PRETERM BIRTH PREDICTION AND INTERVENTION

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OBJECTIVES

- 1. To present an overview of Preterm Birth
- 2. To review current techniques of preterm birth prediction and intervention
- 3. To apply current techniques of prediction and intervention to patients at risk for Preterm Birth

PRETERM BIRTH

- Birth $> 20 \, 0/7$ weeks but $< 34 \, 0/7$ weeks
- Preterm Labor
- Preterm PROM
- Cervical insufficiency (incompetency)
- Not indicated preterm birth

PRETERM BIRTH

- U.S.A. 11.7% (MOD, year 2011)
- Texas 13.1% (Healthy Texas Babies)
- El Paso 14.2% (Healthy Texas Babies)
- 2009: 1,984 preterm births in El Paso

PRETERM BIRTH

- Spontaneous preterm birth is leading cause of prematurity
- 15% of all preterm birth are recurrent
- Etiology of preterm birth is heterogeneous
- Therefore, prevention and treatment must be multifactorial

Short Cervix and Preterm Birth

Women with short cervix (< 20 to 25 mm) by vaginal ultrasound examination have a 75% chance of preterm delivery!!!

Preterm Birth Prediction

- 1. History
- 2. Cervical Length
- 3. Fetal fibronectin

History of Preterm Birth

1. non-Hispanic white: 11%

Hispanic: 11%

African American: 17%

- 2. Number of prior preterm births: each confers a 1.5X risk in subsequent pregnancies
- 3. Gestational age at prior preterm birth less than 32 weeks: an additional 1.5X risk

Clinical Example

African American woman with 2 prior preterm births at 28 and 26 weeks respectively

Now pregnant; determine risk of preterm birth without intervention:

 $17\% \times 1.5 \times 1.5 \times 1.5 \times 1.5 = 86\%$ risk of PTB

Cervical Length

- Vaginal ultrasound between 18-24 weeks
- Shortest of 3 measurements
- CL < 25 mm
- Funneling with short cervix is not significant, CL is utmost importance

Universal Cervical Length Screening

- Controversial
- Pros: may reduce PTB rate, effective interventions are available, cost-effective
- Cons: quality assurance of screening test, geographic restrictions, unnecessary and unproven interventions

ACOG's Position

"...does not mandate universal cervical length screening in women without a prior preterm birth; (however), this screening strategy may be considered."

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Fetal Fibronectin

- Glue-like protein in membranes
- Theory is that "unstable" or "leaky" membranes are associated with an increase in preterm birth risk
- Dacron swab inserted using sterile speculum in the posterior fornix or blindly in the vagina
- Do between 22-35 weeks

Fetal Fibronectin Results

• If negative, less than 1% risk of preterm birth within the next 2 weeks

• If positive, approximately 16% risk of preterm birth within the next 2 weeks

Preterm Birth Interventions

- Recurrent PTB risk: 17-P (Makena)
- Short cervix by scan: Vaginal progesterone or cerclage
- Lifestyle changes
- Preterm Birth Prevention Clinic

17-P

- Weekly injections from 16-36 weeks
- FDA-approved: Makena (start by 20 6/7 weeks)
- ACOG-recommended (start by 24 weeks)
- Standard of Care
- Reduces recurrent preterm birth by 35%

17-P

- History of preterm birth
- Not for short cervix with no history of PTB
- Not for multiple gestations
- Not for use with vaginal progesterone
- Can be used in patients with a cerclage and history of preterm birth

17-P for Your Patients

- Usually a benefit of commercial insurance plans
- Texas Medicaid: if compounding pharmacy within 50 miles, compound 17-P; if not, Makena
- Watson Pharmaceuticals, manufacturer of Makena, will provide Makena to uninsured patients, at least at the present

Know Your Compounding Pharmacy

- 1. Variability in dosage, potency, and purity
- Some patients may be allergic to oils used as the depo agent
- 3. Gluteal abscesses have been reported, some locally and some serious

Progesterone Gel or Suppositories

- No history of preterm birth
- Short CL found on vaginal ultrasound
- Use until 36 weeks
- Reduces preterm birth by 45%

Cervical Cerclage

- "Purse-string" stitch in cervix
- History of incompetent cervix
- Short cervix by vaginal ultrasound without a history of preterm birth
- "Stitch in time, saves nine (months)"

Useful Algorithm

- Prior loss due to CI: cerclage
- Prior preterm birth: 17-P
- Short cervix, no history, < 24 weeks: vaginal progesterone
- Prior preterm birth, on 17-P, short cervix, <24 weeks: consider cerclage

Cervical Cerclage

- McDonald cerclage: easy and effective, 0-Prolene, 1 purse-string stitch
- Shirodkar: difficult to place and remove and not more effective than McDonald
- Best not to use Mersilene band unless you have a desire to remove stitch under anesthesia

Cervical Cerclage

- Prophylactic: 10-14 weeks
- Therapeutic: up to 24-26 weeks
- Remove at 36 weeks
- PPROM: to remove or keep, controversial

Vaginal Progesterone

- 90 mg (gel) or 200 mg (suppository) daily
- Crinone 8%: 1 applicator vaginally q hs
- Crinone gel has a bioadhesive that keeps the gel attached to the vaginal mucosa

Preterm Birth Prevention Clinic

- Focuses on preterm birth prevention
- Uses state of the art diagnostics and therapeutics
- Based on Utah model that showed a 28% reduction in recurrent preterm birth

Manuck et al. Am J Obstet Gynecol 2011;204:320.e1-6

Goals of Preterm Birth Prevention Clinics

- 1. Reduce morbidity and mortality associated with preterm birth
- 2. Educate patients and providers about preterm birth prediction and interventions
- 3. Provide support to patients at risk for preterm birth
- 4. Lower healthcare costs attributable to preterm birth

Summary of ACOG Level A Evidence

- Singleton pregnancy with prior preterm singleton birth, give 17-P
- Short cervix (< 20 mm) before 24 weeks, give vaginal progesterone
- No progesterone in multiple gestations

Inpatient Management of Preterm Labor

- 1. Diagnosis
- 2. Management

Diagnosis of Preterm Labor

- Regular uterine contractions with change in dilatation, effacement, or both
- Regular uterine contractions with cervical dilatation of 2 cm on admission

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Diagnostic Aids for Diagnosing PTL

- Fetal fibronectin
- Cervical length per vaginal ultrasound

No Intervention

- 30% of patients admitted with PTL will have spontaneous resolution
- 50% of patients admitted with PTL will deliver at term
- No interventions < 24 weeks or > 34 weeks (few exceptions)
- If no cervical change, do not treat contractions

Worthless PTL Interventions

- Prophylactic tocolytic therapy
- Home uterine activity monitoring
- Cerclage, unless indicated
- Narcotic sedation
- Bed rest
- Hydration
- IV antibiotics unless documented infection or positive Group B Strept status

Effective PTL Interventions

- Antenatal corticosteroids (24-34 weeks)
- Short course of tocolytic therapy during "steroid window"
- Antibiotics for positive Group B Strept status
- Magnesium sulfate for neuroprotection

Antenatal Corticosteroids

- Standard of care since NIH Consensus Conference (1992)
- Betamethasone or dexamethasone
- Administer between 24-34 weeks
- May give one additional rescue course of steroids if > 7 days since last course (ACOG, 2011)

Short Course of Tocolytic Therapy

- Use for 48 hours or less
- Beta-adrenergic receptor agonists (terbutaline).
- Calcium channel blockers (nifedipine)
- NSAIDS if less than 32 weeks (indomethacin)

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Tocolytics

- Terbutaline: limited use, beware FDA's blackbox designation
- NSAIDS: indomethacin 100 mg rectally then 50 mg q 6 hours
- Nifedipine: preferred agent, 20 mg orally, then every 3-6 hours

Group B Strept

- Treat if positive culture
- If no culture, treat pending culture
- If arrested PTL and positive status, must treat again prior to delivery

Magnesium Sulfate Neuroprotection

- Reduces cerebral palsy by 17-45%
- Use between 24-31 6/7 weeks
- 3 protocols available: Crowther, Marret, and Rouse

Rouse Protocol

- Loading dose: 6 grams
- Maintenance dose: 2 grams/hour
- Use until delivery or for 12 hours, whichever comes first
- If < 6 hours after D/C, restart maintenance dose
- If > 6 hours after D/C, reload
- Generally, do not give additional tocolytics

The End

Thanks for your attention

Questions?