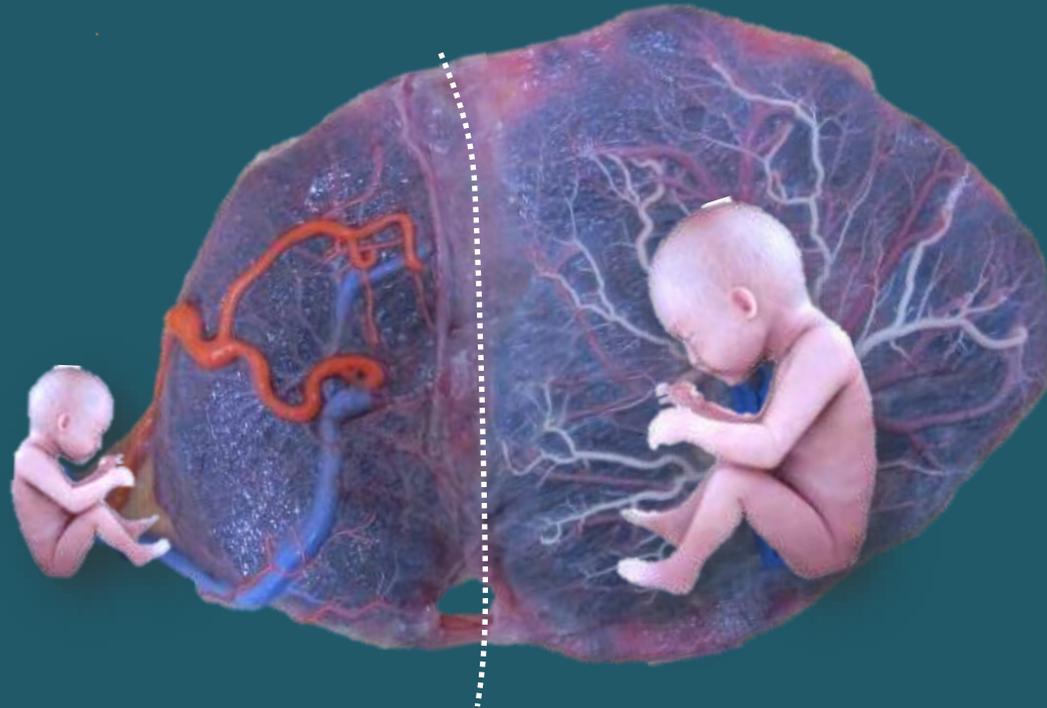
Look for the intertwin septum and bladders



2nd – 3rd trimester: Treatment of TTTS



2nd – 3rd trimester: Treatment of TTTS

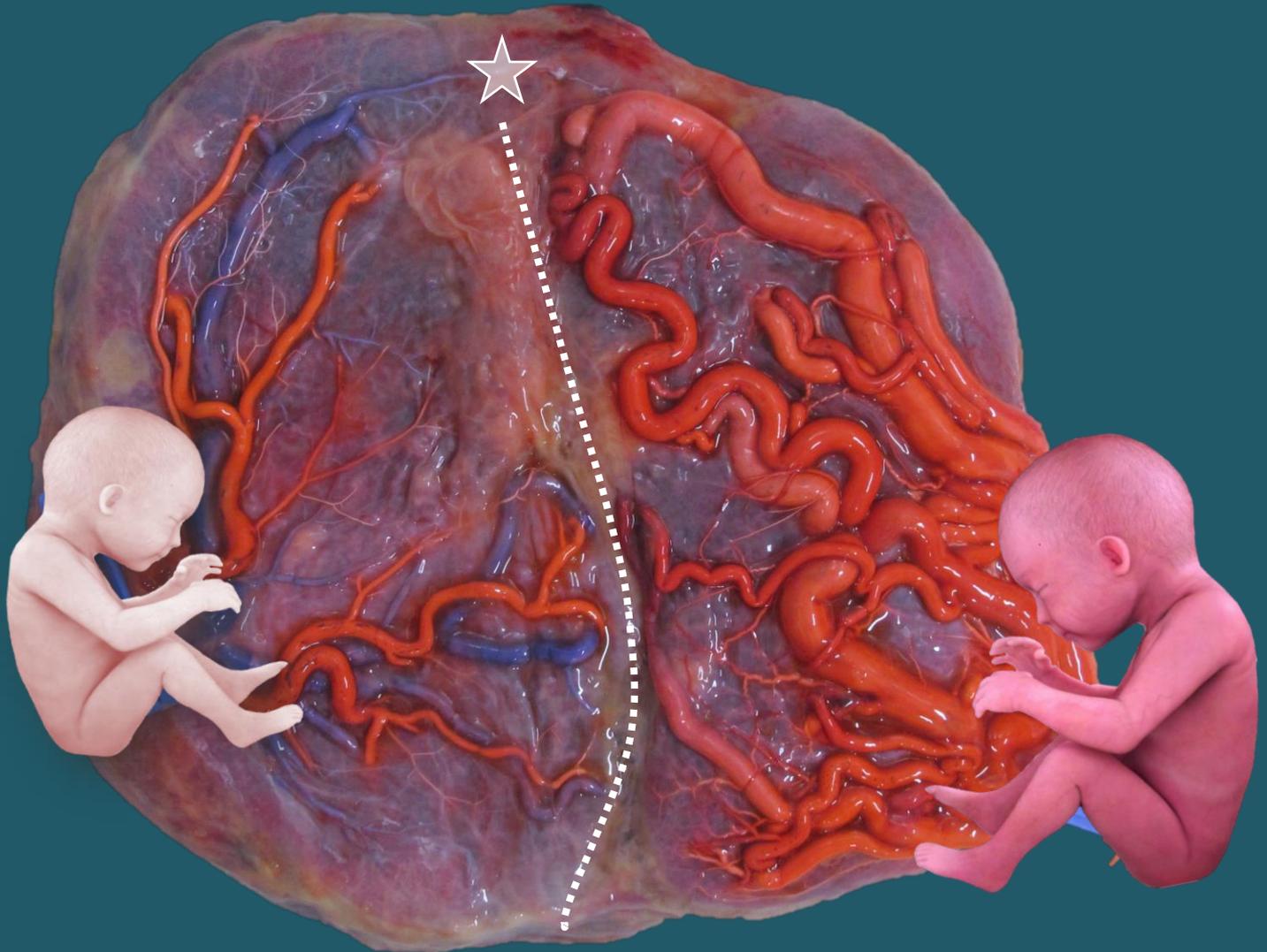


2 survivors	65%
1 survivor	20%
0 survivors	15%

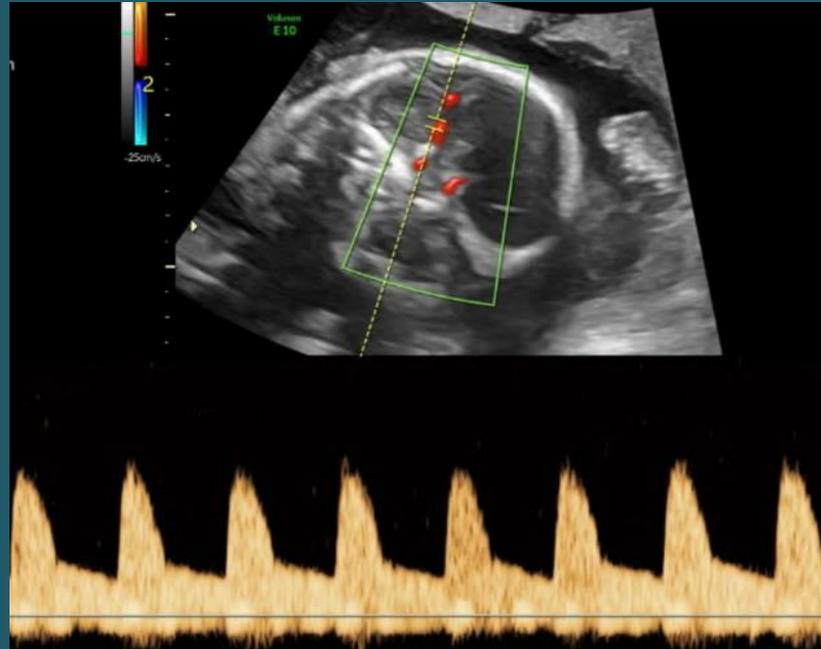
GA at birth 32 weeks, overall 10% risk of impairment

2nd – 3rd trimester: Treatment of TTTS

Missed anastomoses may result in TAPS

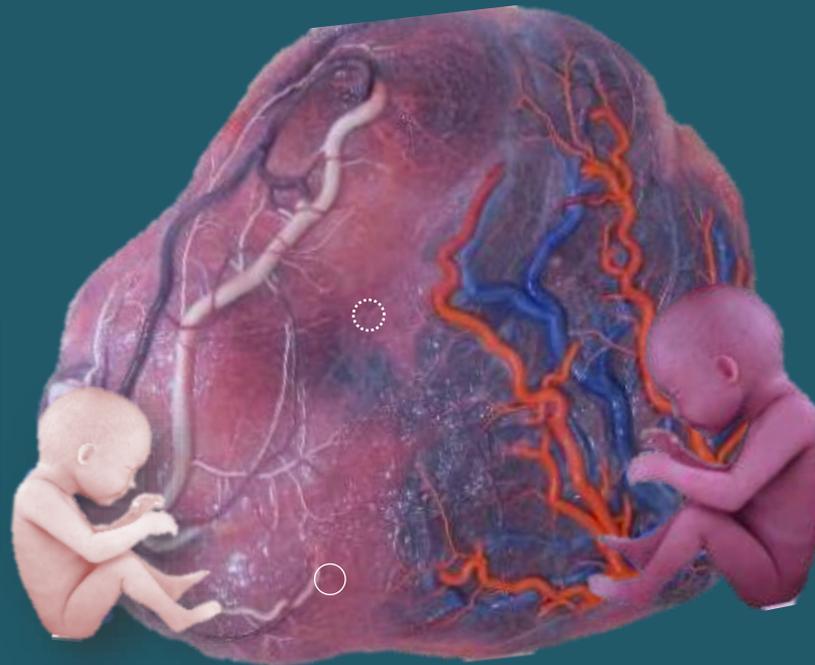


2nd – 3rd trimester: Unique for MCDA twins: Screening for TAPS



2nd – 3rd trimester: Unique for MCDA twins: Screening for TAPS

5% of MCDA twins
Often >26 weeks

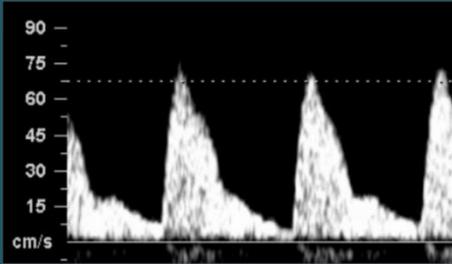


Progressive

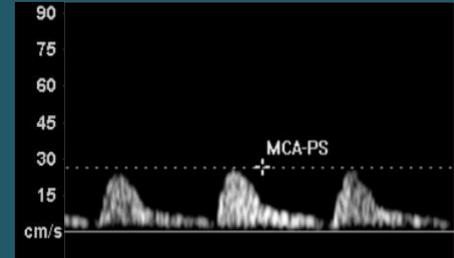
Antenatal stage	Findings at Doppler ultrasound examination
Stage 1	MCA-PSV donor >1.5 MoM <i>and</i> MCA-PSV recipient <1.0 MoM, without other signs of fetal compromise
Stage 2	MCA-PSV donor >1.7 MoM <i>and</i> MCA-PSV recipient <0.8 MoM, without other signs of fetal compromise
Stage 3	as stage 1 or 2, with cardiac compromise of donor, defined as critically abnormal flow ^a
Stage 4	hydrops of donor
Stage 5	intrauterine demise of one or both fetuses preceded by TAPS

Postnatal stage	Intertwin Hb difference, g/dl
Stage 1	>8.0
Stage 2	>11.0
Stage 3	>14.0
Stage 4	>17.0
Stage 5	>20.0

2nd – 3rd trimester: Unique for MCDA twins: Screening for TAPS

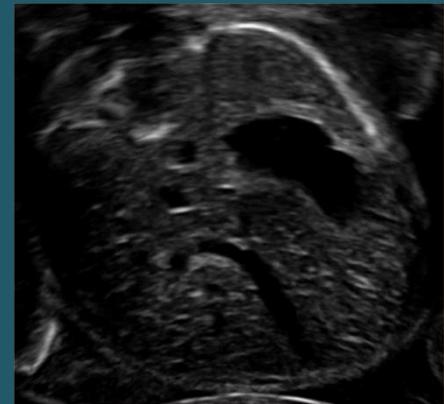
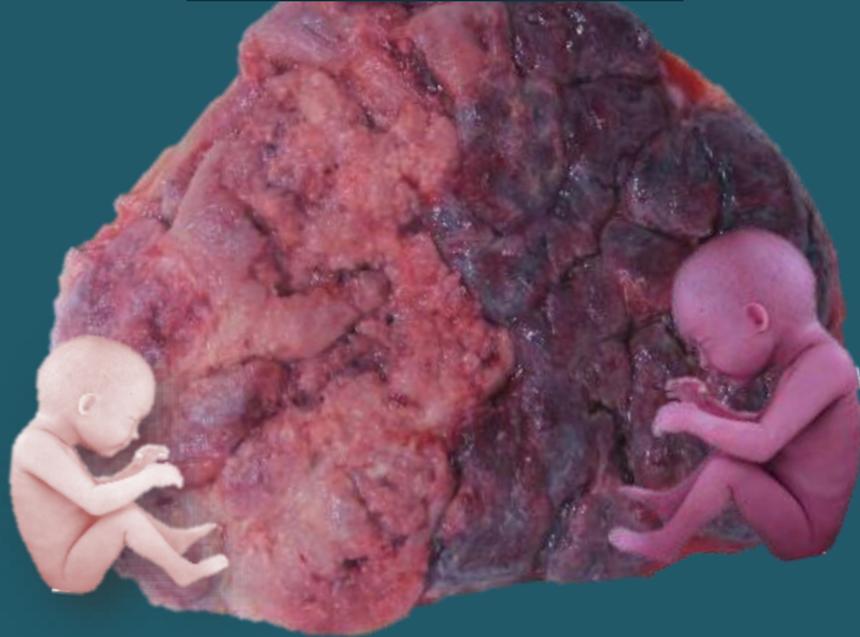


Δ MCA-PSV \geq 1 MoM



MCA-PSV \geq 1.5 MoM

MCA-PSV \leq 0.8 MoM



Most have placental dichotomy

2nd – 3rd trimester: management of TAPS

Individualized management

Gestational age



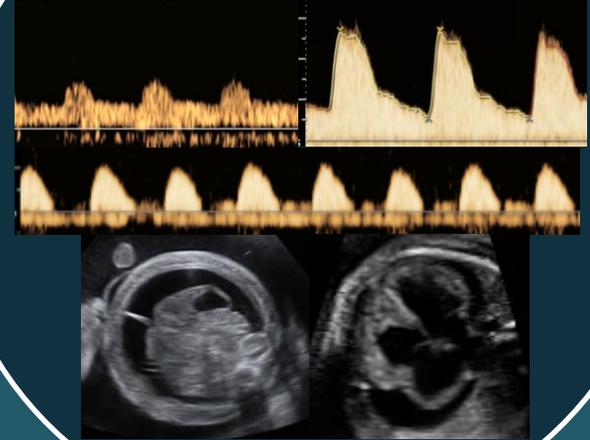
2nd – 3rd trimester: management of TAPS

Individualized management

Gestational age



Severity



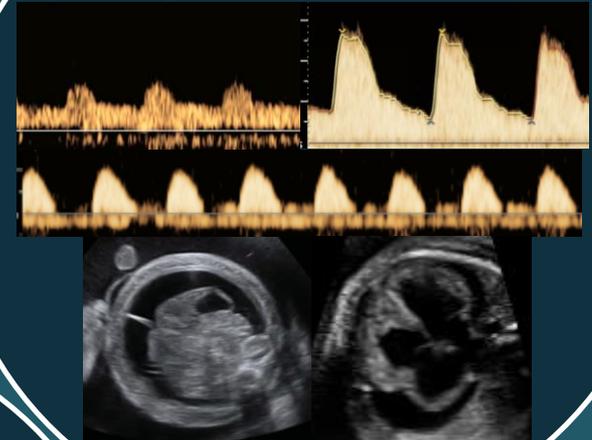
2nd – 3rd trimester: management of TAPS

Individualized management

Gestational age



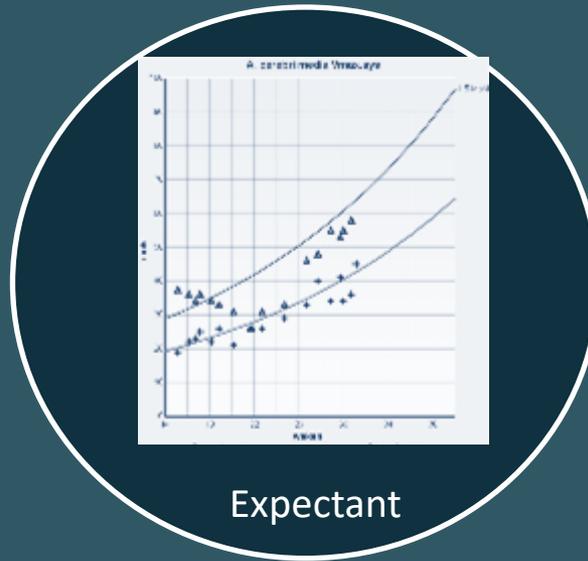
Severity



Technical aspects



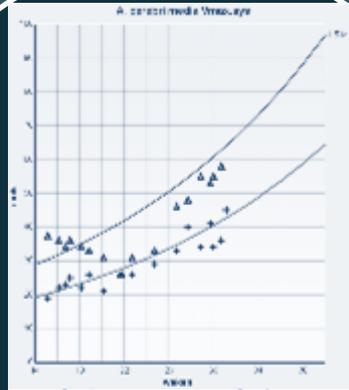
2nd – 3rd trimester: management of TAPS



2nd – 3rd trimester: management of TAPS



Fetoscopic laser surgery

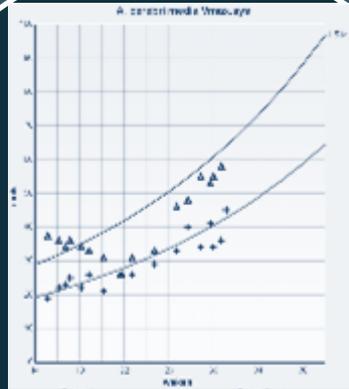


Expectant

2nd – 3rd trimester: management of TAPS



Fetoscopic laser surgery



Expectant

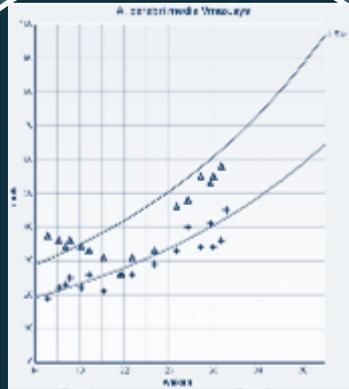


Selective reduction

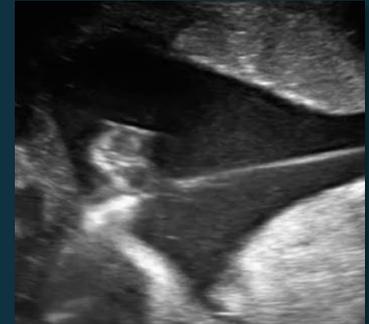
2nd – 3rd trimester: management of TAPS



Fetoscopic laser surgery



Expectant



Selective reduction

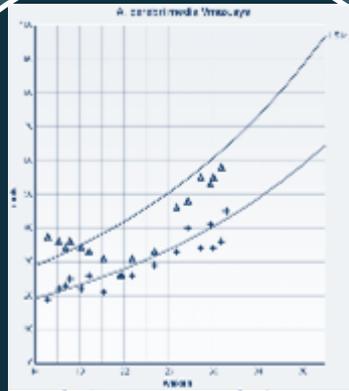


Intrauterine transfusion

2nd – 3rd trimester: management of TAPS



Fetoscopic laser surgery



Expectant



Selective reduction

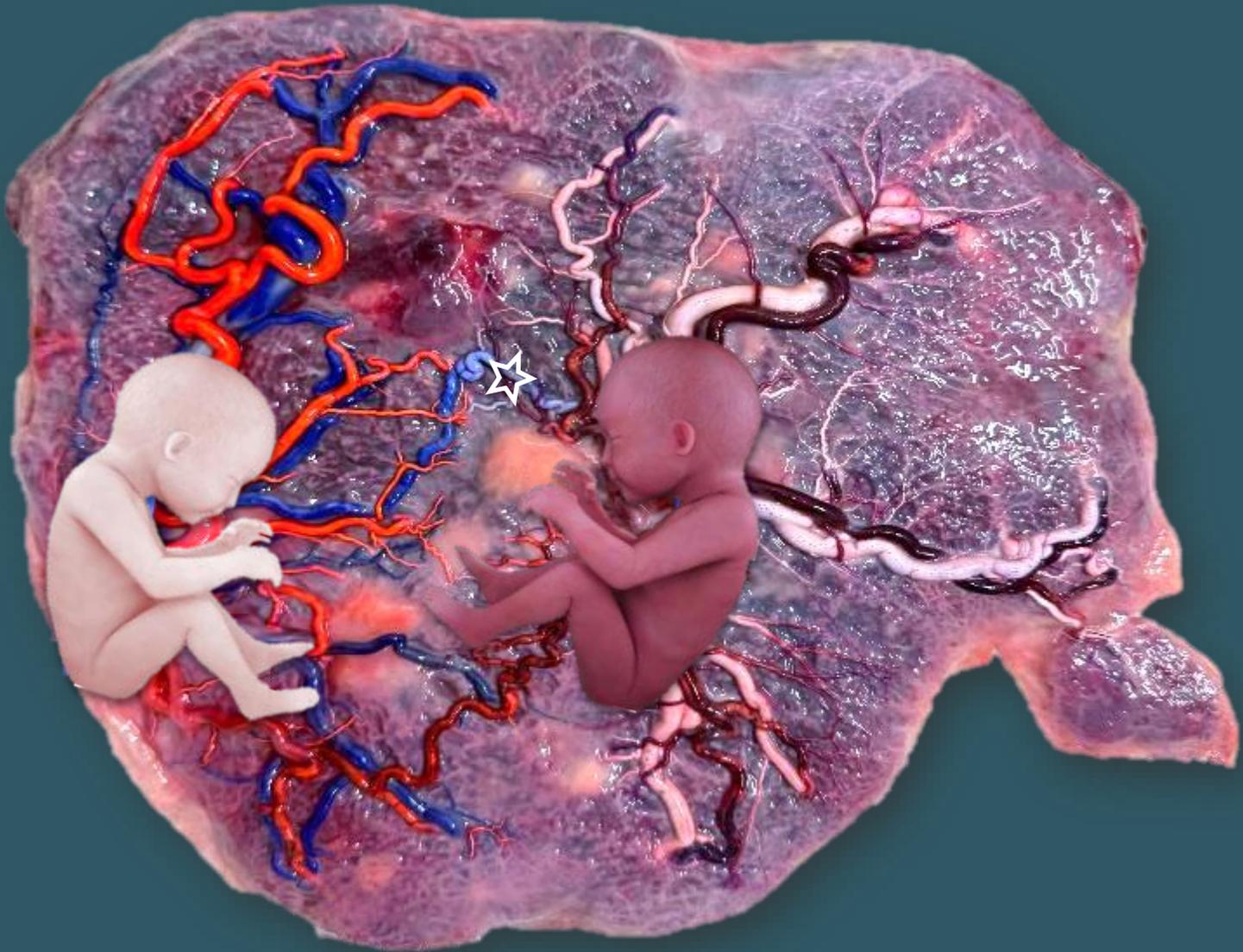


Intrauterine transfusion

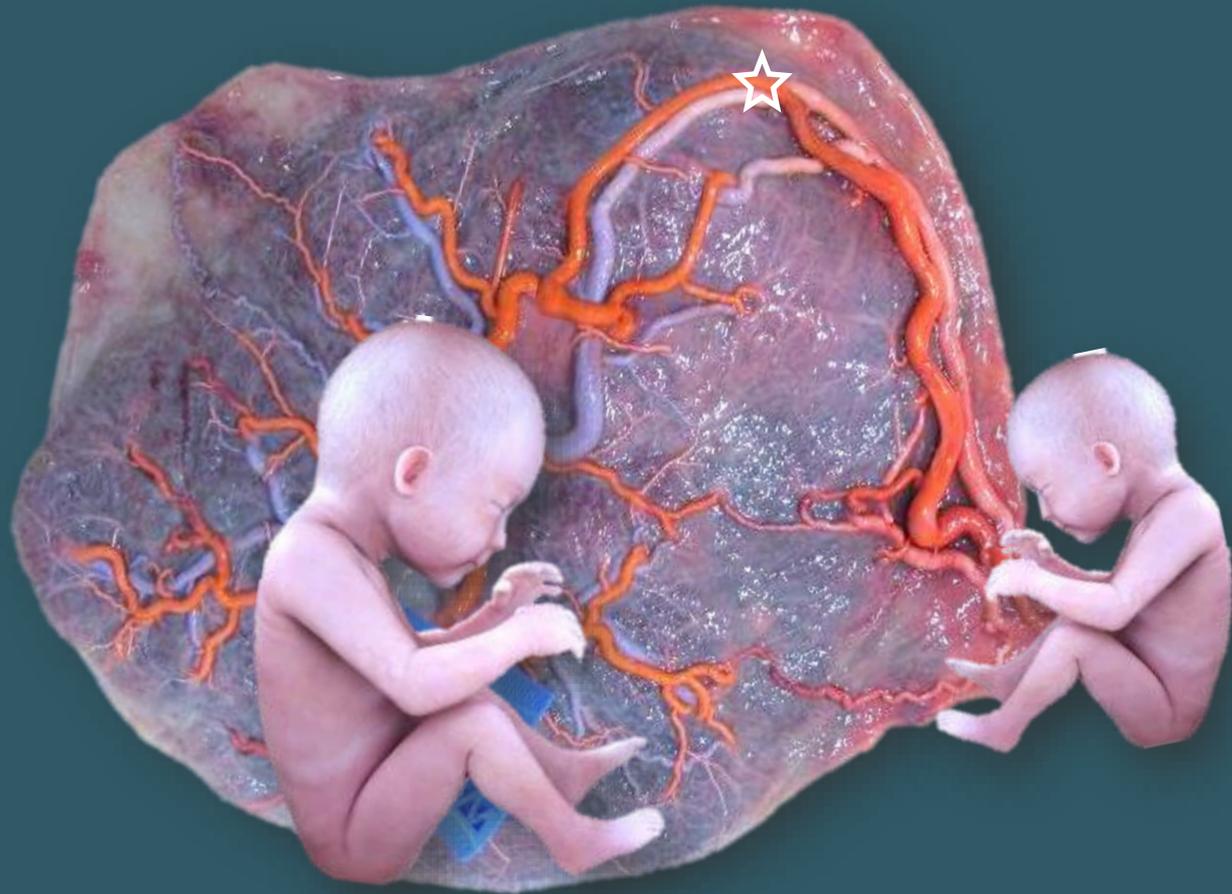


Elective preterm birth

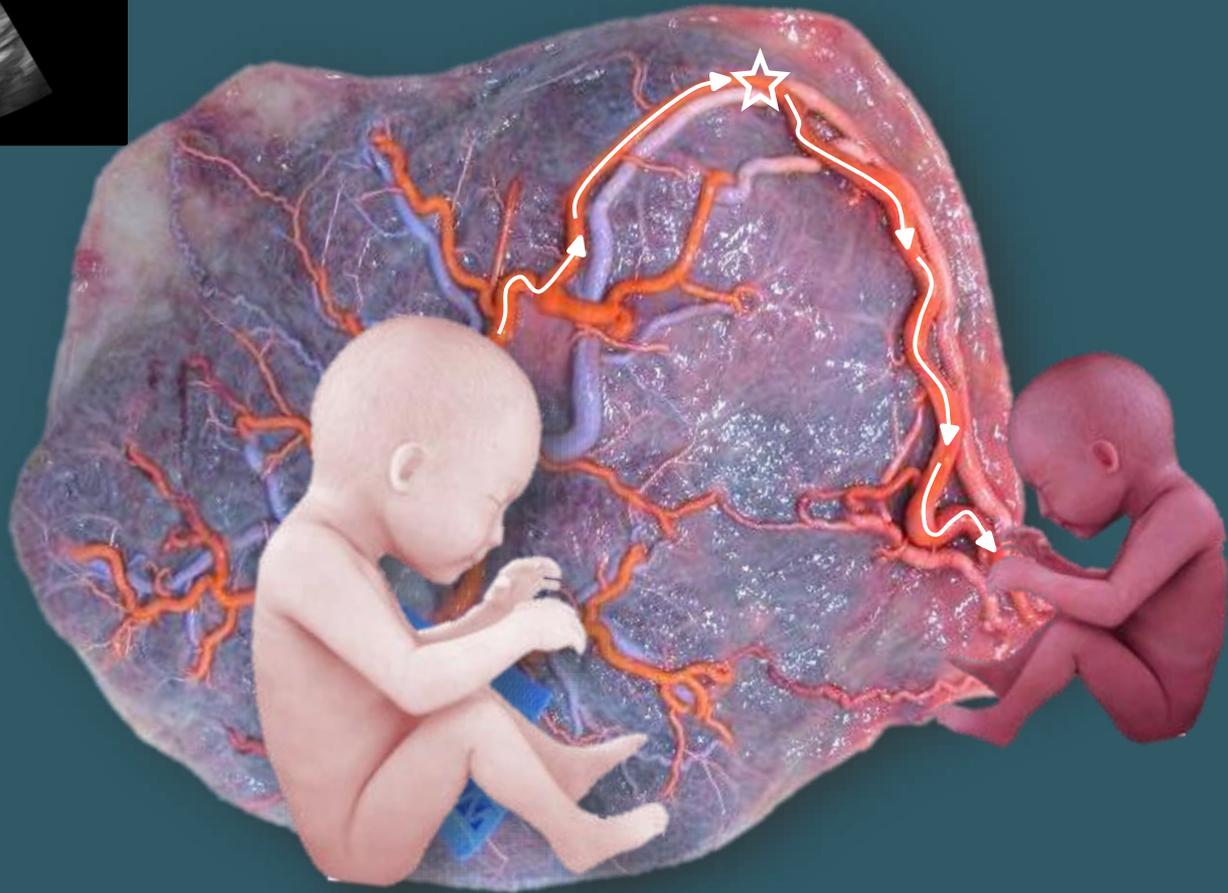
2nd – 3rd trimester: Unique for MCDA twins: Acute intertwin transfusion



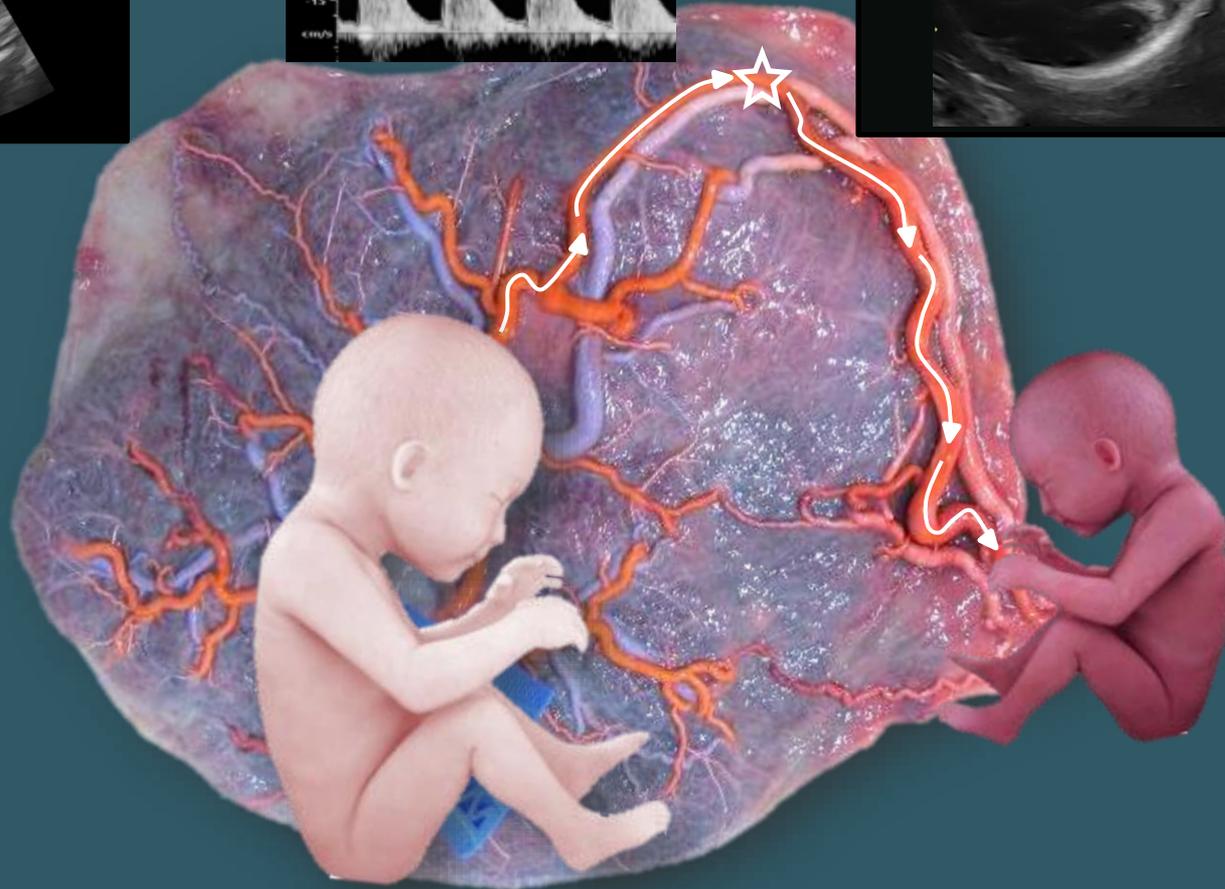
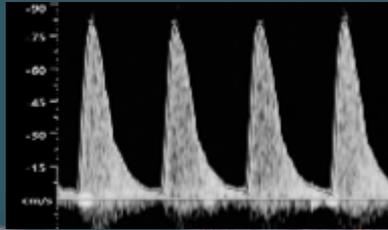
2nd – 3rd trimester: Unique for MCDA twins: Acute intertwin transfusion



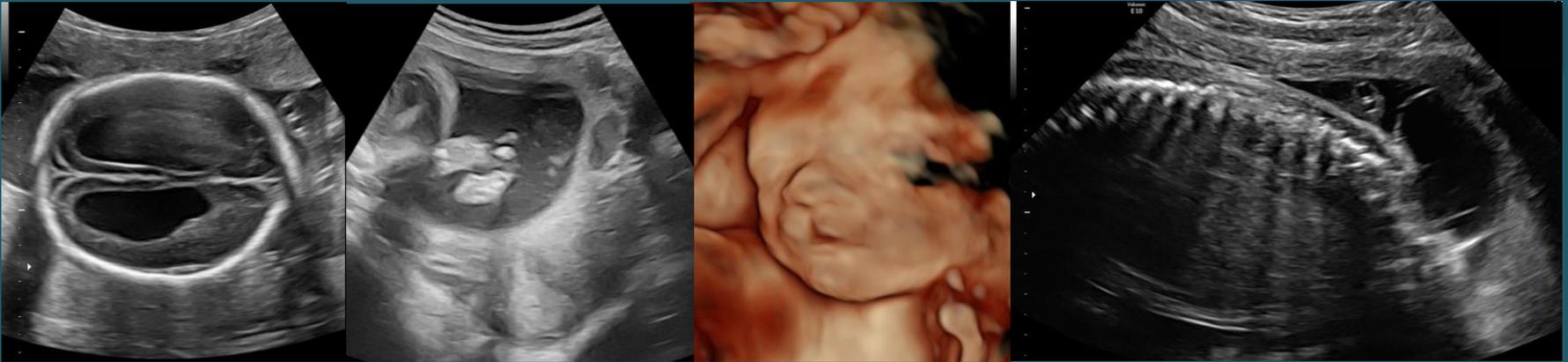
2nd – 3rd trimester: Unique for MCDA twins: Acute intertwin transfusion



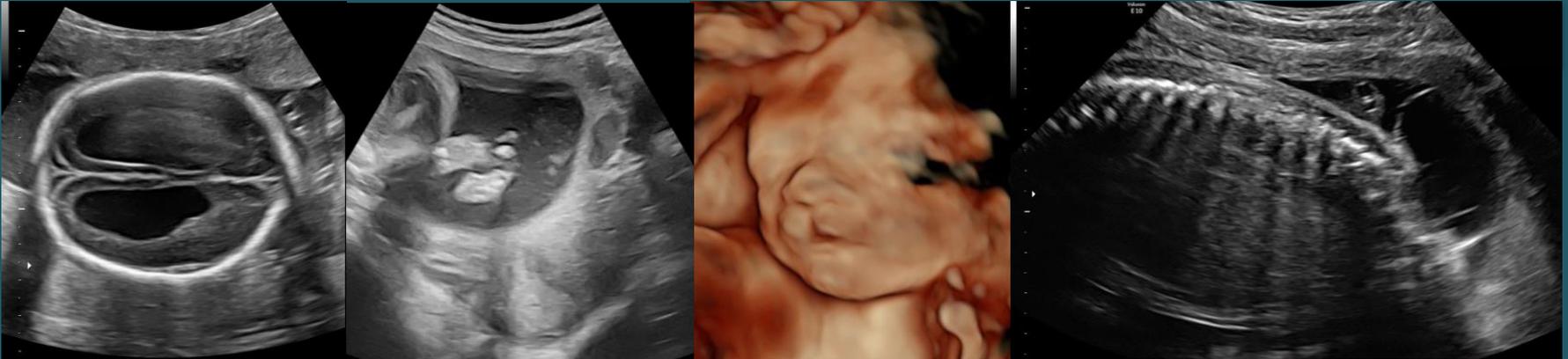
2nd – 3rd trimester: Unique for MCDA twins: Acute intertwin transfusion



2nd – 3rd trimester: Screening for structural anomalies



2nd – 3rd trimester: Screening for structural anomalies



GUIDELINES

ISUOG Practice Guidelines (updated): performance of the routine mid-trimester fetal ultrasound scan

2nd – 3rd trimester: Selective reduction in DC twins



1st TRIMESTER DIAGNOSIS

1st trimester REDUCTION

5% loss rate
5% birth before 32 weeks



2nd – 3rd trimester: Selective reduction in DC twins



1st TRIMESTER DIAGNOSIS

1st trimester REDUCTION

5% loss rate
5% birth before 32 weeks



2nd TRIMESTER DIAGNOSIS

NONLETHAL anomaly

hydrocephaly, spina bifida

2nd trimester REDUCTION

15% loss rate
20% birth before 32 weeks

LATE REDUCTION Between 28-30 weeks

No procedure-related loss
Small risk birth before 32 weeks

LETHAL in fetal life

Severe Ebstein's, lethal monosomy X

EXPECTANT management

No procedure-related loss
No iatrogenic preterm birth

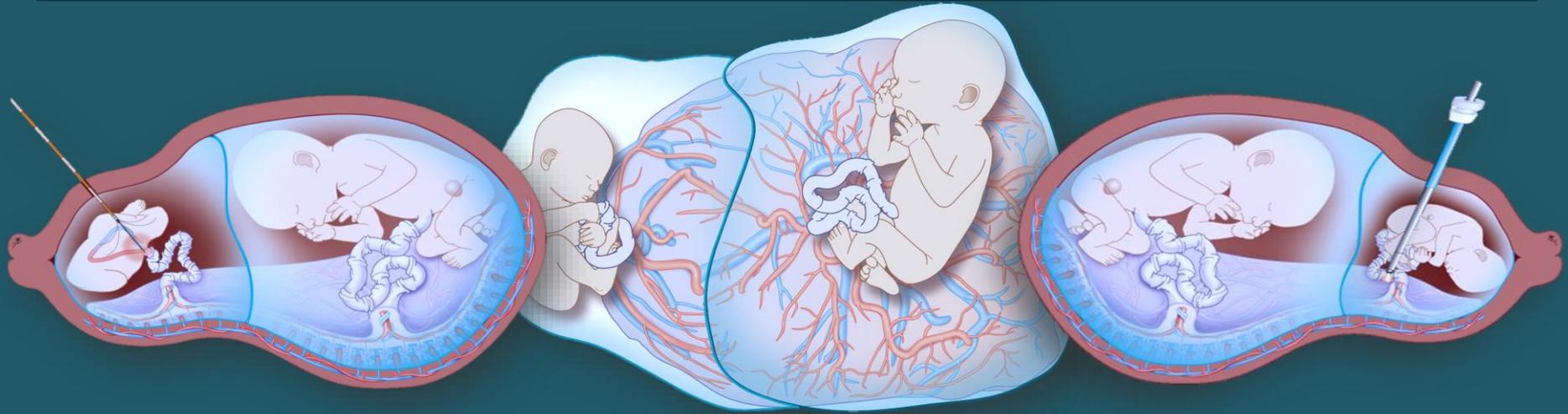
LETHAL in early neonatal period

HLHS, bilateral renal agenesis

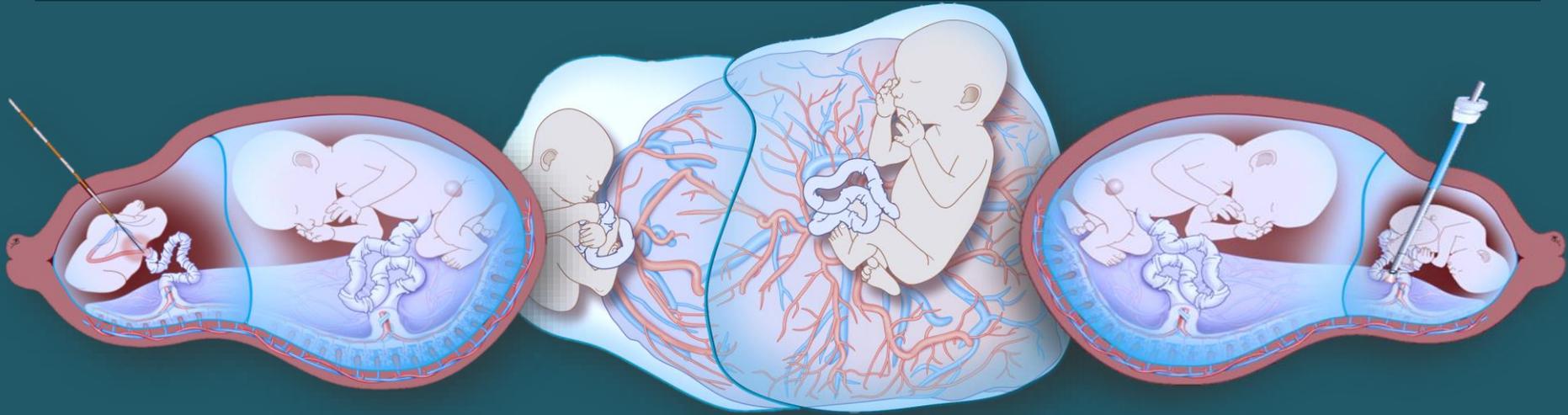
COMFORT CARE

No procedure-related loss
No iatrogenic preterm birth

2nd – 3rd trimester: Selective reduction in MC twins



2nd – 3rd trimester: Selective reduction in MC twins



RISK co-twin

TRAP, lethal monosomy X, Ebstein's, TTTS

IMMEDIATE REDUCTION

From 16 weeks onward

80% survival

10% fetal demise

10% miscarriage



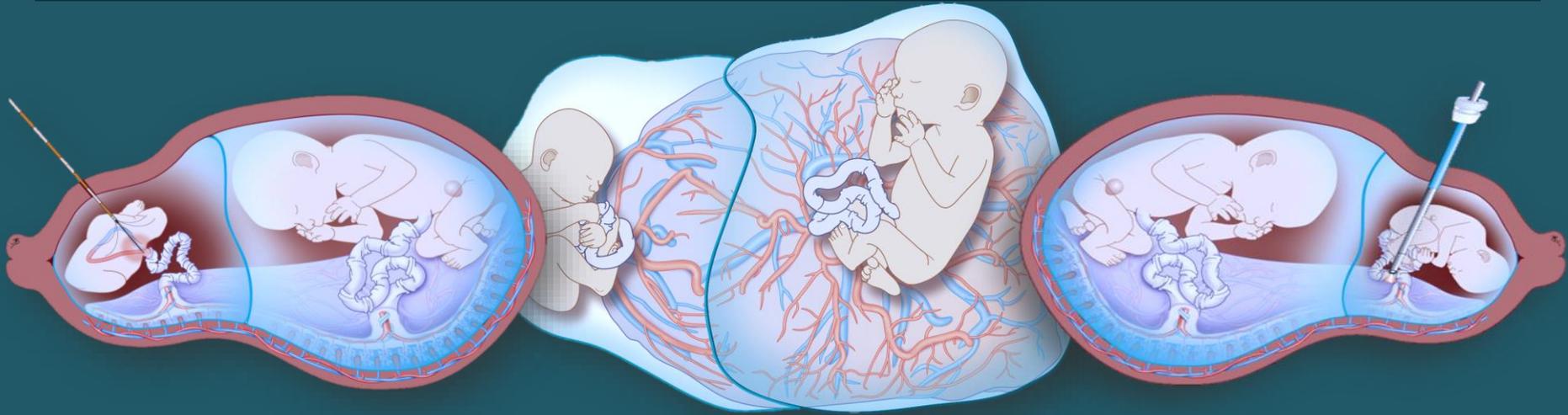
20% birth before 32 weeks

10% 24-28 weeks

10% 28-32 weeks



2nd – 3rd trimester: Selective reduction in MC twins



RISK co-twin

TRAP, lethal monosomy X, Ebstein's, TTTS

IMMEDIATE REDUCTION From 16 weeks onward

80% survival

10% fetal demise
10% miscarriage

20% birth before 32 weeks

10% 24-28 weeks
10% 28-32 weeks



NO RISK co-twin and NONLETHAL

hydrocephaly, spina bifida, Dandy walker

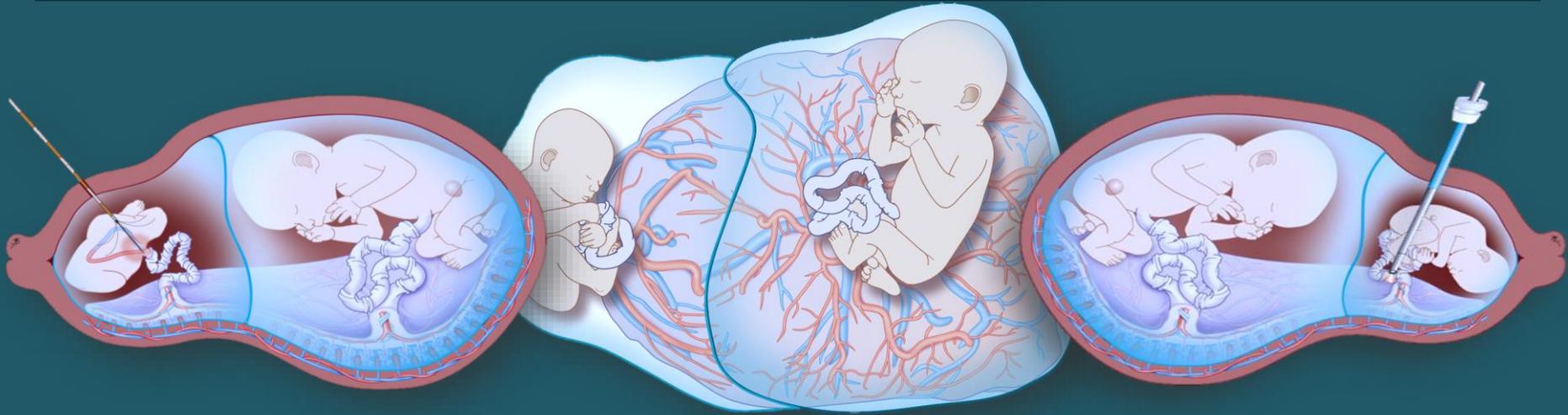
LATE REDUCTION Between 28-30 weeks

No procedure-related loss

20% birth before 32 weeks

0% 24-28 weeks
20% 28-32 weeks

2nd – 3rd trimester: Selective reduction in MC twins



RISK co-twin

TRAP, lethal monosomy X, Ebstein's, TTTS

IMMEDIATE REDUCTION From 16 weeks onward

80% survival

10% fetal demise
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20% birth before 32 weeks

10% 24-28 weeks
10% 28-32 weeks

NO RISK co-twin and NONLETHAL

hydrocephaly, spina bifida, Dandy walker

LATE REDUCTION Between 28-30 weeks

No procedure-related loss

20% birth before 32 weeks

0% 24-28 weeks
20% 28-32 weeks

NO RISK co-twin and LETHAL in early neonatal period

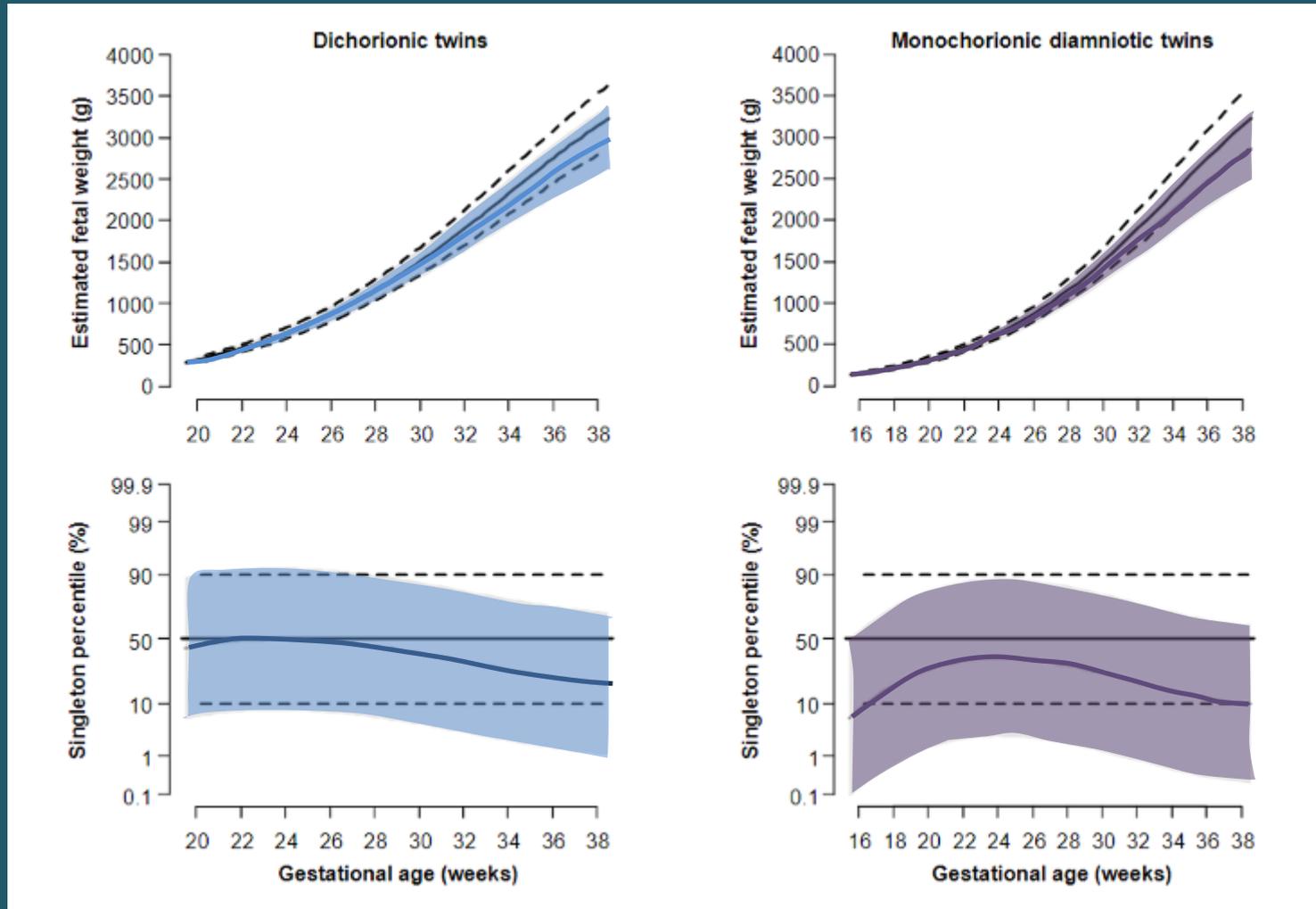
HLHS, bilateral renal agenesis

COMFORT CARE After birth

No procedure-related loss

No iatrogenic preterm birth

2nd – 3rd trimester: Poor growth in twins



2nd – 3rd trimester: sIUGR in DC twins

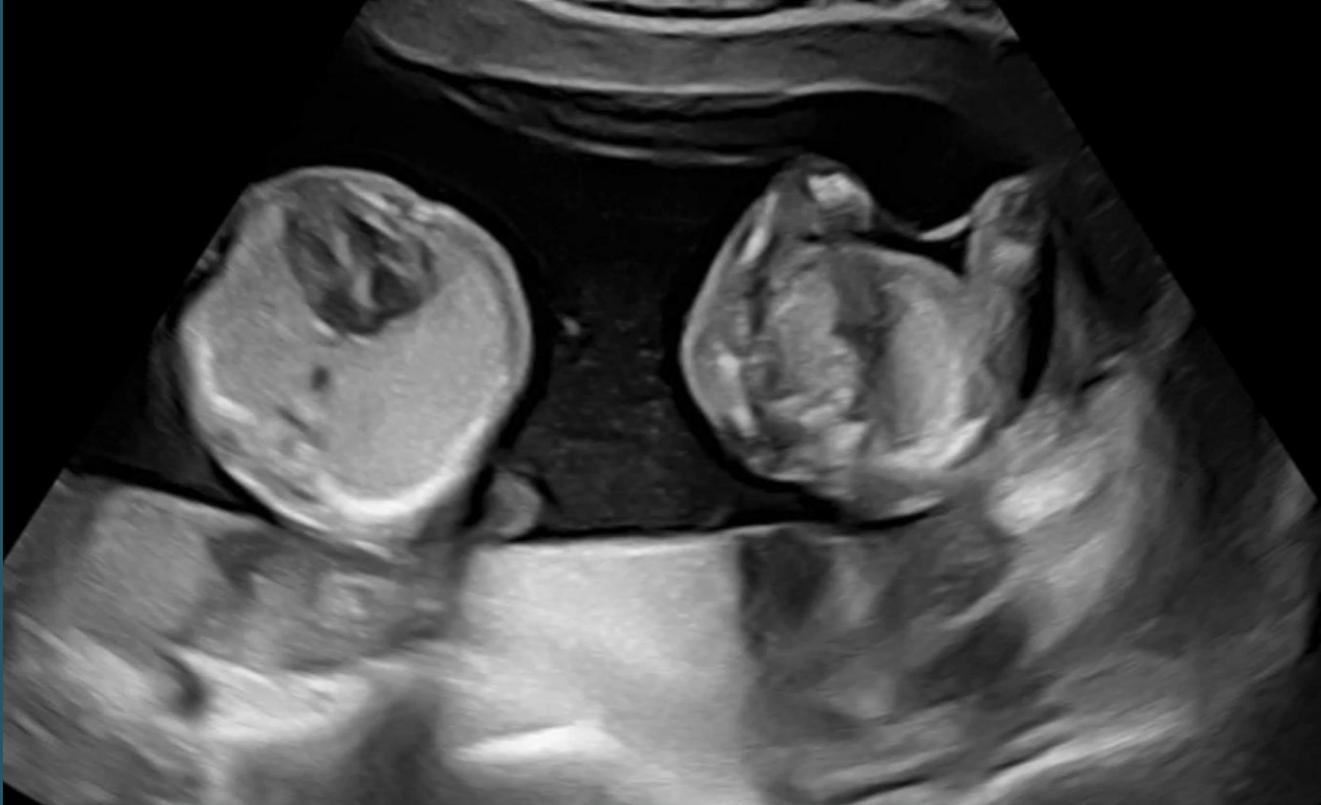


2nd – 3rd trimester: sIUGR in DC twins

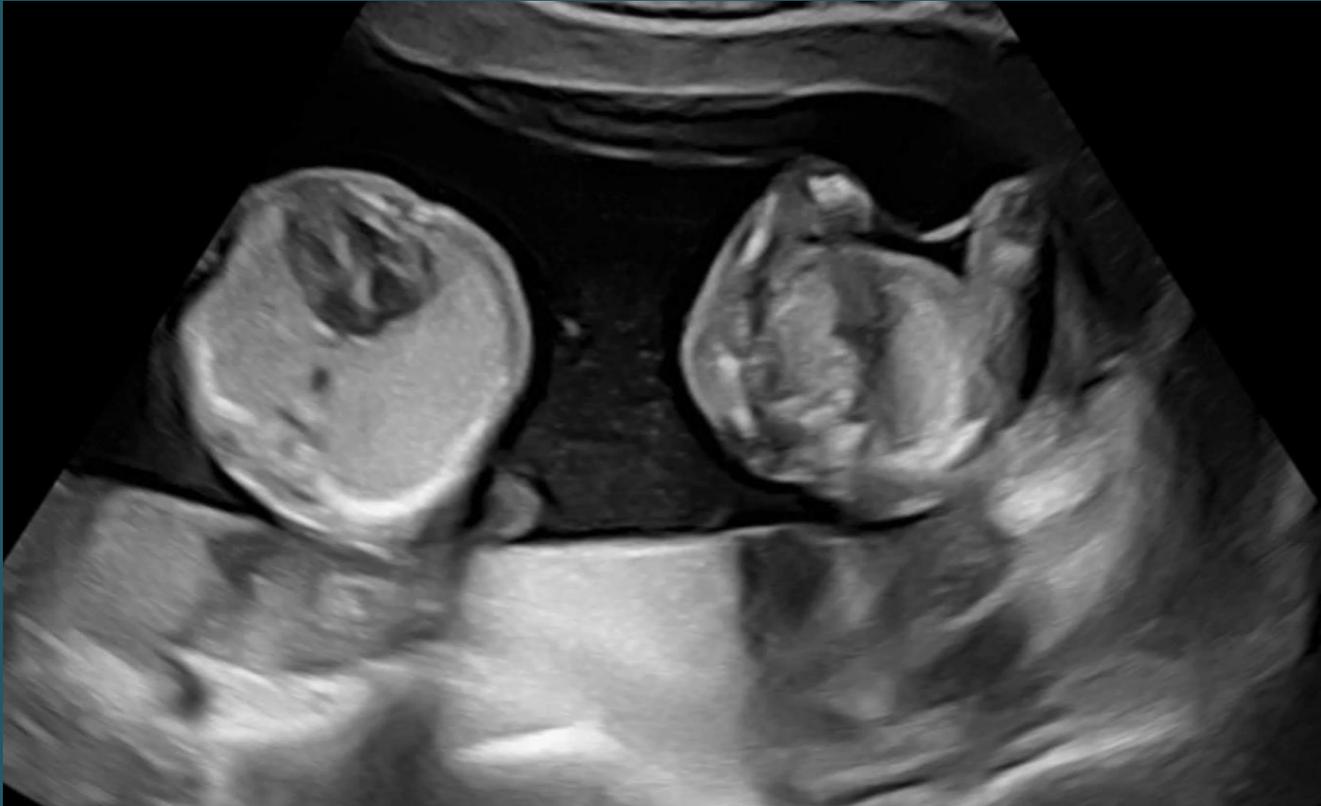


- Aneuploidy or infection may explain poor growth of 1 twin
- Fetal demise of the small twin should not directly affect the large twin
- Timing of birth will depend on the chances of survival of both twins

2nd – 3rd trimester: sIUGR in MC twins



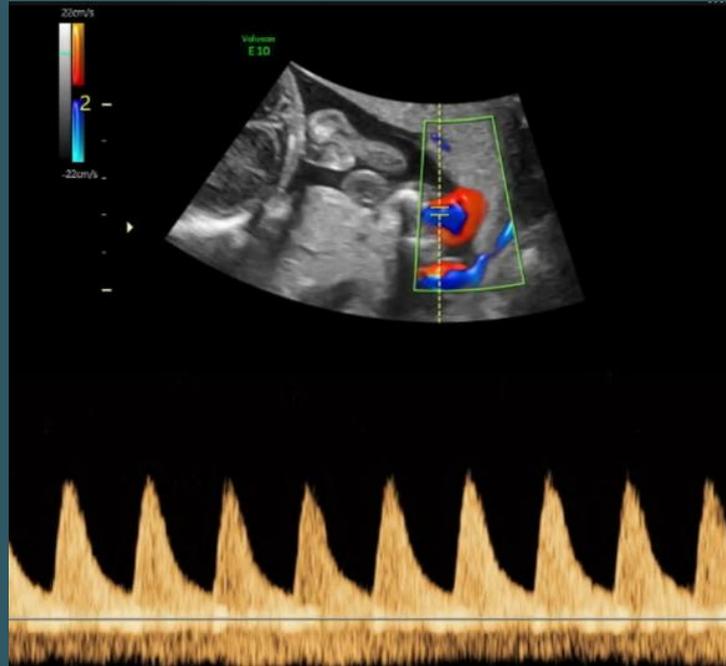
2nd – 3rd trimester: sIUGR in MC twins



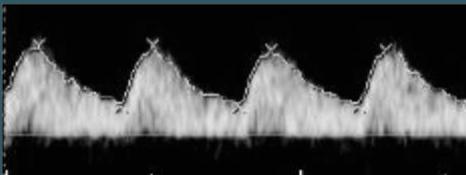
- Aneuploidy or infection is unusual to explain poor growth of 1 twin
- Fetal demise of the small twin will directly affect the large twin
- Timing of birth will depend on the risk of demise of the smaller twin

2nd – 3rd trimester: sIUGR in MC twins

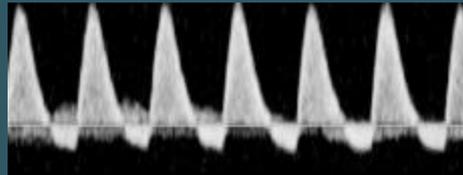
20% ≠ in size



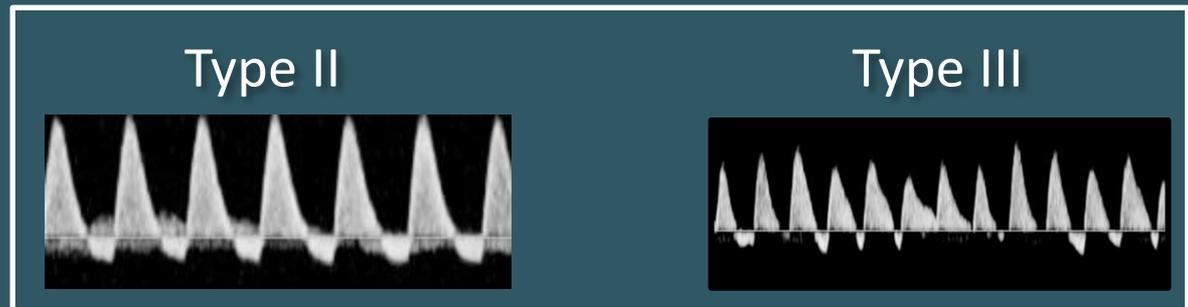
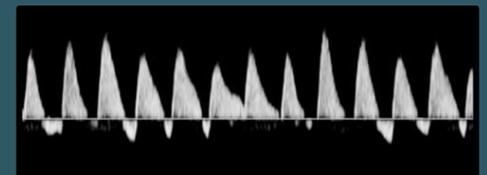
Type I



Type II



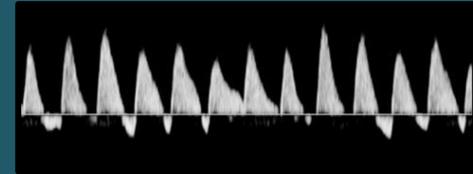
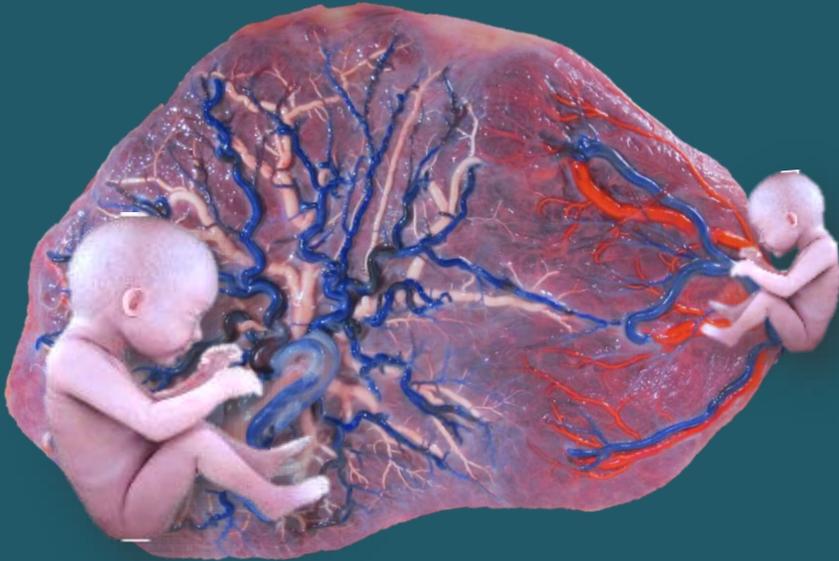
Type III



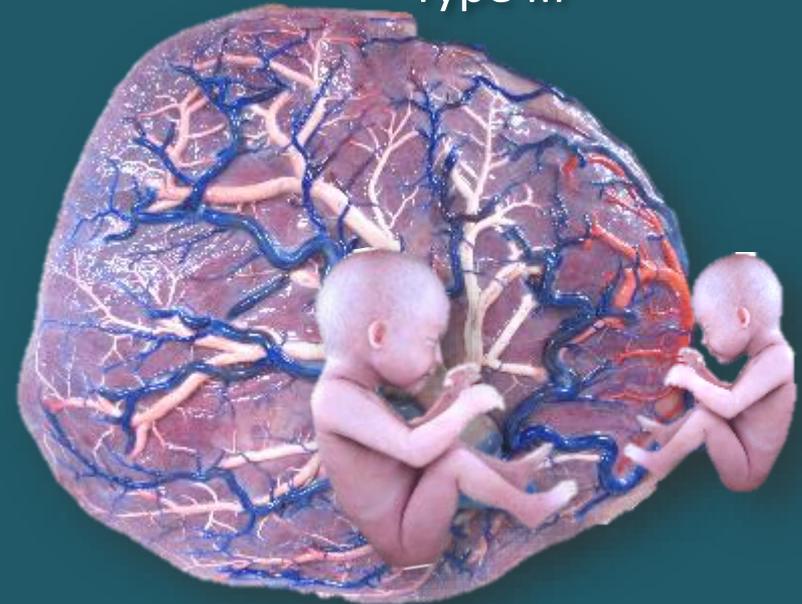
2nd – 3rd trimester: sIUGR in MC twins



Type II



Type III



2nd – 3rd trimester: management of sIUGR in MC twins



Expectant

Type II
60% survival

Type III
85% survival

2nd – 3rd trimester: management of sIUGR in MC twins

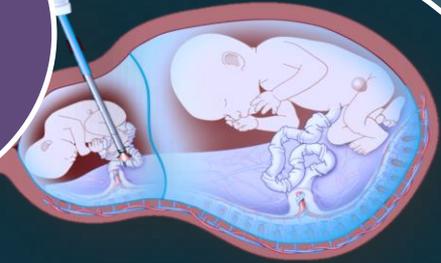


Expectant

Type II
60% survival

Type III
85% survival

40% survival



Selective reduction

- Discordant anomalies
- Worsening type II - III
- Parental request

2nd – 3rd trimester: management of sIUGR in MC twins

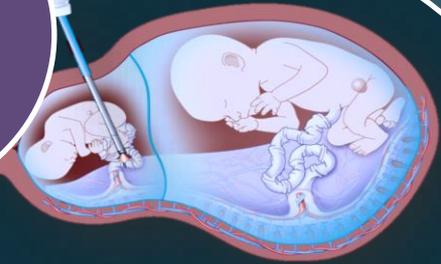


Expectant

Type II
60% survival

Type III
85% survival

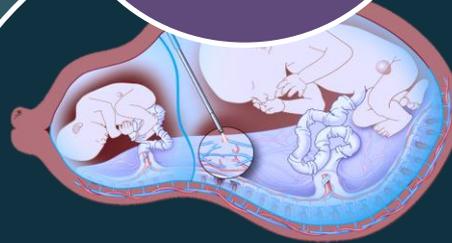
40% survival



Selective reduction

- Discordant anomalies
- Worsening type II - III
- Parental request

25-60% survival



Fetoscopic laser surgery

- To protect the larger twin

2nd – 3rd trimester: management of sIUGR in MC twins

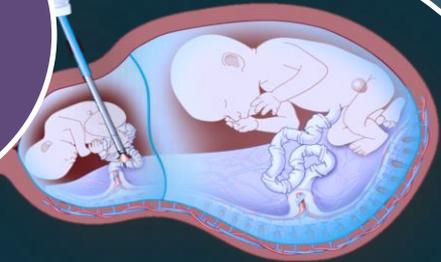


Expectant

Type II
60% survival

Type III
85% survival

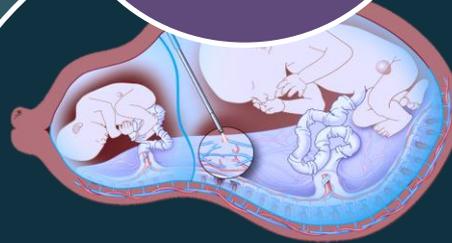
40% survival



Selective reduction

- Discordant anomalies
- Worsening type II - III
- Parental request

25-60% survival



Fetoscopic laser surgery

- To protect the larger twin

2nd – 3rd trimester: cervical length management

10% delivers before 32 weeks



2nd – 3rd trimester: cervical length management

10% delivers before 32 weeks



CL \leq 25 mm
TWIN- cerclage trial



CL \leq 15 mm or dilated cervix

Take home message



DICHORIONIC

11-14 wks

- Chorionicity
- Dating
- Labeling
- Early anatomy
- Screening T21

MONOCHORIONIC

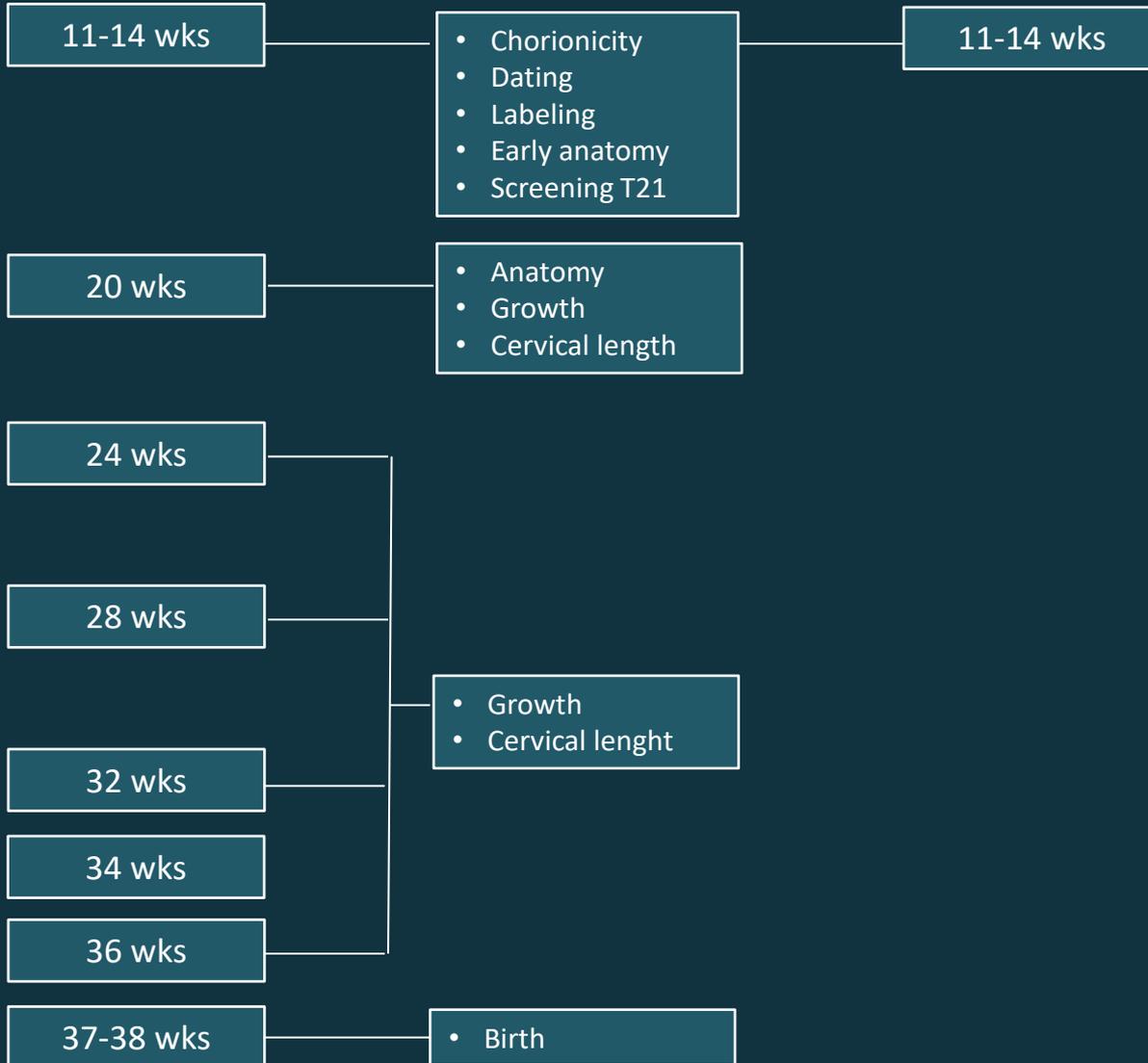
11-14 wks

Take home message



DICHORIONIC

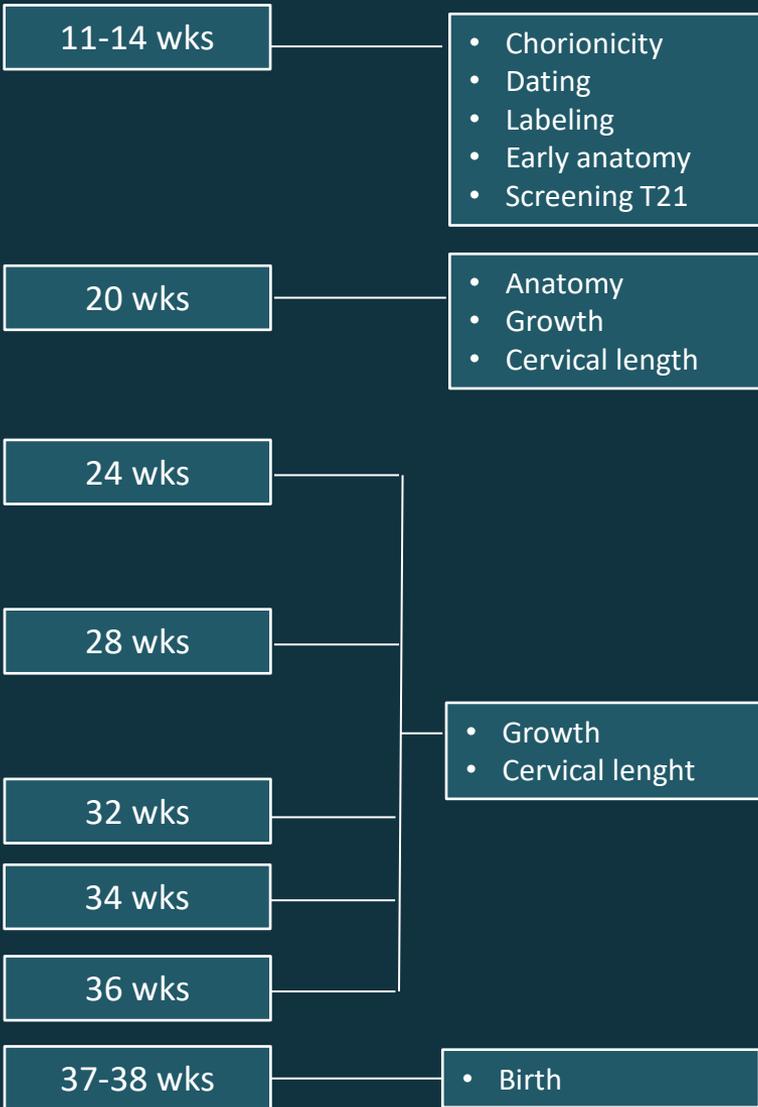
MONOCHORIONIC



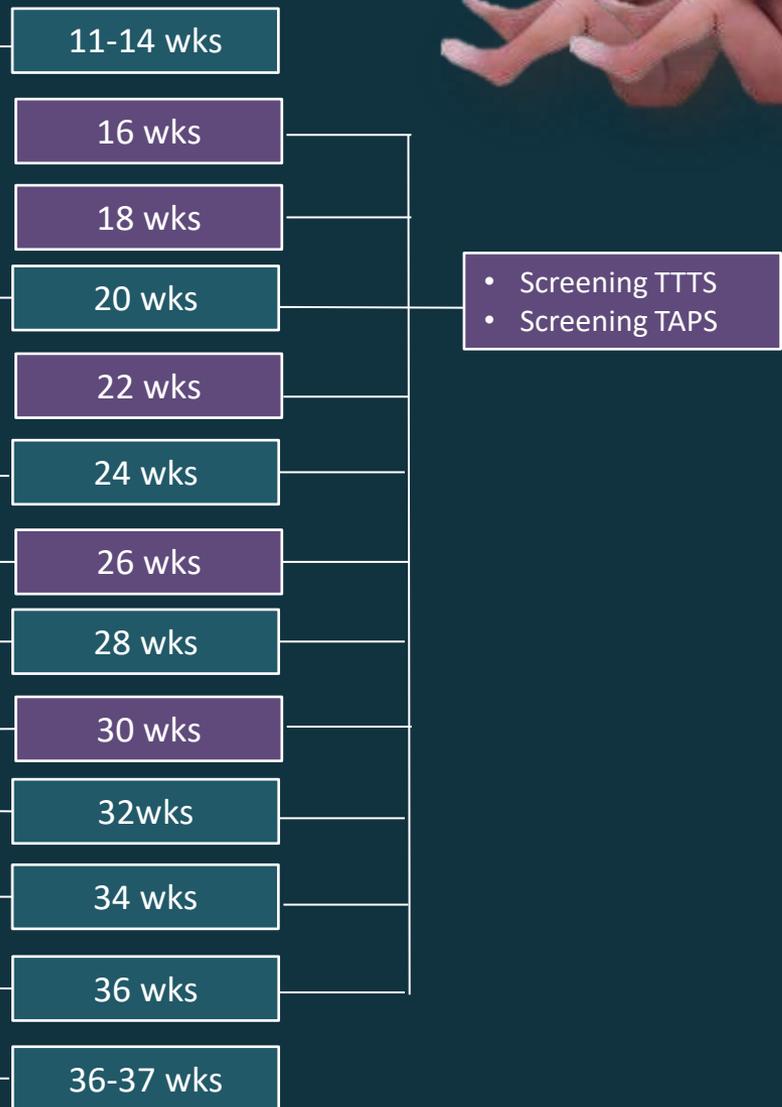
Take home message



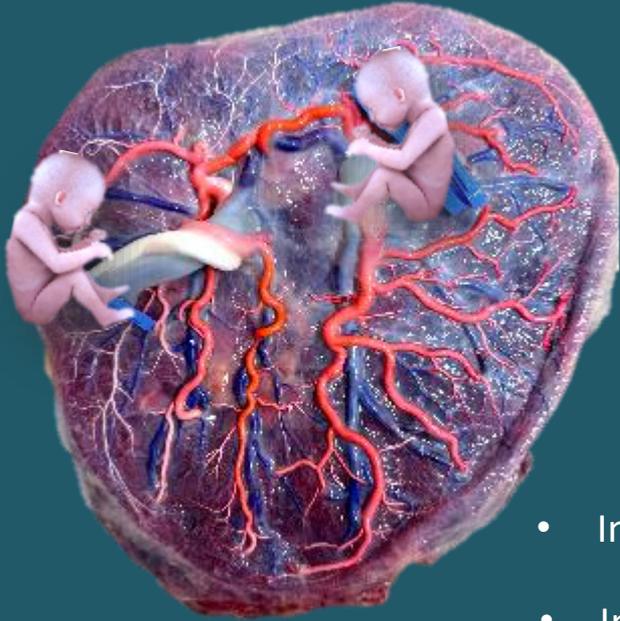
DICHORIONIC



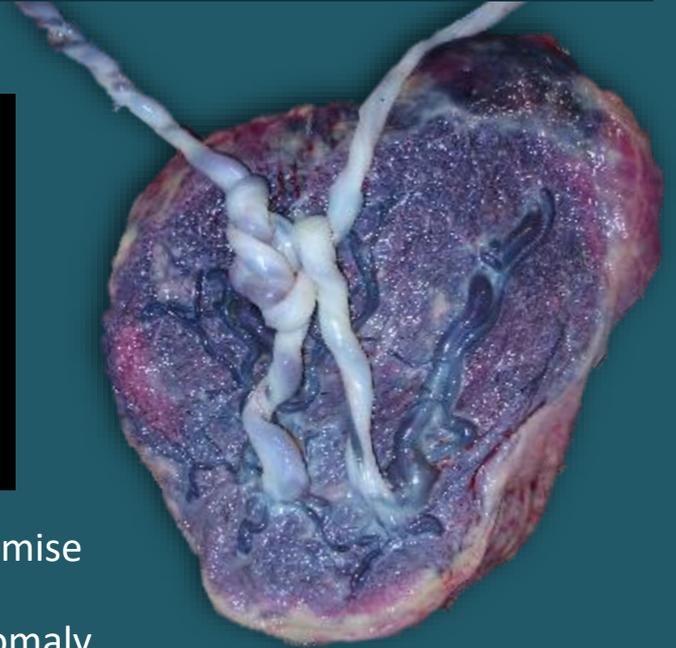
MONOCHORIONIC



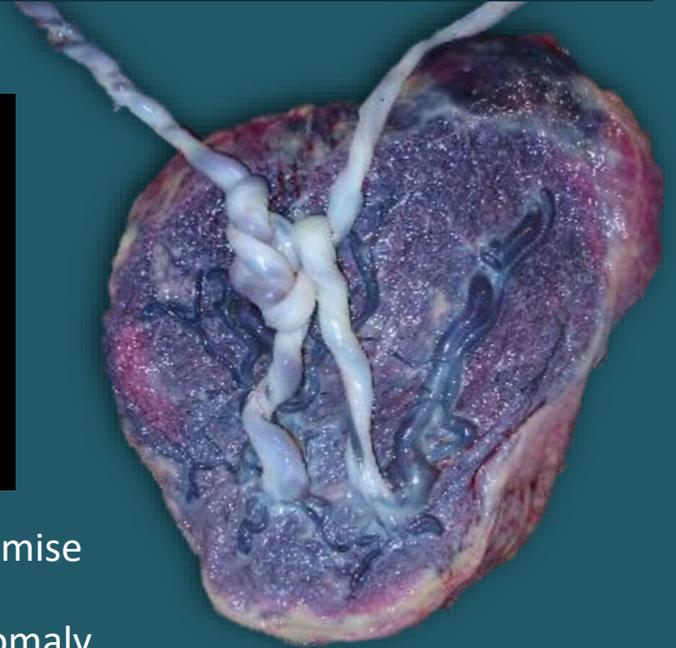
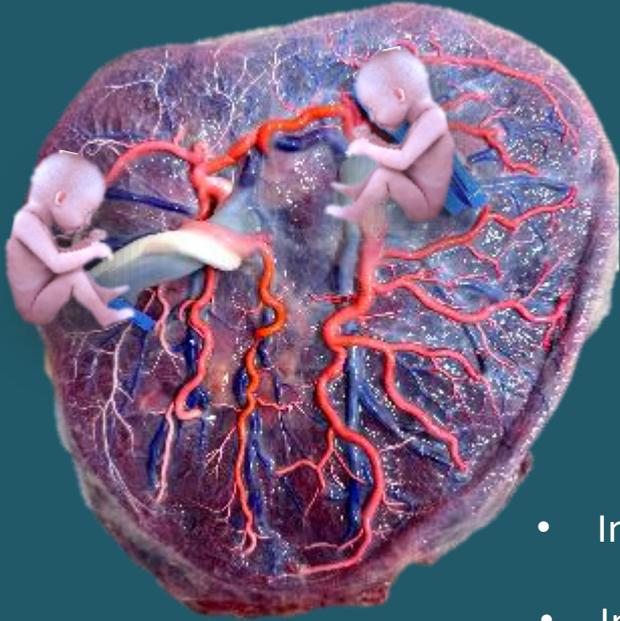
MCMA TWINS



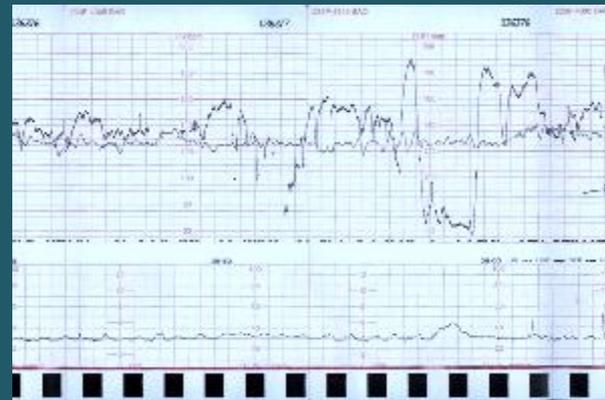
- Increased risk of sudden fetal demise
- Increased risk of a structural anomaly



MCMA TWINS



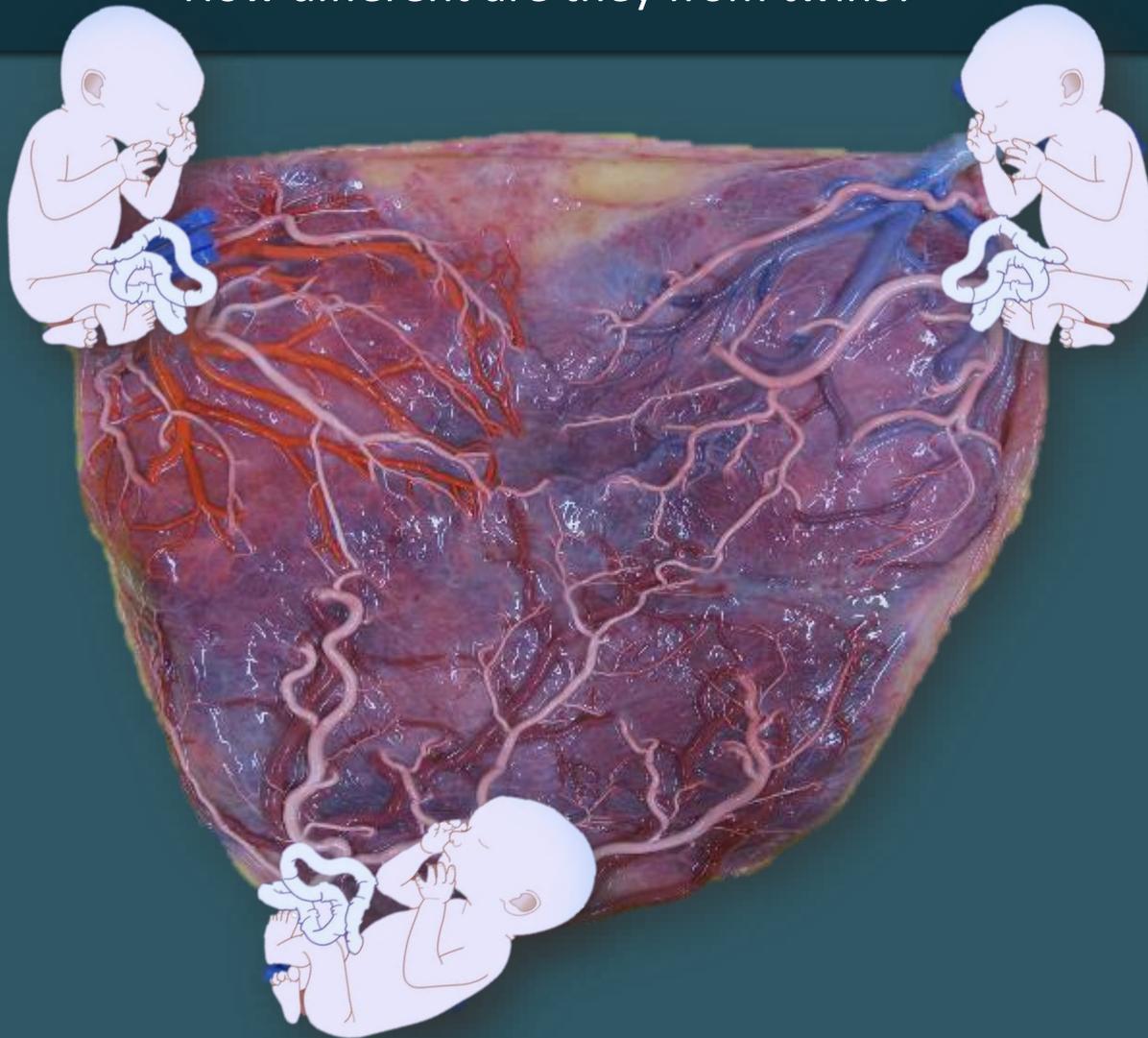
- Increased risk of sudden fetal demise
- Increased risk of a structural anomaly



Survival in MCMA is 78% *versus* 88% in MCDA twins

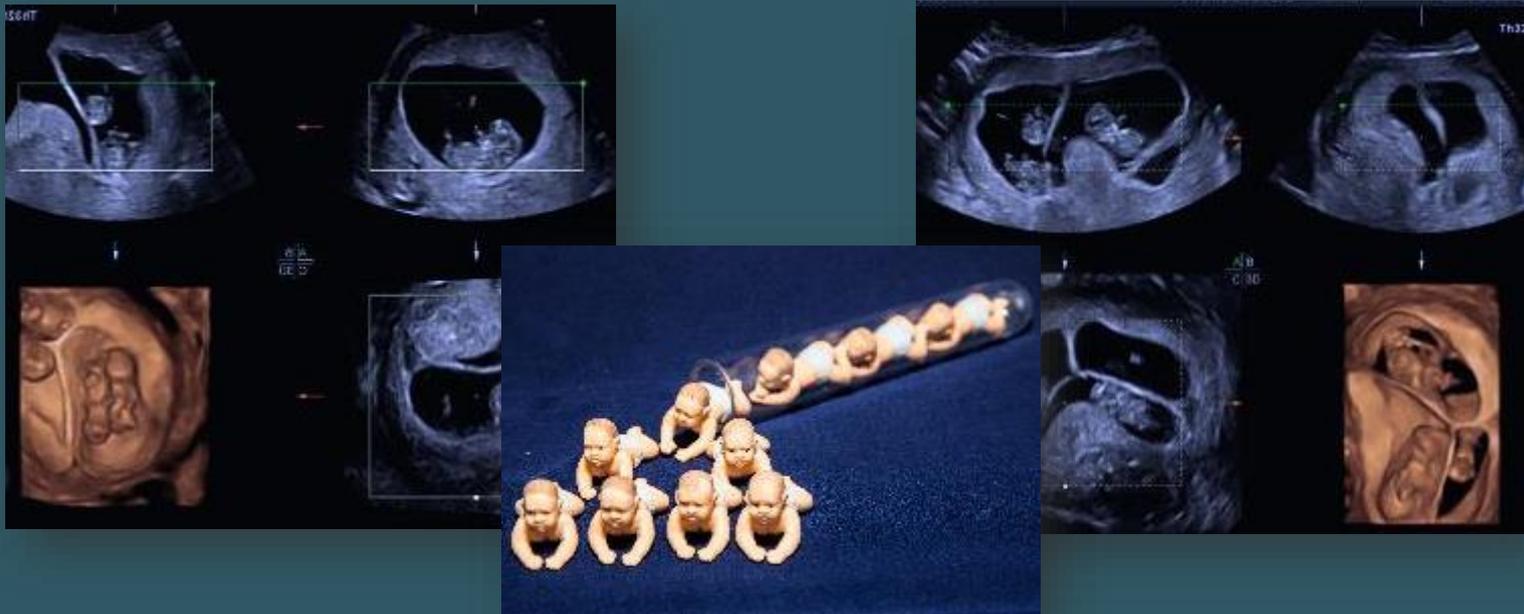
Triplet Pregnancies

How different are they from twins?



Epidemiology of multiple pregnancies

Natural incidence	twins	1 in 89	pregnancies
	triplets	1 in 89 ² or ≈8000	pregnancies
	quadruplets	1 in 89 ³ or ≈700000	pregnancies
<i>Hellin's law</i>	N-tuplets	1 in 89 ⁿ⁻¹	pregnancies



USA 2021 triplets 1 in 1315 pregnancies

Flanders 2023 triplets 1 in 4307 pregnancies

Typical features of triplet pregnancies from a fetal perspective



Singleton: 39-40 weeks

Increased risk of preterm birth



Twin: 35-36 weeks



Triplet: 33-34 weeks

Increased risk of poor growth and growth discordance, esp. if nulliparous or DC-MC

Largest - Smallest
—————
Largest



30% ≥ 20% ≠
10% ≥ 40% ≠

From 28 weeks, growth velocity starts to decline so that by 35 weeks, triplets are on average growth restricted

3 babies at risk of ELBW (<1000g), neonatal mortality/morbidity and long-term handicap

- ✓ Perinatal mortality= 7% in triplets *versus* 3% in twins *versus* 0.7% in singletons
- ✓ Risk of cerebral palsy is 1 in 20 in triplets; 1 in 100 in twins and 1 in 500 in singletons

Typical features of triplet pregnancies from the mother's perspective

1 in 25 risk of major obstetric hemorrhage or other acute life-threatening morbidity

Need for caesarean section to be offered from 35.0 weeks onward if not yet delivered

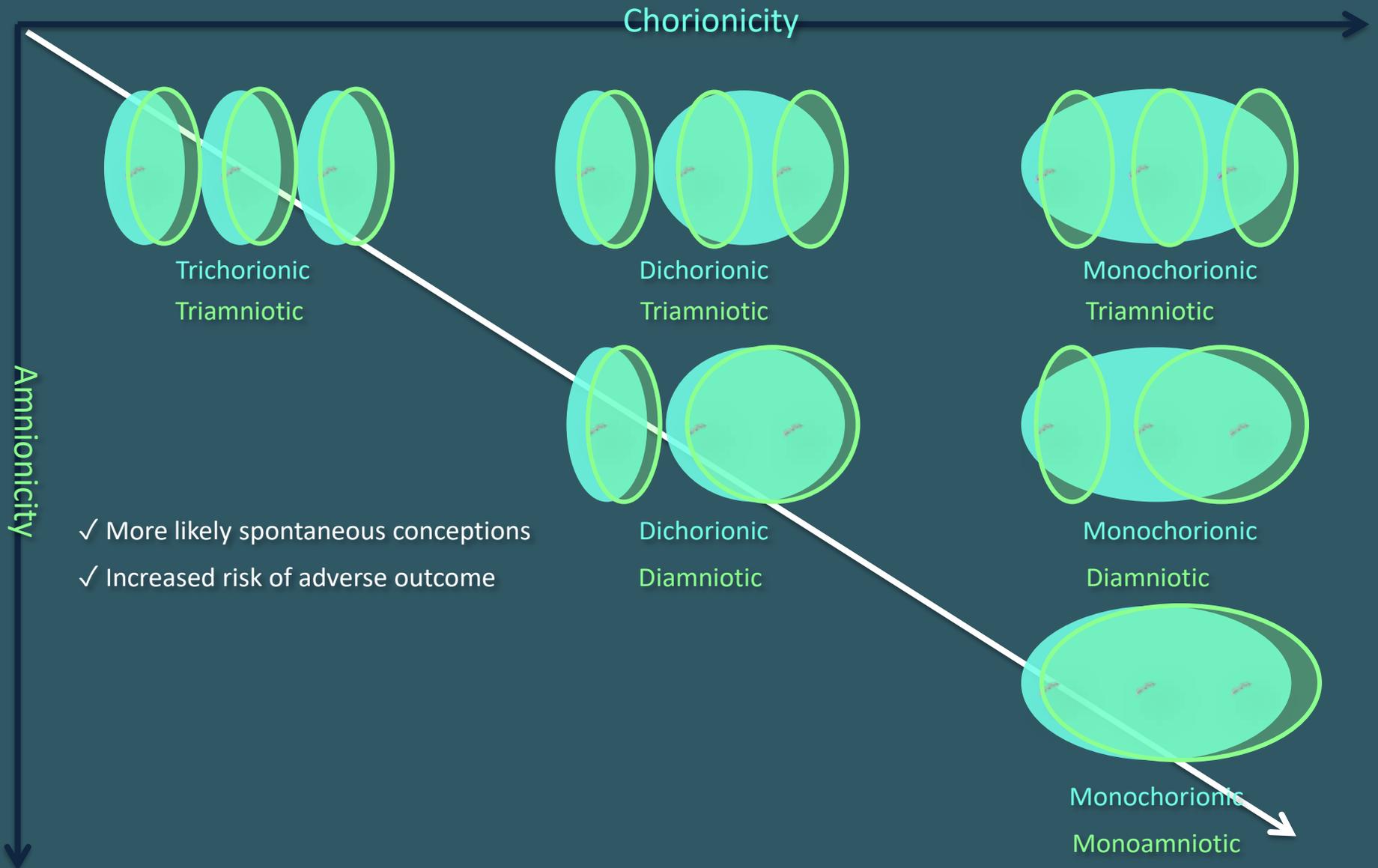


Increased risks of pre-eclampsia, diabetes, cholestasis, anemia

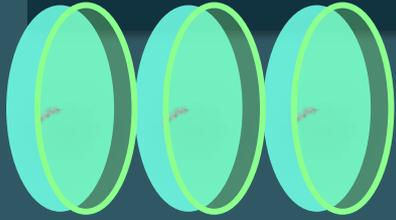
Physical discomfort, especially from 24 weeks onwards

Financial and emotional burden to look after 3 baby's: depression – marital strain

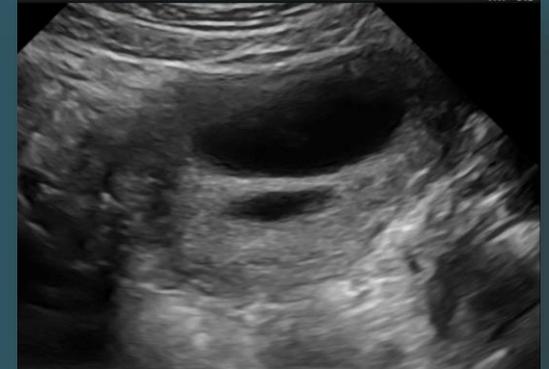
Different type of triplet pregnancies



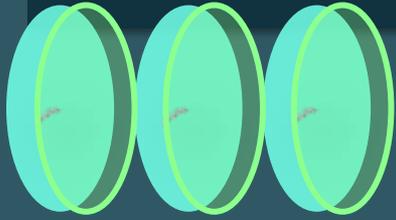
Outcome and management of trichorionic triplets



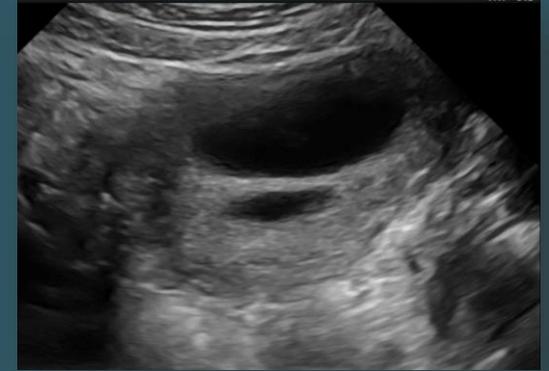
	TCTA N=411
% ART	90-95%
Survival	90%
GA at birth	34 wks
Birth < 32 wks	30%
Birth < 28 wks	10%



Outcome and management of trichorionic triplets



	TCTA N=411
% ART	90-95%
Survival	90%
GA at birth	34 wks
Birth < 32 wks	30%
Birth < 28 wks	10%



✓ 1st trimester screening for aneuploidy usually done by maternal age and US variables

NT measurement and other markers for aneuploidy – allocate sufficient time!

✓ 1st trimester discussion of pro's and con's of fetal reduction, usually from 3 to 2

Birth < 32 wks from 30% to 10% with extra miscarriage of about 5%



Take home messages!

✓ Most TCTA triplets result from ART

✓ Triplets are born at 33-34 wks with perinatal death in 7% and cerebral palsy in 5%

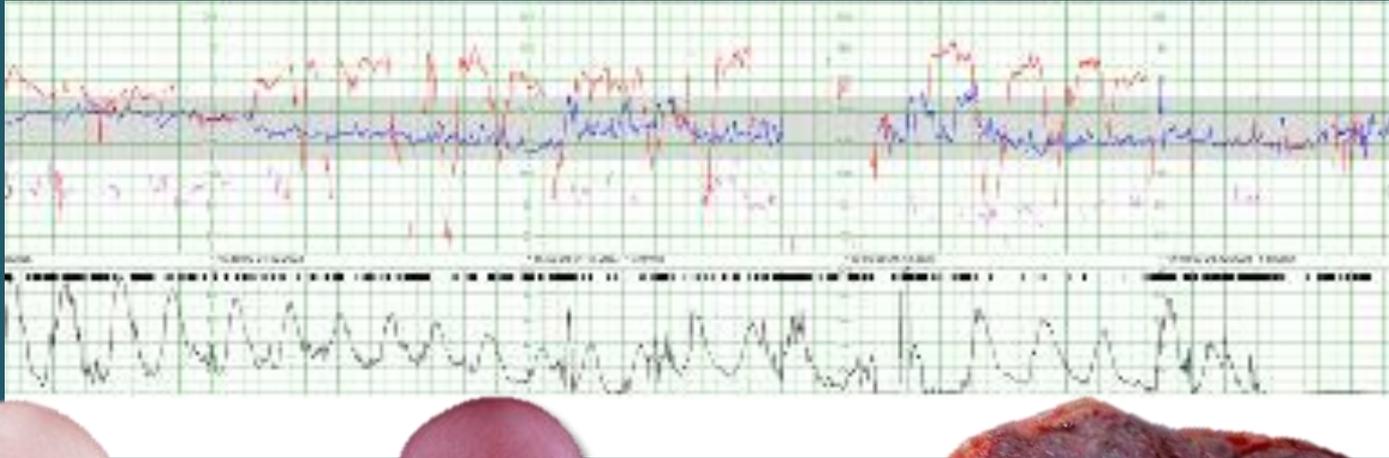
✓ Mothers have a 4% risk of life-threatening morbidity

✓ The more the triplets share - the higher the risks

✓ What is the long-term outcome? [East Flanders Prospective Twin Survey on Triplets](#)

What is your diagnosis? Case 1

- Induction @36w1d



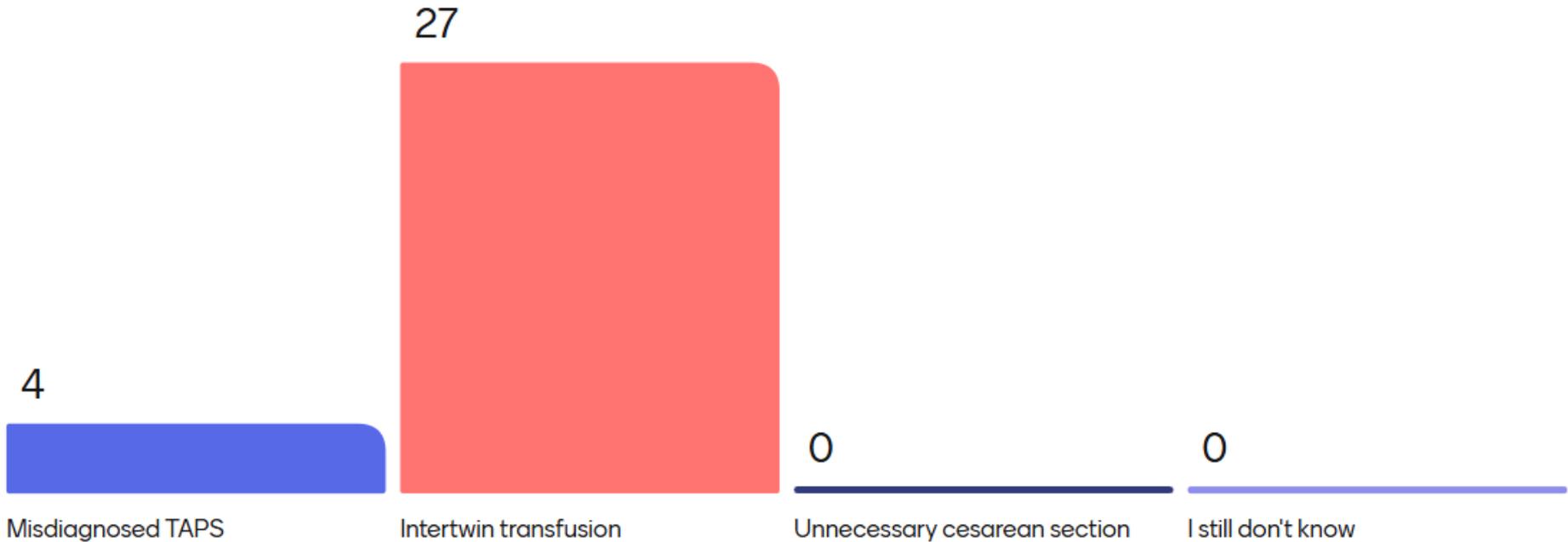
- pH 7.18
- Hb 12 gr/dL
- pH 7.22
- Hb 21 gr/dL

What is your diagnosis now? Case 1



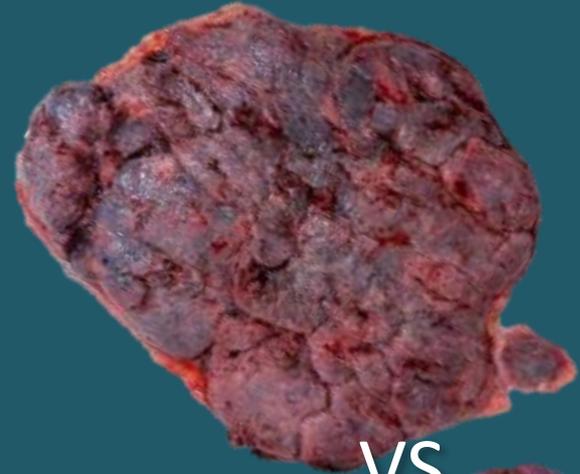
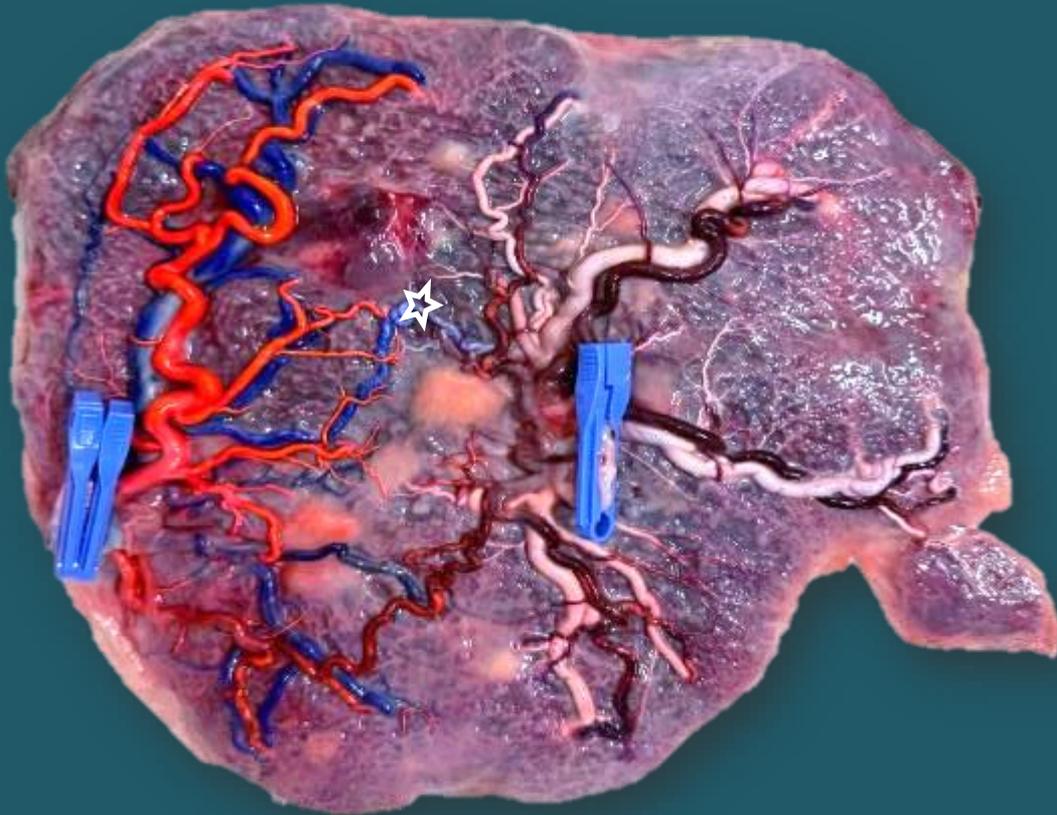
What is your diagnosis now? Case 1

What is your diagnosis now?



What is your diagnosis? Case 1

Acute intertwin transfusion



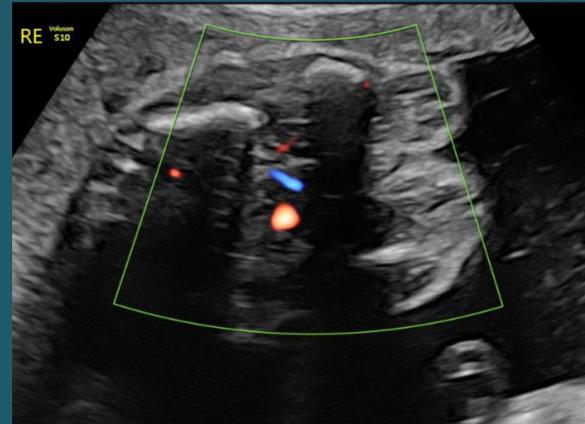
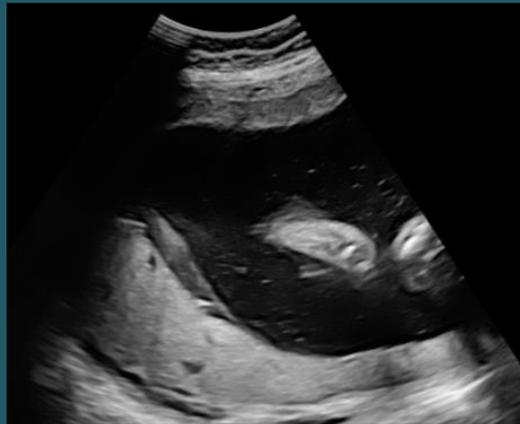
VS



2x low reticulocyte count

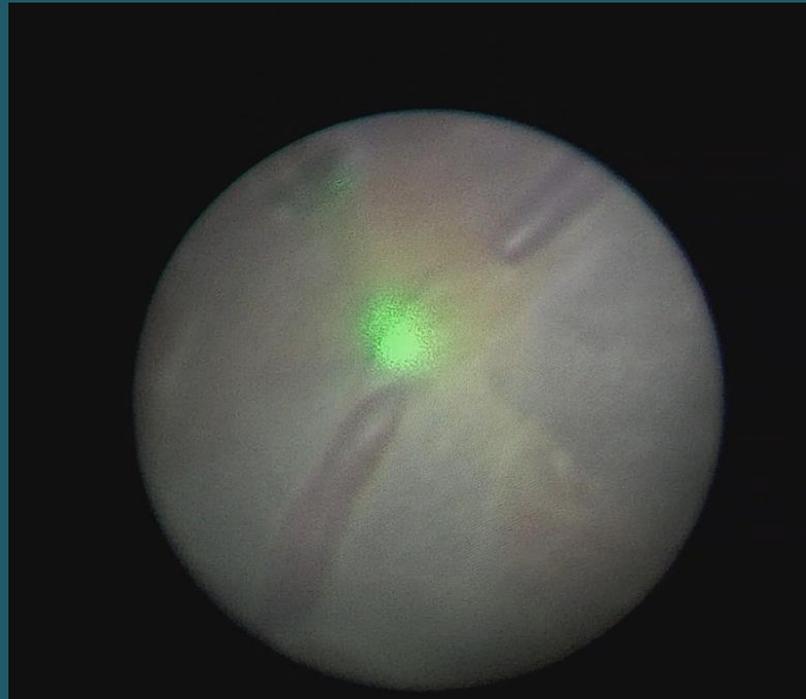
What would you do? Case 2

- G2P1, spontaneous MCDA twins
- Concordant growth and amniotic fluid
- @25w2d: TTTS st 2



What would you do? Case2

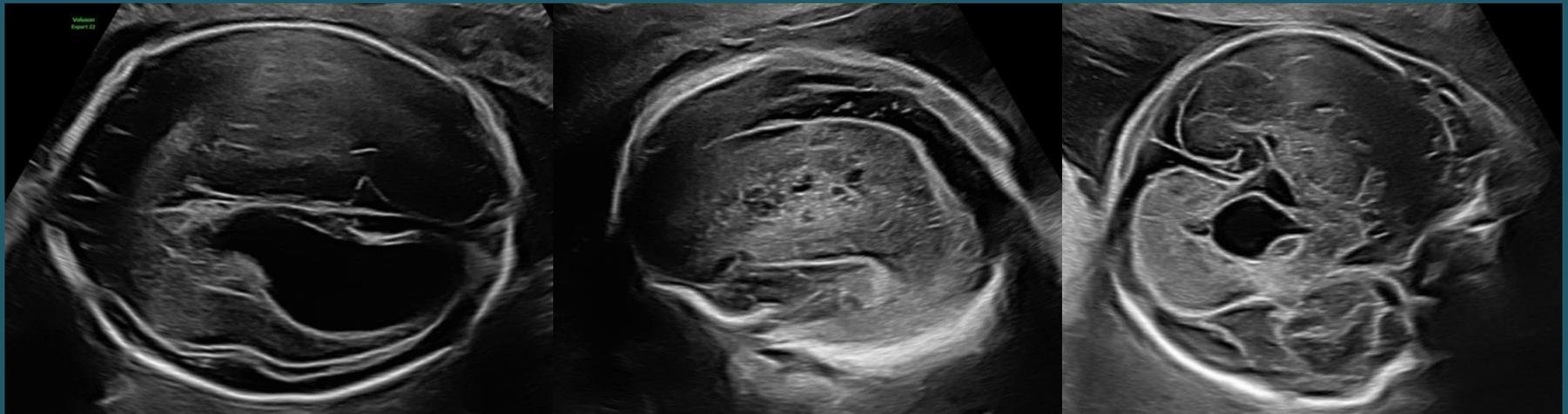
- fMRI: no signs of bleeding or ischemia
- Parents opt for laser therapy
- @25w3d:



What would you do? Case 2

US @27w5d

- Fetus 1 (ex-donor): normal amniotic fluid and dopplers
- Fetus 2 (ex-recipient): severe ventriculomegaly right with atrophy of the cortex, periventricular cysts, abnormal corpus callosum and hyperechogenic brain



- Broad CMS and septostomy

What would you do? Case 2

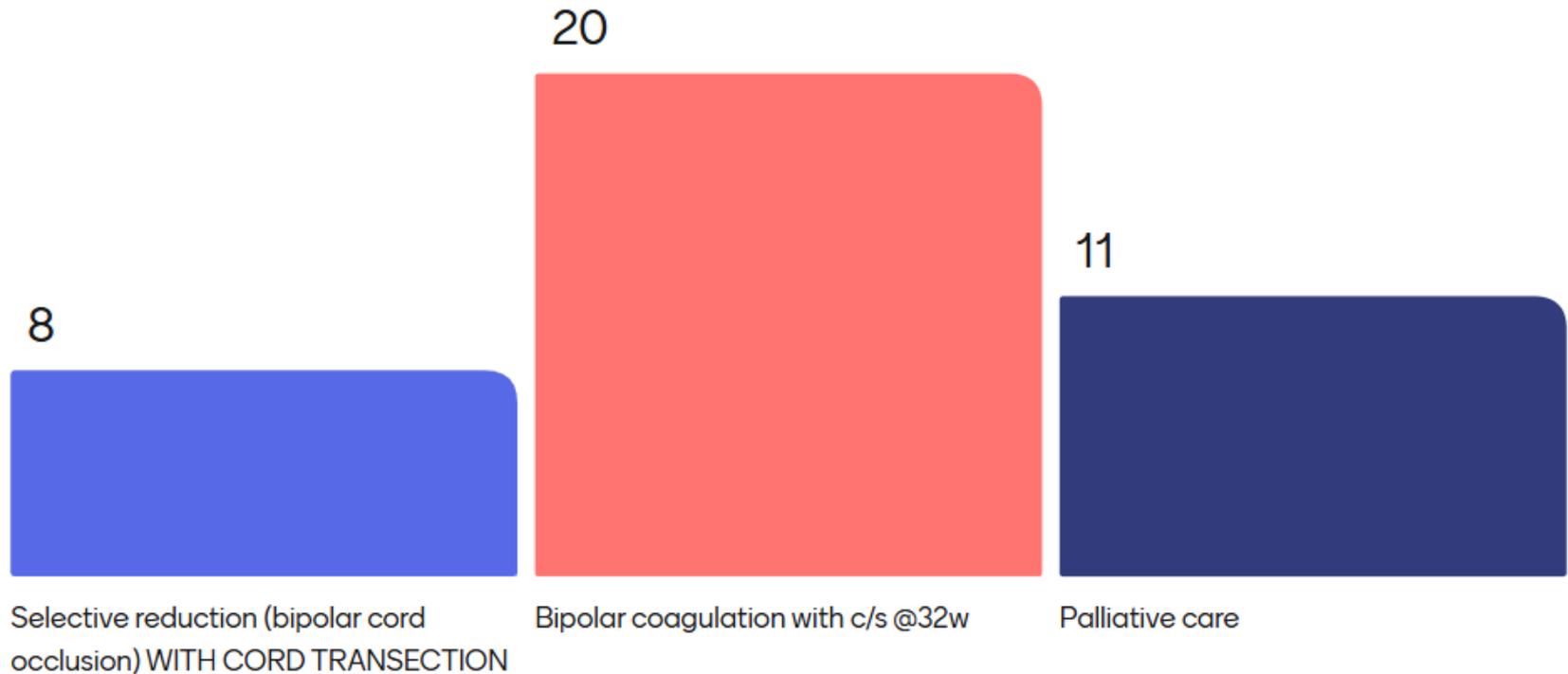
- MRI @27w5d:
Fetus 2: Broad parenchyma loss, compatible with ischemia,
Severe ventriculomegaly right
- The parents ask for sTOP
 - Selective reduction (bipolar cord occlusion) WITH CORD TRANSECTION
 - Bipolar coagulation followed by c/s @32w
 - Palliative care

What would you do? Case 2



What would you do? Case 2

What would you do? Case 2



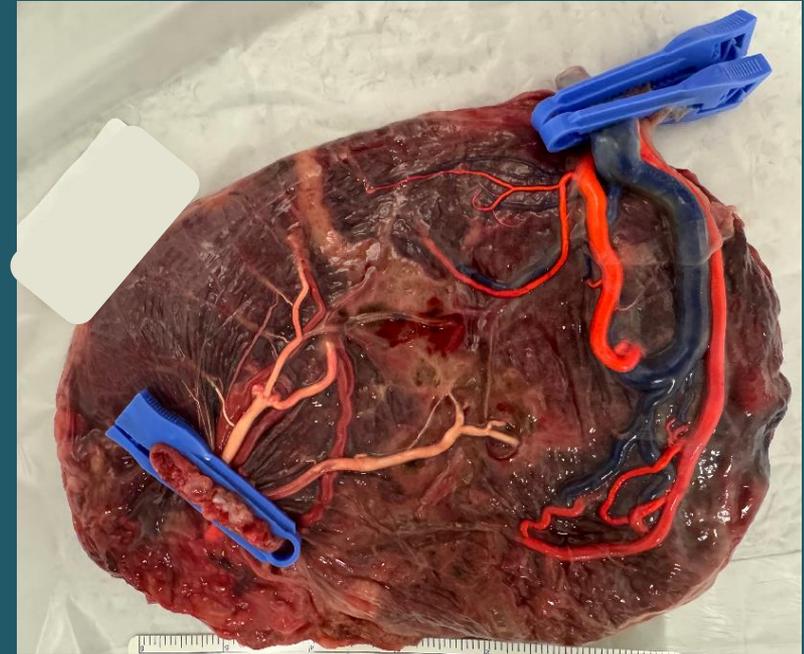
What would you do? Case 2

- Hospitalization @29w0d due to PPROM
- Management: section + reduction planned @32w, palliative care for fetus 2 if earlier labor

- @30w0d: C/S for abnormal CTG fetus 1
 - Baby 1: 1100 gr, good start, CPAP
 - Baby 2: 1370 gr, good start, palliative care
 - Removal of amniotic bands during extraction

What would you do? Case 2

- Baby 1:
Transfer to peripheric hospital @20d
High flow 4L
Vena porta thrombosis
Full enteral feeding @14d
- Baby 2
Good respiratory adaptation after birth
Deceased @d3





Liesbeth Lewi



KU LEUVEN



Roland Devlieger



Hannes Vandermerwe



Jan Deprest

Thank you!