Prenatal management of Lower Urinary Tract Obstruction





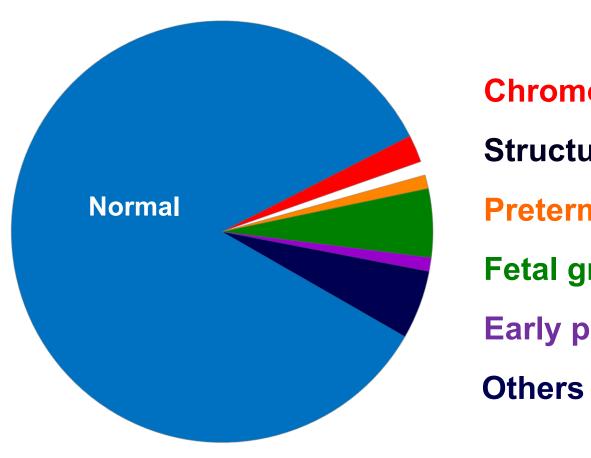
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Pregnancy outcome

Average European birth rate: ~ 5 million per year



Chromosomal abnormalities	1-2%
Structural defects	1%
Preterm pre-eclampsia	0.5%
Fetal growth restriction	5%
Early preterm delivery	1%
Others	5%

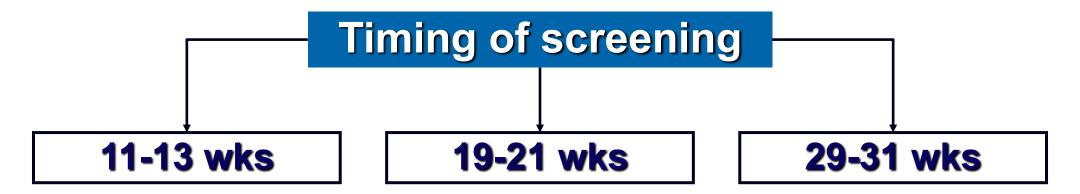


Prenatal Ultrasound











Early detection of fetal defects

Ultrasound at 11-13 wks (n=44,859)

Major defects 1%

Detected 44%

Always detected 30%

Acrania
Holoprosencephaly
Exomphalos
Gastroschisis
Megacystis
Body stalk anomaly















Early detection of fetal defects

Undetectable at 11-13w

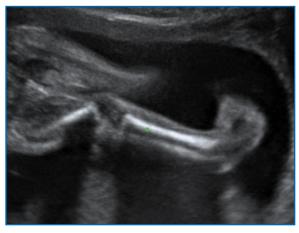
Corpus callosum agenesis
CCAM /sequestration
Esophageal / duodenal atresia
Bowel obstruction
Hydronephrosis
Dysplastic kidneys
Talipes











Syngelaki et al 2011



Fetal urinary tract on ultrasound

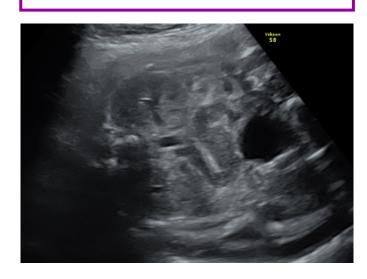
1st Trimester



2nd Trimester



3rd Trimester



Criteria for normality

Normal echogenicity

No hydronephrosis

Visible bladder, not enlarged

Ureters not visible

Normal amniotic fluid after 15 weeks



Early detection of fetal defects

Megacystis





- Prevalence: 1/1,600
- Bladder length

7-15 mm resolution 90%

>15 mm resolution 0%

•Trisomy 13 or 18 (31%)

35 of 57,119 singleton pregnancies

Kagan *et al.*, 2010



Invasive diagnostic testing

CVS



Amnio



Fetal blood

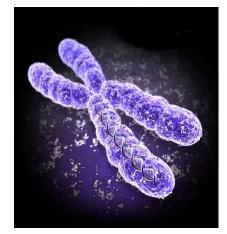


Fetal karyotype

Microarray

Targeted genetic testing







Lower Urinary Tract Obstruction

Prevalence 1 in 5,000 pregnancies

- Chromosomal defects
- Genetic syndromes
- Other structural defects

Isolated LUTO

Posterior urethral valves Urethral stenosis/atresia

High neonatal mortality

Pulmonary hypoplasia End-stage renal disease

Ultrasound findings

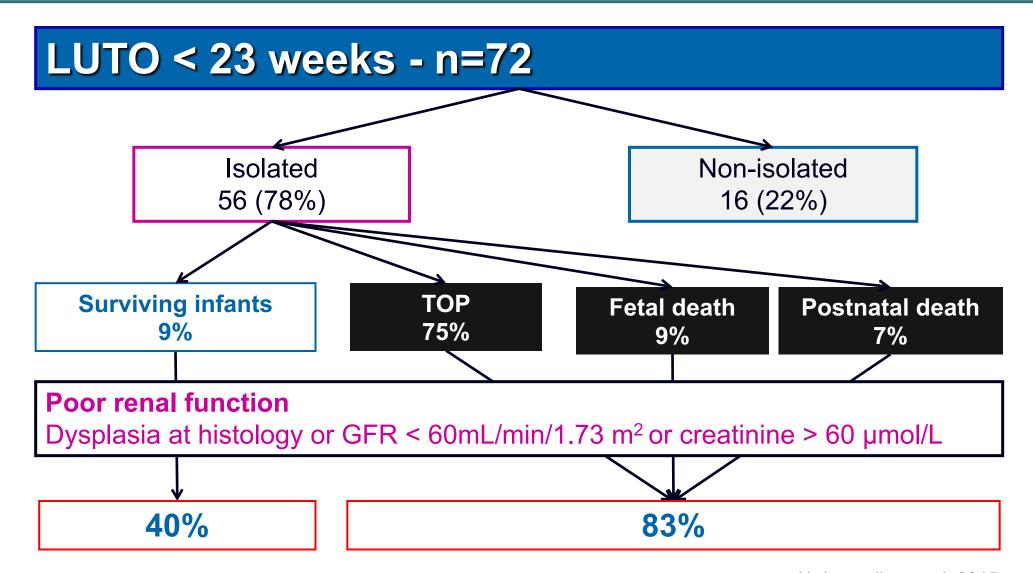
Distended bladder with 'keyhole sign' Oligo-anhydramnios Hydro-ureteronephrosis Ecogenic kidneys







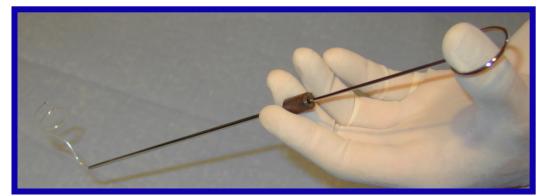
Natural history





Vesicoamniotic shunting

Procedure – 16-28 weeks

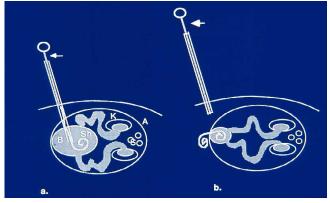






- Local anesthesia
- 3 mm KCH catheter or other devices
- Antibiotic prophylaxis
- Tocolisis (Indomethacine, Nitrate)





Nicolini et al, 1987



Vesicoamniotic shunting

Perinatal survival

Shunt n=112

Expectant n=134

OR

95%CI

57%

39%

2.54

1.14-5.67

12 months survival

Shunt n=43

44%

Expectant n=91

42%

OR

1.77

95%CI

0.25-12.71

Good renal function

Shunt n=41

68%

Expectant

n=64

48%

OR

2.09

95%CI

| 0.74-5.94

Shunt complications

Blockage or migration Preterm delivery

25%

20%



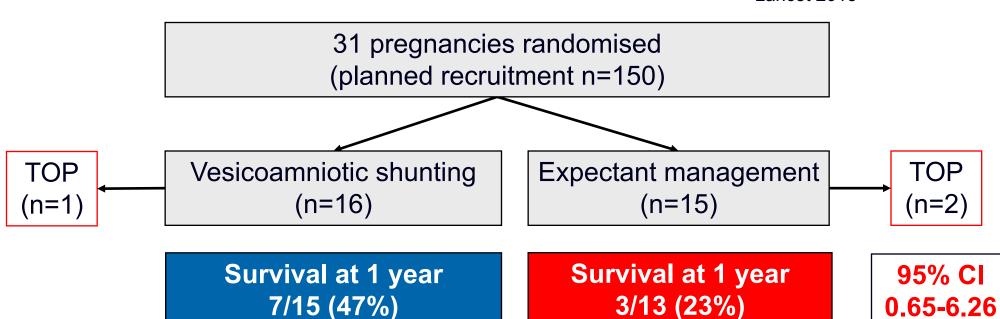
PLUTO trial

0.65-6.26

Percutaneous vesicoamniotic shunting versus conservative management for fetal lower urinary tract obstruction (PLUTO): a randomised trial

Rachel K Morris, Gemma L Malin, Elisabeth Quinlan-Jones, Lee J Middleton, Karla Hemming, Danielle Burke, Jane P Daniels, Khalid S Khan, Jon Deeks, Mark D Kilby, for the Percutaneous vesicoamniotic shunting in Lower Urinary Tract Obstruction (PLUTO) Collaborative Group

Lancet 2013

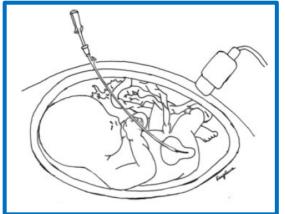




Cystoscopic laser

Fetal cystoscopy





Laser ablation of posterior urethral valves

- Fetal intramuscular anesthesia
- 1-3 mm curved fetoscope
- ND:YAG or Diode laser

	Shunt (n=16)	Laser (n=34)	Nothing (n=61)
Survival at 6m	44%	38%	20%
Good renal function	60%	75%	39%

Laser complications

Urological fistulas 9% Preterm delivery 20%





Ruano et al, 2014



Invasive testing

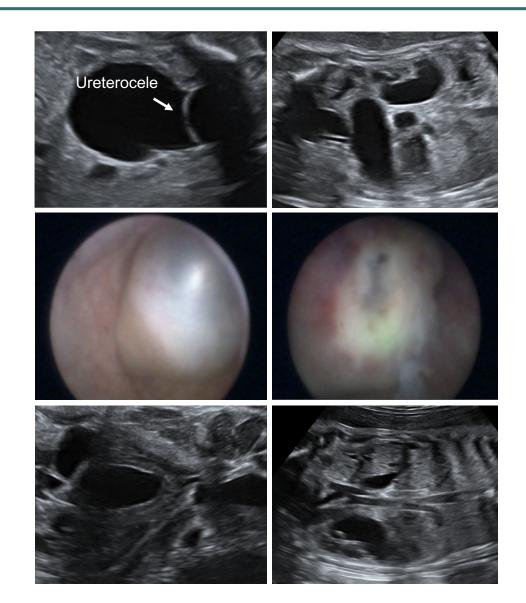
Rupture of membranes after invasive procedures

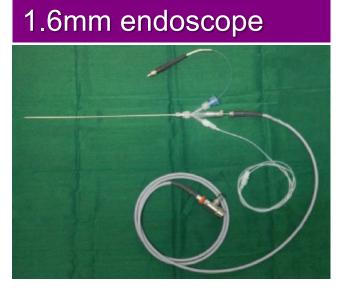


Entry (mm)	PROM (%)	Procedure
<1	0.2	Amniocentesis
2	10	Fetoscopy
3	20	Fetoscopy, Feto-amniotic shunt
4	40	Fetoscopy



Cystoscopic laser for fetal ureterocele







Persico et al, UOG 2017



Amniotic fluid and renal function

Normal amniotic fluid



Olygohydramnios



Anhydramnios



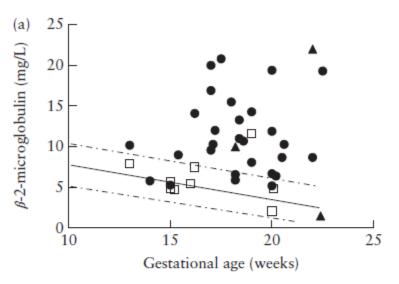
Systematic review - n=215

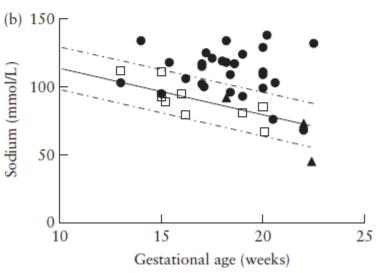
Outcome measures: Serum creatinine Renal histology

	Sensitivity	FPR
Olygo-anhydramnios	63%	24%
Echogenic kidneys	57%	16%

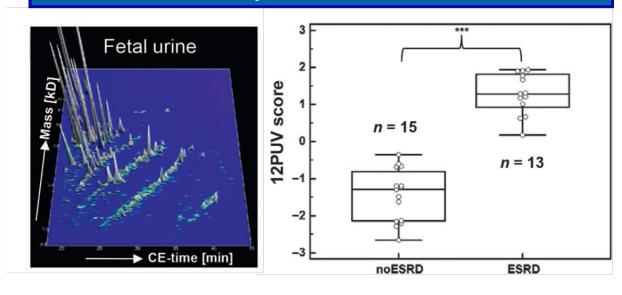


Fetal urine analysis





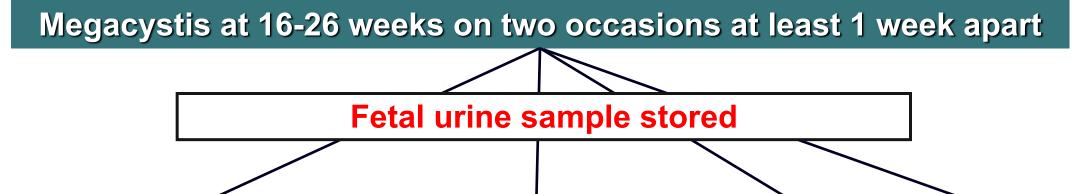
Proteomic analysis



	Sensitivity	FPR
ß-2 microglobulin	81%	11%
Sodium	61%	0%
Proteomic	88%	5%



Proteomic urine analysis



Vesicoamniotic shunt

Cystoscopic laser

Expectant

TOP

Postnatal renal function or histology

Proteomic fetal urine analysis



Heidelberg
University Hospital











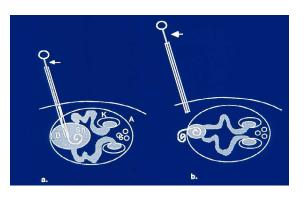
Conclusions

Prenatal management of LUTO: a lot more to do

- Early screening and diagnosis
- Optimal timing for fetal intervention
- Selection criteria based on reliable assessment of fetal renal function
- Further investigation on the best technique







Conclusions

Centralization of cases

Bonelli Centre for the prevention of renal disease

- Obstetrics
- Pediatric Urology
- Pediatric Nephrology
- Neonatal Intensive Care
- Genetics
- Psychology



Mangiagalli Centre, Milan, Italy

Next Webinar: 8th January 2019

Access for PD and HD in children

Rukshana Shroff London, UK

