South Australian Perinatal Practice Guidelines

Preterm Labour & Birth

Prevention, Diagnosis & Management

© Department for Health and Wellbeing, Government of South Australia. All rights reserved.

Note:

This guideline provides advice of a general nature. This statewide guideline has been prepared to promote and facilitate standardisation and consistency of practice, using a multidisciplinary approach. The guideline is based on a review of published evidence and expert opinion.

Information in this statewide guideline is current at the time of publication.

SA Health does not accept responsibility for the quality or accuracy of material on websites linked from this site and does not sponsor, approve or endorse materials on such links.

Health practitioners in the South Australian public health sector are expected to review specific details of each patient and professionally assess the applicability of the relevant guideline to that clinical situation.

If for good clinical reasons, a decision is made to depart from the guideline, the responsible clinician must document in the patient's medical record, the decision made, by whom, and detailed reasons for the departure from the guideline.

This statewide guideline does not address all the elements of clinical practice and assumes that the individual clinicians are responsible for discussing care with consumers in an environment that is culturally appropriate and which enables respectful confidential discussion. This includes:

- The use of interpreter services where necessary,
- Advising consumers of their choice and ensuring informed consent is obtained,
- Providing care within scope of practice, meeting all legislative requirements and maintaining standards of professional conduct, and
- Documenting all care in accordance with mandatory and local requirements

Explanation of the aboriginal artwork

The Aboriginal artwork used symbolises the connection to country and the circle shape shows the strong relationships amongst families and the aboriginal culture. The horse shoe shape design shown in front of the generic statement symbolises a woman and those enclosing a smaller horse shoe shape depicts a pregnant woman. The smaller horse shoe shape in this instance represents the unborn child. The artwork shown before the specific statements within the document symbolises a footprint and demonstrates the need to move forward together in unison.

Australian Aboriginal Culture is the oldest living culture in the world yet Aboriginal people continue to experience the poorest health outcomes when compared to non-Aboriginal Australians. In South Australia, Aboriginal women are 2-5 times more likely to die in childbirth and their babies are 2-3 times more likely to be of low birth weight. The accumulative effects of stress, low socio economic status, exposure to violence, historical trauma, culturally unsafe and discriminatory health services and health systems are all major contributors to the disparities in Aboriginal maternal and birthing outcomes. Despite these unacceptable statistics the birth of an Aboriginal baby is a celebration of life and an important cultural event bringing family together in celebration, obligation and responsibility. The diversity between Aboriginal cultures, language and practices differ greatly and so it is imperative that perinatal services prepare to respectfully manage Aboriginal protocol and provide a culturally positive health care experience for Aboriginal people to ensure the best maternal, neonatal and child health outcomes.

Purpose and Scope of PPG

This PPG is intended to guide primary care, emergency, specialist and midwifery practitioners in the care of women at risk of or experiencing preterm labour. It provides a standardised, evidence-based approach to the assessment and management of South Australian women presenting with threatened preterm labour and includes a statewide referral pathway for women presenting in regional/lower acuity centres. For the first time, this PPG includes recommendations for how to identify women at risk of preterm birth and strategies for prevention. It forms the basis of "The Whole Nine Months SA" campaign to safely reduce rates of preterm birth in South Australia.

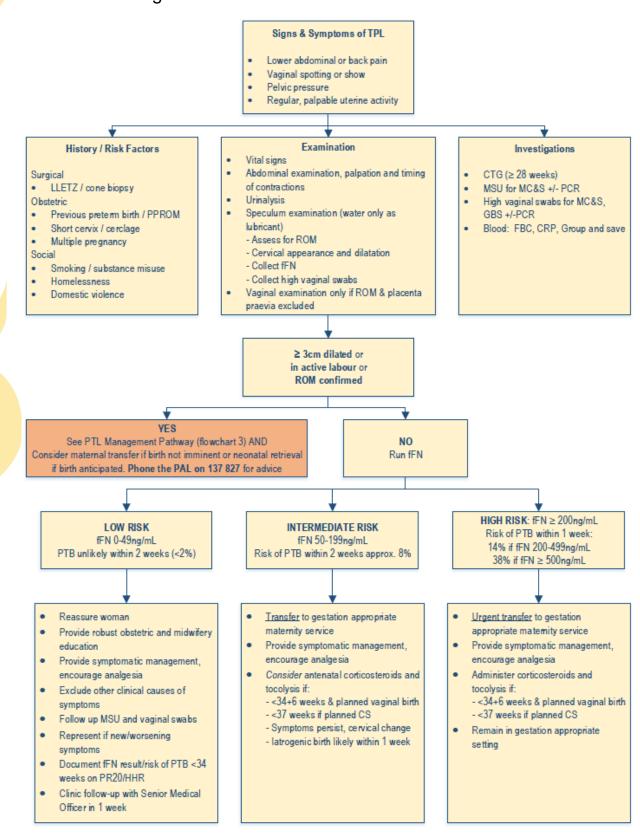


INFORMAL COPY WHEN PRINTED Page 1 of 33

Public-I4-A4

Prevention, Diagnosis & Management

Flowchart 1: Threatened Preterm Labour Assessment and Transfer – Regional Sites

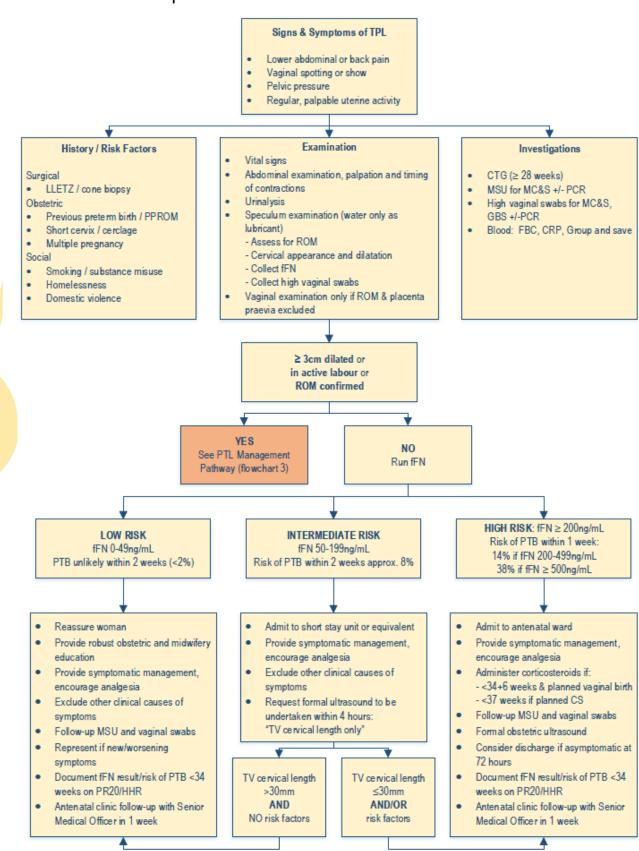




INFORMAL COPY WHEN PRINTED Page 2 of 33

Prevention, Diagnosis & Management

Flowchart 2: Assessment of Threatened Preterm Labour – Metropolitan Sites

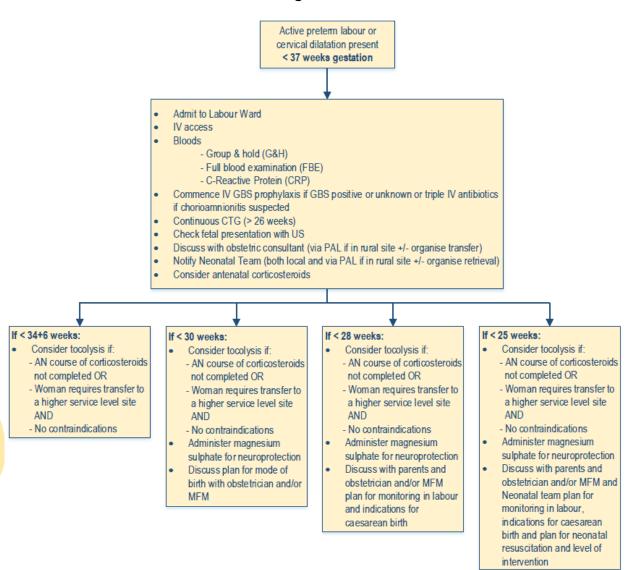




INFORMAL COPY WHEN PRINTED Page 3 of 33

Prevention, Diagnosis & Management

Flowchart 3: Preterm Labour Management



See body of PPG for detailed information and treatment regimens for:

- Antenatal Corticosteroids
- Tocolysis with nifedipine (first line) or salbutamol (second line)
- Magnesium Sulphate for Neuroprotection of the Fetus



INFORMAL COPY WHEN PRINTED Page 4 of 33

Prevention, Diagnosis & Management

Table 1: Risk Factors for Preterm Birth and Recommended Actions – Quick Reference

| Risk Factors | | Action |
|---|-----------------------------------|--|
| Maternal | | |
| Age | <20 | Continuity of Carer |
| | | Postnatal LARC |
| | >35 | Consider aspirin (Appendix 1) |
| Ethnicity | ATSI | Refer to AFBP |
| | Indian, African, Indo-Caribbean | Continuity of Carer |
| Cervical Surgery | Especially >10mm, | Cervical length at morphology |
| On a self-all the day Assessible | repeated LLETZ or Cone Biopsy | On Continue that are collected |
| Congenital Uterine Anomalies | 10 1 00 | Cervical length at morphology |
| BMI | <18 and >30 | Optimise BMI pre-pregnancy |
| Medical Comorbidities | Hypertension, DM, Renal | Consider aspirin (Appendix 1) Optimise pre-pregnancy |
| Medical Comorbidities | Disease, SLE, APLS, | Consider aspirin (Appendix 1) |
| | Scleroderma | Multi-disciplinary Team Care |
| Nutrition | Vegetarian/Non-Fish Diet | Omega 3 & Zinc Supplements |
| 1 danieli | Malabsorption/Inflammatory | omoga o a zine cappiomonio |
| | Bowel Disease/Gastric Banding | |
| | Previous PTB/At risk woman | Screen for Vitamin D deficiency |
| Smoking | | Screen at booking & every third |
| - | | visit as per SAPR |
| | | Refer to Quitline |
| Obstetric History | | |
| Previous preterm birth/PPROM/ | | Refer to Obstetrician/MFM |
| cerclage/shortened cervix | | |
| Previous fully dilated CS, STOP, GTOP | | Cervical length at morphology |
| Pregnancy Features | | |
| Shortened Cervix | <25mm on TVUS, especially | Urgent referral to Obstetrician/ |
| | <10mm or funnelling | MFM |
| Short Interpregnancy Interval | Especially <6 but up to 18 months | Continuity of Carer |
| | months | Optimise nutrition & medical comorbidities |
| | | Postnatal LARC |
| ART/IVF | | Single Embryo Transfer |
| 74(171) | | Consider aspirin (Appendix 1) |
| Urogenital Infections | All Women | Screen, culture & treat UTI |
| - 3 | Symptomatic Women | Culture & treat urogenital |
| | | infections |
| | History of infection associated | Refer to Obstetrician/MFM |
| | losses and PTB | |
| NA MILE | e.g. chorioamnionitis | |
| Multiple pregnancy | | Refer to Obstetrician. Refer to |
| | | MFM if MCDA, DCDA complexity |
| | | or higher multiple Consider aspirin (Appendix 1) |
| Social Factors | | Consider aspirin (Appendix 1) |
| | | |
| Low SES/Intimate Partner Violence | | Continuity of Carer |
| Low SES/Intimate Partner Violence | | Continuity of Carer |
| Low SES/Intimate Partner Violence | | Any available enhanced antenatal |
| Low SES/Intimate Partner Violence | | Any available enhanced antenatal care programs |
| Low SES/Intimate Partner Violence ATSI | | Any available enhanced antenatal care programs Refer to Social Work if indicated |
| | | Any available enhanced antenatal care programs |

Prevention, Diagnosis & Management

Table of Contents

| Purpose and Scope of PPG | |
|---|----|
| Flowchart 1: Threatened Preterm Labour Assessment and Transfer – Regional Sites | 2 |
| Flowchart 2: Assessment of Threatened Preterm Labour – Metropolitan Sites | 3 |
| Flowchart 3: Preterm Labour Management | 4 |
| Table 1: Risk Factors for Preterm Birth and Recommended Actions - Quick Reference | 5 |
| Summary of Practice Recommendations | 8 |
| Abbreviations | 9 |
| Definition | 9 |
| Background | 10 |
| Prevention | |
| Maternal Risk Factors | 10 |
| Non-modifiable Maternal Risk Factors | 10 |
| Age | 10 |
| Ethnicity | 10 |
| Cervical Surgery | |
| Congenital Uterine Anomalies | |
| Modifiable Maternal Risk Factors | |
| BMI | |
| Medical Comorbidities | |
| Nutrition | |
| Smoking | |
| Obstetric History | |
| Previous Preterm Birth | |
| Obstetric Cervical Trauma | |
| Pregnancy Features | |
| Shortened Cervix | |
| Interpregnancy Interval | |
| Assisted Reproductive Technologies | |
| Urogenital Infections | |
| | |
| Systemic Infections | |
| Multiple Pregnancies | |
| Social Determinants | |
| Aboriginal and Torres Strait Islander Women | |
| Socially Disadvantaged and Vulnerable Women | |
| Antenatal Care Provision | |
| Midwifery Continuity of Carer | |
| Dedicated Preterm Birth Clinics | |
| Threatened Preterm Labour | |
| Cervical Length | |
| Fetal Fibronectin | |
| Combined Cervical Length and Fetal Fibronectin | |
| Threatened Preterm Labour - Assessment (Flowcharts 1 & 2) | |
| Signs and Symptoms | |
| History and Risk Factors | |
| Examination | |
| Investigations | |
| Threatened Preterm Labour Management (Flowcharts 1 & 2) (Appendix 2) | |
| Fetal fibronectin 0-49 ng/mL | |
| Fetal fibronectin 50-199 ng/mL | |
| Fetal fibronectin >200 ng/mL | 19 |
| Fetal fibronectin >500 ng/mL | 19 |

Prevention, Diagnosis & Management

| Antenatal Corticosteroids | . 20 |
|--|-----------------|
| Tocolysis (First Line) – Nifedipine | . 21 |
| Tocolysis (Second Line) - Salbutamol | |
| Preterm Labour Management (Flowchart 3) | |
| Antibiotics | |
| Magnesium Sulphate | . 24 |
| Mode of Birth | |
| References | |
| Appendices | . 30 |
| Appendix 1. Major Risk Factors for Preeclampsia and Aspirin Prophylaxis | |
| Appendix 2: Stratification of Preterm Birth Risk by fFN Concentration in Symptomatic Won | |
| | . 31 |
| Appendix 3: Table 2. Management of Cervical Head Entrapment at Vaginal Breech Delivery | [,] 31 |
| Acknowledgements | |



INFORMAL COPY WHEN PRINTED

Prevention, Diagnosis & Management

Summary of Practice Recommendations

No pregnancy should be ended before 39 weeks without obstetric or medical indication.

Women with chronic hypertension, type 1 or type 2 diabetes, renal disease or autoimmune conditions such as antiphospholipid syndrome, systemic lupus erythematous and scleroderma should commence low dose (100mg) oral aspirin nocte in early pregnancy.

Consider omega-3 and zinc supplementation in women at risk of dietary insufficiency.

Screen and treat vitamin D insufficiency in women at risk of preterm birth.

Ask about smoking in pregnancy initially and at every third visit if still smoking; refer women to Quitline and the "Quit for You, Quit for Two" app.

Vaginal progesterone from 16 to 36 weeks should be considered for women with a singleton pregnancy and a history of preterm birth.

Women with a history of preterm birth, PPROM, recurrent mid-trimester loss, shortened cervix and cervical cerclage should be referred to an obstetrician and/or MFM Unit, ideally for pre-conception counselling or as early as possible in pregnancy.

Measurement of cervical length is recommended at all mid-trimester morphology scans for women with risk factors. It should be considered for all other women.

- A closed length of ≥35mm is adequate.
- A closed cervical length of ≤25mm (via transvaginal ultrasound) is considered shortened and must be immediately referred to an obstetrician/MFM Unit.

Vaginal progesterone is recommended for women with a shortened cervix of ≤25mm at mid-trimester ultrasound screening.

Pregnancies should be spaced at least 12 months apart.

ART should be used carefully and judiciously, including single embryo transfer only, to minimise the risk of multiple pregnancy.

Routine urogenital antenatal screening for infection must include syphilis serology, a mid-stream urine sample to detect asymptomatic bacteriuria and screening for Chlamydia in women <25 years of age or otherwise at high risk.

There is insufficient evidence for routine screening and treatment of bacterial vaginosis; women at high risk of preterm birth should be referred to an obstetrician or MFM subspecialist to consider their individual risk profile.

All pregnant women should have the influenza vaccine.

Multiple pregnancies are at very high risk of preterm birth and should be managed by an obstetrician or MFM subspecialist.

Aboriginal and Torres Strait Islander women should be offered specialised and culturally appropriate antenatal care featuring continuity of carer and community stewardship.

Pregnant women with identified social risk factors for preterm birth such as young age, substance abuse and exposure to intimate partner violence should be allocated to enhanced models of antenatal care where available and referred for supporting services.

Women at high risk of preterm birth should be prioritised to receive antenatal care in a model that affords midwifery continuity of carer.

Antenatal corticosteroids are most effective if given between 48 hours and 7 days before birth.

A fFN reading of between 10-49ng/mL modestly increases the woman's overall risk (8.2%) of experiencing a preterm birth before 34 weeks, therefore she should be counselled to represent with persistent, worsening or new symptoms.

Consider tocolysis for women who have not completed a full course of corticosteroids or for inutero transfer of a woman to a higher level service.

Do not continue nifedipine beyond 48 hours. A repeat course of nifedipine should only be considered if the woman represents with PTL. Nifedipine is NOT to be used prophylactically.

Commence IV antibiotic prophylaxis for GBS for women in active preterm labour.

Intravenous magnesium sulphate must be administered to any woman considered at imminent risk of preterm birth or where birth is planned at <30 weeks.



INFORMAL COPY WHEN PRINTED Page 8 of 33

Prevention, Diagnosis & Management

Abbreviations

| AFI | Amniotic Fluid Index |
|-------|---|
| APH | Antepartum haemorrhage |
| APLS | Anti-Phospholipid Syndrome |
| ATSI | Aboriginal and Torres Strait Islander |
| BMI | Body Mass Index |
| BV | Bacterial vaginosis |
| CoC | Continuity of Care |
| CS | Caesarean Section |
| CTG | Cardiotocography |
| DASSA | Drug and Alcohol Services South Australia |
| DCDA | Dichorionic Diamniotic (Twin Pregnancy) |
| DM | Diabetes Mellitus |
| fFN | Fetal fibronectin |
| GTOP | Genetic Termination of Pregnancy |
| IBD | Inflammatory Bowel Disease e.g. Crohn's, Ulcerative Colitis |
| IPV | Intimate Partner Violence |
| LARC | Long Acting Reversible Contraceptive |
| LLETZ | Large loop excision of the transformation zone |
| MC&S | Microscopy, Culture and Sensitivities |
| MCDA | Monochorionic Diamniotic (Twin Pregnancy) |
| MDT | Multi-Disciplinary Team |
| MFM | Maternal Fetal Medicine |
| MSU | Mid-stream urine |
| NAAT | Nucleic Acid Amplification Test |
| PPG | Perinatal Practice Guideline |
| PPH | Postpartum haemorrhage |
| PPROM | Preterm prelabour rupture of membranes |
| PTB | Preterm Birth |
| RCT | Randomised controlled trial |
| RDI | Recommended daily intake |
| SALHN | Southern Adelaide Local Health Network |
| SA PR | South Australian Pregnancy Record |
| SES | Socioeconomic Status |
| SET | Single Embryo Transfer |
| SLE | Systemic Lupus Erythematosus |
| STOP | Surgical termination of pregnancy |
| TVUS | Transvaginal Ultrasound |
| UTI | Urinary tract infection |
| WCHN | Women's and Children's Health Network |

Definition

Preterm birth A birth occurring before 37 weeks completed gestation



Prevention, Diagnosis & Management

Background

Late preterm births, occurring between 34⁺⁰ and 36⁺⁶ weeks, account for over 70% of all preterm births, but early preterm births occurring before 34 weeks are responsible for the majority of associated morbidity and mortality.¹

The rate of preterm birth is steadily rising in most jurisdictions including South Australia, where 7.2% of live births and 9.6% of all births in 2017 were preterm; the rate of preterm birth in South Australian Aboriginal women is much higher at 19.1%.² It is important for clinicians to note that **the rate of** *iatrogenic* preterm births has doubled in the last 20 years.¹ This is despite the incidence of hypertensive disorders of pregnancy, small for gestational age babies and smoking in pregnancy all decreasing over the same time; instead, South Australian women are birthing at older ages with higher body mass indices (BMIs) and we have a more ethnically diverse population.¹

Prevention

A simple way of minimising the burden of prematurity in South Australia is to avoid further 'obstetric creep' or iatrogenic delivery at increasingly earlier gestations. Therefore:

No pregnancy should be ended before 39 weeks without obstetric or medical indication.

Spontaneous preterm birth by contrast, is the culmination of woman- and fetus-specific risk factors unique to each pregnancy and therefore, is more helpfully considered a syndrome rather than a diagnosis. By extension, the prevention of spontaneous preterm birth requires a multidisciplinary and multidimensional approach.

It is important for clinicians to be able to identify early in a woman's pregnancy if she is at increased risk of preterm birth, in order to implement risk minimisation strategies and assign an appropriate model of care. Risk factors can be grouped by relation to the woman, her obstetric history, features of the current pregnancy and social determinants of health (see Table 1).

Maternal Risk Factors

Non-modifiable Maternal Risk Factors

Age

Young women (<18 years old) may be physiologically immature and are more likely to be experiencing disadvantageous social circumstances,³ whilst older women (>35 years old) are more likely to have medical comorbidities such as diabetes and obesity contributing to their risk.⁴

Ethnicity

Aboriginal, South Asian, African, African American and Indo-Caribbean women are at higher risk of experiencing preterm and early preterm birth compared to other ethnicities.^{2,5} Emerging evidence suggests at least some of this risk may be attributable to genetic polymorphic differences in immune response and differences in the vaginal microbiome.^{6,7}

Cervical Surgery

A history of cervical procedures such as LLETZ and cervical conisation is associated with increased risk of preterm birth, especially with increased size of excision (>10mm), repeated procedures and conisation.⁸ As a minimum standard of care these women should be identified and referred for a cervical length measurement at morphology ultrasound.



INFORMAL COPY WHEN PRINTED Page 10 of 33

Prevention, Diagnosis & Management

Congenital Uterine Anomalies

Women with congenital uterine anomalies are at increased risk of preterm birth and should have a mid-trimester cervical length measured. Whilst the risk of preterm birth appears greatest in women with major fusion defects (unicornuate, bicornuate and didelphys anomalies), the value of combined cervical length and fetal fibronectin (fFN) in predicting preterm birth is greater for those with resorptive anomalies (septate and arcuate anomalies). 10

Women with a congenital uterine anomaly or a history of cervical procedures such as LLETZ and conisation are at increased risk of preterm birth and should have a cervical length measured and reported at their mid-trimester morphology ultrasound.

Modifiable Maternal Risk Factors

BMI

A powerful modifiable risk factor is BMI; women with low BMIs (<18) are at greater risk of spontaneous preterm birth, whereas women with high BMIs (>30) are more likely to experience an iatrogenic preterm birth secondary to oxidative stress-related complications such as preeclampsia and fetal growth restriction.¹¹ Where possible, women should be encouraged and assisted to optimise their BMI before conceiving.

Medical Comorbidities

Medical comorbidities such as hypertension, diabetes, renal disease and autoimmune conditions should also be optimised prior to conception and managed in a multidisciplinary team, together with relevant specialists and obstetric physicians. A simple but powerful prevention strategy in addition to optimising these medical comorbidities is commencing low dose aspirin 100mg nocte once pregnancy is confirmed (preferably after confirmation of fetal heart beat), prior to 16 weeks' gestation. (Appendix 1)

Women with chronic hypertension, type 1 or type 2 diabetes, renal disease or autoimmune conditions such as antiphospholipid syndrome, systemic lupus erythematous and scleroderma should commence low dose (100mg) oral aspirin nocte in early pregnancy.

Nutrition

Nutrition is integral to a healthy pregnancy outcome and even in a high resource setting such as South Australia, some groups of women are at risk of malnutrition. Women with extremes of BMI (low and high), a history of bariatric procedures, inflammatory bowel disease, those who follow a vegan or vegetarian diet, recent migrants and women of low socioeconomic status are at particular risk of micronutrient insufficiencies in pregnancy. Specific nutrients that have been linked with preterm birth prevention include omega 3, vitamin D and zinc.

Omega 3

A Cochrane systematic review of omega 3 supplementation in pregnancy found an 11% reduction in preterm births and a 42% reduction in early preterm births, however, a single large randomised controlled trial (RCT) (Omega-3 to Reduce the Incidence of Preterm Birth – ORIP) in the Australian setting showed no benefit of *universal* high-dose omega 3 supplementation on preterm birth rates. ^{13,14} Of note, ORIP enrolled multiple pregnancies, including triplets, as well as women who were already supplementing with low-dose omega 3. Further exploration of the ORIP data has revealed that in singleton pregnancies, a low baseline total omega 3 concentration (<4.2% total blood fatty acids) in early pregnancy was associated with increased risk of preterm birth and that *targeted* supplementation of these pregnancies significantly reduced the risk of preterm birth. ¹⁵ As we do not routinely screen for total omega 3 concentrations in early pregnancy, we suggest that women who do not eat *fatty* fish such as salmon, herring, mackerel and sardines once a week or *any* fish 2-3 times a week should consider supplementing with a good quality marine (fish or algal) omega 3 source.



INFORMAL COPY WHEN PRINTED Page 11 of 33

Prevention, Diagnosis & Management

Zinc

A Cochrane systematic review of zinc supplementation in pregnancy showed a 14% relative reduction in preterm birth, though the majority of data came from resource poor settings, making general application in an Australian setting difficult.¹⁶ The Australian recommended daily intake (RDI) of zinc in pregnancy is 11mg/day, however, as absorption is greater from animal rather than plant sources, vegans and strict vegetarians require intakes that are 50% higher. The main sources of zinc are meat, fish and poultry and to a lesser extent, dairy, nuts and cereals.¹⁷

Consider omega-3 and zinc supplementation in women at risk of dietary insufficiency.

Vitamin D

As regards vitamin D, the latest Cochrane systematic review has not shown universal supplementation in pregnancy to decrease preterm birth directly, however, there remains evidence that vitamin D adequacy decreases the risk of preeclampsia, gestational diabetes, low birth weight and postpartum haemorrhage (PPH). Therefore, we recommend screening for and optimising cholecalciferol levels in women at risk of preterm birth, particularly those with a history of prior preterm birth, in addition to women at risk of vitamin D insufficiency (see *Vitamin D Status in Pregnancy* PPG available at www.sahealth.sa.gov.au/perinatal). 18

Screen and treat vitamin D insufficiency in women at risk of preterm birth.

Smoking

Smoking remains an important modifiable risk factor for numerous pregnancy complications, including preterm birth. All pregnant women should be asked about their smoking status at booking and if still smoking, at every third antenatal visit in pregnancy as per the South Australian Pregnancy Record. Women who continue to smoke should be offered referral to the National Quitline Number (13 7848) (or complete the online referral form) and encouraged to use tools such as the "Quit for You, Quit for Two" app.

Ask about smoking in pregnancy initially and at every third visit if still smoking; refer women to Quitline and the "Quit for You, Quit for Two" app.

Obstetric History

Previous Preterm Birth

Clinically, the single greatest predictor of preterm birth is a history of preterm birth; the earlier the gestation at which birth occurred, the greater the risk. A woman with one previous preterm birth has a 4-fold increased risk in her second pregnancy, rising to 6.5-fold after two preterm births. Women with a history of preterm birth should be referred to an obstetrician and/or Maternal Fetal Medicine Unit, ideally preconceptionally, to assess contributing factors and optimise modifiable risk factors, as well as to identify women that may benefit from early interventions such as vaginal progesterone, serial cervical length measurement and prophylactic cervical cerclage.

There is much controversy regarding the use of vaginal progesterone for the prevention of preterm birth in women with a history of preterm birth, with two large recent RCTs contradicting the earlier results of systematic reviews and meta-analyses. ²⁰⁻²² Until updated meta-analyses are performed, current advice remains as per the Royal Australian and New Zealand College of Obstetricians and Gynaecologists (RANZCOG). ²³

Vaginal progesterone from 16 to 36 weeks should be *considered* for women with a singleton pregnancy and a history of preterm birth.

Where a woman's history of preterm birth is suggestive of cervical insufficiency, she should have serial ultrasound cervical length surveillance implemented from 14-16 weeks under the supervision of an obstetrician and/or MFM subspecialist.^{24,25} Further shortening may then justify an ultrasound indicated cervical cerclage.

Prophylactic cerclage is considered for some women with a history of recurrent mid-trimester losses and preterm birth but decisions around timing, approach and the implementation of a cervical cerclage should be undertaken by, or in consultation with, MFM.



INFORMAL COPY WHEN PRINTED Page 12 of 33

Prevention, Diagnosis & Management

Women with a history of preterm birth, PPROM, recurrent mid-trimester loss, shortened cervix and cervical cerclage should be referred to an obstetrician and/or MFM Unit, ideally for pre-conception counselling or as early as possible in pregnancy.

Obstetric Cervical Trauma

As with gynaecological cervical procedures, an obstetric history of cervical trauma including surgical termination of pregnancy (STOP), induced abortions, genetic terminations of pregnancy (GTOP) and previous fully dilated caesarean sections increase future risk of preterm birth.²⁶⁻²⁸

Women with any history of obstetric cervical trauma are at increased risk of preterm birth and should have a cervical length measured at the mid-trimester morphology ultrasound.

Pregnancy Features

Shortened Cervix

Whilst this document has so far focused on women with risk factors to be specifically referred for a cervical length measurement at the mid-trimester morphology ultrasound, we know that many women experiencing pregnancy loss and preterm birth have no identifiable risk factors. Cervical length in mid-trimester is highly correlated with gestation at birth.²⁵ We therefore strongly advocate that it become the standard of care that all South Australian radiology practices performing mid-trimester morphology scans report on the cervical length of **all** pregnant women.

Measurement of cervical length is recommended at all mid-trimester morphology scans. Where the cervix can be imaged clearly on trans-abdominal scan, a closed length of ≥35mm is adequate. In all other cases, trans-vaginal scanning is required, where a closed cervical length of ≤25mm is considered shortened and must be immediately referred to an obstetrician/MFM Unit.

Trials have consistently shown that where a woman is found to have a shortened cervix on ultrasound, treatment with progesterone prolongs the duration of pregnancy and improves pregnancy outcomes. Vaginally administered progesterone appears to have better evidence than intramuscular formulations. As per RANZCOG²³:

Vaginal progesterone is *recommended* for women with a shortened cervix of ≤25mm at mid-trimester ultrasound screening.

Interpregnancy Interval

A short interpregnancy interval, particularly <6 months but also between 6-18 months after the last, increases the risk of a subsequent preterm birth.^{29,30} Where possible, women should be counselled to delay subsequent pregnancies for 12 months and offered appropriate contraception to achieve this

Pregnancies should be spaced at least 12 months apart from the end of one pregnancy to the beginning of the next.

Assisted Reproductive Technologies

Women who required assisted reproductive technology (ART) to conceive, in particular *in vitro* fertilisation (IVF) and intracytoplasmic sperm injection (ICSI), are at increased risk of preterm birth, however, other than limiting the number of embryos transferred (single embryo transfers only), there is no specific intervention to mitigate this risk.³¹ These technologies, including ovulation induction, should be used carefully and judiciously to minimise the numbers of multiple pregnancies that occur as a result. Optimisation of weight, nutrition and medical comorbidities should occur prior to use of ART and consideration be given to whether women meet criteria for low dose aspirin prophylaxis after conceiving.

ART should be used carefully and judiciously, including single embryo transfer only, to minimise the risk of multiple pregnancy.



INFORMAL COPY WHEN PRINTED Page 13 of 33

Prevention, Diagnosis & Management

Urogenital Infections

The initiation of labour is an inflammatory event and many cases of preterm birth are associated with systemic or urogenital infection. Urogenital infection is a strong risk factor for preterm birth. Routine screening and treatment of urogenital infection in pregnancy currently consists of:

- Syphilis serology screening with antenatal screening bloods (with additional screening in the third trimester (28 and 36 weeks), at birth and 6 weeks postnatally for women in outbreak areas or otherwise at high risk);
- A mid-stream urine (MSU) microscopy, culture and sensitivity performed together with antenatal screening bloods;
- Screening for chlamydia (+/- gonorrhoea) in pregnant women <25 years old or at high risk of a sexually transmitted infection.

Routine urogenital antenatal screening for infection must include syphilis serology, a midstream urine sample to detect asymptomatic bacteriuria and screening for Chlamydia in women <25 years of age or otherwise at high risk.

Whilst periodontal disease and bacterial vaginosis in pregnancy have been associated with preterm birth, causation has not been established and universal treatment of these conditions has not reduced the incidence of preterm birth. 32,33 There is some evidence that for women with a history of preterm birth, prophylactic treatment with antibiotics from the mid-trimester where bacterial vaginosis is detected, may improve outcomes. 34 Also, in a separate systematic review, treatment with oral clindamycin (rather than vaginal clindamycin or metronidazole) before 22 weeks of gestation for asymptomatic bacterial vaginosis decreased preterm births <37 weeks and late miscarriages. 35 Women who are *symptomatic* with bacterial vaginosis or other genital tract infections should be treated.

Recent research has identified changes in the vaginal microbiome related to pregnancy, as well as ethnic variation. Some vaginal microbiomes and taxa are associated with increased risk of preterm birth, such as BV-associated bacterium-1, and may present a screening and therapeutic target, although this research is still underway.^{6,36}

There is insufficient evidence for routine screening and treatment of bacterial vaginosis; women at high risk of preterm birth should be referred to an obstetrician or MFM subspecialist to consider their individual risk profile.

Systemic Infections

Intercurrent systemic infections with pathogens such as malarial parasites or influenza are associated with preterm birth. These should be identified and treated. In the case of influenza, the Australian Government recommends and provides free vaccination to all pregnant women in recognition of their status as a vulnerable population.³⁷

All pregnant women should have the influenza vaccine.

Multiple Pregnancies

Three-quarters (75%) of multiple births are preterm in South Australia.³⁸ The increased risk of general (hypertensive, cholestatic, diabetic) and multiple-specific (e.g. twin-twin transfusion, selective intra-uterine growth restriction) pregnancy complications, as well as the risk of stretch mechanoreceptor initiation of labour, contribute to both iatrogenic and spontaneous preterm births in multiple pregnancies, respectively.³⁹ Aside from minimising the risk of multiple gestations arising from ART, twin and higher order multiple pregnancies require specialised care with either an obstetrician or MFM subspecialist.

Multiple pregnancies are at very high risk of preterm birth and should be managed by an obstetrician or MFM subspecialist.



INFORMAL COPY WHEN PRINTED Page 14 of 33

Prevention, Diagnosis & Management

Social Determinants

Social disadvantage has consistently been linked with preterm birth and is associated with a range of independent risk factors such as poor nutrition, substance abuse, periodontal disease, lower educational attainment and psychological stressors. 40-42 Women in lower socioeconomic groups are also more likely to be over-represented in other at-risk groups related to ethnicity, extremes of age and BMI. High-grade evidence to support enhanced antenatal care programs for socially disadvantaged women to reduce preterm births in high-income countries is lacking and methodologically challenging, with many studies showing decreases in preterm birth outcomes that were not statistically significant or of borderline statistical significance when compared to standard care models. 43 This, however, does not necessarily translate to a lack of clinical significance, particularly where individual outcomes and the cost, financial and otherwise, of a single early preterm birth are taken into account.

Aboriginal and Torres Strait Islander Women

Australian studies looking at antenatal care delivery to Aboriginal and Torres Strait Islander women have similarly suffered a lack of strong evidence, likely related to small sample sizes, heterogeneity of care delivery and reporting of outcomes.⁴⁴

A recent prospective cohort study conducted in Brisbane, Australia, found an impressive 50% reduction in preterm birth in Aboriginal women receiving 'culturally safe continuity of carer within a holistic service with high levels of community investment-ownership-activation and health service leadership across partner organisations.'45 We note that 25% of identified Indigenous births at the health service were excluded from the analysis, including women allocated to other specialised care such as MFM and drug and alcohol dependency services, nevertheless, the translatable absolute reduction in preterm births suggests this to be a worthwhile model of antenatal care for a group of women known to be at significantly increased risk of preterm birth.

Closer to home, Aboriginal women participating in the South Australian Aboriginal Family Birthing Program highly value receiving care from another Aboriginal woman, paving the way for greater engagement with antenatal care and improved outcomes. These women achieved similar birth outcomes, including preterm birth rates, to Aboriginal women in other models of care, despite having greater medical and social complexity.⁴⁶

Aboriginal and Torres Strait Islander women should be offered specialised and culturally appropriate antenatal care featuring continuity of carer and community stewardship.

Socially Disadvantaged and Vulnerable Women

A lack of high-quality evidence similarly pertains to enhanced antenatal care provision to other socially disadvantaged women such as teenagers, pregnant women with substance abuse issues and women in low socioeconomic groups: whilst studies suggest improved outcomes including preterm birth rates, they suffer from selection bias and other methodological limitations.⁴³

Women experiencing domestic or intimate partner violence (IPV) are represented among all socioeconomic groups and are at high risk of preterm birth; the importance of screening for IPV has been recognised by RANZCOG and the Australian Government by linking it to a number of antenatal (and postnatal) Medicare item numbers.⁴⁷

Pragmatically, despite low grade evidence, socially vulnerable women should be offered enhanced antenatal and adjunct service provision including prioritisation to midwifery continuity of care models and other specialised antenatal care streams (e.g. Strengthening Links (WCHN), Early Links (SALHN)), as well as access to Drug and Alcohol Services South Australia (DASSA), social work and mental health services, to optimise their individual health and pregnancy outcomes, as any reduction in preterm births in terms of absolute numbers will still be clinically and economically meaningful.

Pregnant women with identified social risk factors for preterm birth such as young age, substance abuse and exposure to intimate partner violence should be allocated to enhanced models of antenatal care where available and referred for supporting services.



INFORMAL COPY WHEN PRINTED Page 15 of 33

Prevention, Diagnosis & Management

Antenatal Care Provision

Midwifery Continuity of Carer

There is good quality evidence that midwifery continuity of carer in pregnancy reduces preterm birth, with a reported risk reduction of 24%. Although none of the included studies had adequate blinding, this is unlikely to have affected outcomes, but only low-risk and mixed-risk women were included; women with 'significant maternal disease and substance abuse' were excluded from some trials, subgroups of women at high risk of experiencing preterm birth. Nevertheless, the findings raise important considerations regarding the value of therapeutic relationship continuity, which supersedes management and informational continuity, and how this may improve the pregnancy outcomes of women engaging in various models of antenatal care.

Women at high risk of preterm birth should be prioritised to receive antenatal care in a model that affords midwifery continuity of carer.

Dedicated Preterm Birth Clinics

The provision of antenatal care in dedicated preterm birth clinics helps to facilitate the streamlining of multidisciplinary care and predictive and preventive measures such as urogenital tract infections screening, second trimester alkaline phosphatase measurement, serial fetal fibronectin quantification, access to serial transvaginal cervical length measurement and midwifery and medical continuity of carers. Evidence from such clinics in the UK and Australia, both alone and as part of a comprehensive state-wide intervention, have demonstrated significant reductions in the incidence of preterm birth and improved prediction of women requiring inpatient management and timing of interventions such as antenatal corticosteroids^{49,50} This despite systematic reviews failing to show statistically significant improvements in preterm birth prevention as with other models of antenatal care.^{51,52} The high personal and societal cost of each preterm birth together with the array of interventions requiring individualisation to each at-risk woman's circumstances justify the need for dedicated preterm birth clinics and parent's anxiety and satisfaction with care must be factored.⁵¹

Women at high-risk of preterm birth benefit from specialised antenatal care to provide timely and evidenced access to predictive and preventive interventions; local health networks should strongly consider the establishment of dedicated preterm birth prevention clinics to facilitate this.



INFORMAL COPY WHEN PRINTED

Prevention, Diagnosis & Management

Threatened Preterm Labour

Presentations with threatened preterm labour are common and rarely result in preterm birth within 1-2 weeks.

Nevertheless, many of these women will remain at higher risk of delivering preterm later in pregnancy. The timing of interventions such as antenatal corticosteroids, magnesium sulphate and antibiotics is crucial to improving outcomes for those babies that do go on to be born prematurely – the earlier the gestation, the more important the timing of these interventions. ^{53,54}

Antenatal corticosteroids are most effective if given between 48 hours and 7 days before birth.

Repeated doses of antenatal corticosteroids are associated with less clear evidence of benefit and are associated with poorer long-term learning and behavioural outcomes; the optimal dosing regime is also uncertain. The greatest clinical challenge, therefore, is determining which women presenting with signs and symptoms of threatened preterm labour are at greatest risk of progressing to preterm birth in the next one to two weeks.

Cervical Length

Cervical length in the mid-trimester has now for some time been recognised as highly correlated with risk of preterm birth.^{55,56} Even in symptomatic women at later gestations, a longer cervical length is still protective in terms of predicting who will progress to preterm birth.⁵⁷

Fetal Fibronectin

After identification of the glycoprotein fetal fibronectin (fFN) in 1988, it was quickly identified as being clinically useful in helping to predict which women presenting with symptoms of threatened preterm labour were at greatest risk of preterm birth.⁵⁸ Fetal fibronectin promotes adhesion between the fetal chorion and maternal decidua and is typically absent from cervicovaginal secretions between 24 and 36 weeks' gestation. In 2013 it was shown that quantifying the level of fFN in symptomatic women enhances its predictive value over qualitative measurement alone.⁵⁹ Other biomarkers present in vaginal secretions are promoted as predictors of preterm birth but revert to a qualitative measurement and have not been found to have as high a negative predictive value, of great import when deciding not to administer interventions to improve neonatal outcomes.⁶⁰

Combined Cervical Length and Fetal Fibronectin

Combining the use of cervical length measurement and quantitative fFN further improves the predictive ability of clinicians to determine which women are at highest risk of progressing to preterm birth in the next seven days and therefore more accurately and beneficially targeting interventions such as the administration of antenatal corticosteroids, magnesium sulphate and antibiotics. 61-64 A risk threshold of 5% is generally used as a cut-off for determining who requires intervention, however, the gestation of the pregnancy and context may factor into clinical decision making.

Recognising that access to timely (<4 hours from presentation) and accurate cervical length measurement is challenging in most maternity settings and presentations, this guideline focuses on the use of quantitative fFN for triaging women presenting with signs and symptoms of threatened preterm labour to determine who requires transfer and intervention. Where available, the addition of cervical length can further assist with clinical decision making around administration of antenatal corticosteroids and magnesium sulphate and assist with discharge planning to local and distant settings.



INFORMAL COPY WHEN PRINTED Page 17 of 33

Prevention, Diagnosis & Management

Threatened Preterm Labour - Assessment (Flowcharts 1 & 2)

Signs and Symptoms

Women may present with:

- Lower abdominal cramping, regular, painful contractions or tightenings;
- Lower back pain;
- Vaginal or rectal pressure;
- · Vaginal discharge including bleeding;
- PPROM: one-third of women will experience premature prelabour rupture of membranes (PPROM) before regular, painful tightenings. Intrauterine infection is a major concern – manage as per the *Preterm Prelabour Rupture of Membranes* PPG available at www.sahealth.sa.gov.au/perinatal)

History and Risk Factors

A brief history should review the woman's risk factors for preterm birth including:

- · Congenital uterine anomalies;
- Previous LLETZ or cone biopsies;
- Any previous mid-trimester losses, PPROM and preterm births and the gestations at which these occurred;
- Women with a cervical cerclage, multiple pregnancy or polyhydramnios: these women are at higher risk of preterm birth and require MFM input;
- Women with social risk factors: a lower threshold for admission should be considered to obtain assistance and minimise ongoing risk where possible.

Examination

Initial triage and midwifery assessment should incorporate:

- Maternal vital signs including temperature;
- Fetal heart rate assessment either by Doppler auscultation or CTG at >28/40;
- · Assessment of vaginal losses;
- Abdominal examination including presentation;
- Timing of uterine activity;
- Urinalysis.

Further assessment (within a midwife's or doctor's scope of practice) should incorporate:

- A speculum examination using only water as lubrication, assessing for:
 - PPROM, including any ancillary tests/glass slide for ferning;
 - Cervical length, dilatation and appearance;
 - Cervical and vaginal swabs for MC&S and Chlamydia/Gonorrhoea NAAT;
 - A fFN swab from the posterior fornix.

Please note that whilst recent digital or ultrasound vaginal examination, coitus and the presence of blood in the vagina are relative contraindications to performing a fFN measurement and increase the risk of both false and true positive results, a negative fFN in these situations is still clinically valid and should be performed if it will help appropriately time intervention and/or discharge planning.

If PPROM and placenta praevia have been excluded:

 A digital vaginal examination to assess the dilatation of the cervix; this may be repeated in 4 hours' time to help determine if the woman is in active labour.

If bedside ultrasound is available:

- Confirm presenting part of fetus;
- Assess AFI and umbilical artery Dopplers.



INFORMAL COPY WHEN PRINTED Page 18 of 33

Prevention, Diagnosis & Management

Investigations

- Urine MC&S;
- Cervical and vaginal swabs for NAAT and MC&S, respectively;
- Bloods including full blood examination (FBE), group and hold and C reactive protein (CRP);
- fFN quantification
- TVUS cervical length if fFN >50ng/mL
- A formal ultrasound assessment of growth should form part of the work up if the woman requires admission.

Threatened Preterm Labour Management (<u>Flowcharts 1</u> & <u>2</u>) (<u>Appendix 2</u>)

Fetal fibronectin 0-49 ng/mL

These women have a very low risk (<2%) of experiencing a preterm birth in the next 2 weeks and can therefore be managed symptomatically and discharged with timely (within 1 week) antenatal follow up.

Be aware that a reading of between 10-49ng/mL modestly increases the woman's overall risk (8.2%) of experiencing a preterm birth before 34 weeks, therefore she should be counselled to represent with persistent, worsening or new symptoms and her antenatal care provider should remain vigilant to the risk of a preterm birth later in pregnancy.

Fetal fibronectin 50-199 ng/mL

These women are at a modestly increased risk (7.7%) of progressing to a preterm birth within 14 days. They should be transferred to a gestation appropriate maternity setting for a full assessment including growth ultrasound and transvaginal cervical length assessment (Flowchart 3). Antenatal corticosteroids should be *considered* but not automatically commenced as very few babies will benefit from them within 7 days. Any urogenital infections should be screened for and treated if clinically indicated or confirmed.

Fetal fibronectin >200 ng/mL

These women are increased risk of delivering within 1 week (14%) and must be transferred to a gestation appropriate maternity and nursery setting (Flowchart 3) with commencement of an antenatal corticosteroid course (with tocolysis) if less than 34+6 weeks gestation. They may need to remain in this setting even on discharge as their risk of delivery within 2 weeks is as high as 29%.

Fetal fibronectin >500 ng/mL

These women are at very high risk of preterm birth: 38% within 1 week and 47% within 2 weeks. They must be urgently transferred to a gestation appropriate maternity and nursery setting (Flowchart 3) and remain there for the duration of their pregnancy or until they attain a gestation suitable to be transferred to a lower acuity centre. Antenatal corticosteroids must be commenced as a priority with tocolysis if appropriate (see below).



INFORMAL COPY WHEN PRINTED Page 19 of 33

Prevention, Diagnosis & Management

Antenatal Corticosteroids

| Initial course | Indications: | | |
|-----------------------|--|--|--|
| | Gestational age is between 23⁺⁰ and 34⁺⁶ weeks; Risk of imminent preterm birth, even if within 24 hours; Preterm birth is planned or expected within the next seven days; Elective caesarean section at less than 37⁺⁰ weeks. Dosage: Administer IM betamethasone in two doses of 11.4 mg 24 hours apart to the woman; | | |
| | If betamethasone is unavailable, give IM dexamethasone in two doses of 12 mg, 24 hours apart. | | |
| Repeat dose or course | Indications: When the gestational age is 32 ⁺⁶ days or less, a repeat antenatal corticosteroid dose may be given 7 days or more after the first course in women still considered at risk of early preterm birth | | |
| | Dosage: | | |
| | A single repeat dose of IM betamethasone 11.4 mg IM OR; A repeat course of IM betamethasone in two doses of 11.4 mg 24 hours apart. If betamethasone is unavailable, give IM dexamethasone 12 mg. | | |
| Further doses | Seven days after the first, single repeat dose (and less than 14 days since the initial course), if the woman is still considered to be at risk of preterm birth at a gestation 32⁺⁶ days or less within the next seven days, a further, single repeat dose of antenatal corticosteroids (IM betamethasone 11.4 mg IM) can be given, up to a maximum of three, single repeat doses only. | | |
| | Do not give any further repeat courses if a repeat course of IM betamethasone in two doses (of 11.4 mg 24 hours apart) has already been given. | | |



INFORMAL COPY WHEN PRINTED

Prevention, Diagnosis & Management

Tocolysis (First Line) - Nifedipine

Oral nifedipine is the preferred available tocolytic for suppression of preterm labour (reduced side effects, ease of administration and greater efficacy compared to betamimetics).

Indication:

Tocolysis with nifedipine is **ONLY** indicated if:

- The woman is less than 34+6 weeks pregnant and;
- Requiring transfer to a gestation appropriate maternity service and/or;
- Yet to complete an initial course of antenatal corticosteroids.

Contraindications:

Maternal:

- A systolic blood pressure of <90mmHg
- Cervical dilatation >3cm, especially in the context of PPROM
- Hypersensitivity to nifedipine or any excipients within the formulation
- Cardiac comorbidities including congestive cardiac failure and aortic stenosis
- Hepatic dysfunction
- Preeclampsia/eclampsia
- Concurrent use of IV salbutamol.

Fetal:

- Chorioamnionitis
- Placental abruption/Antepartum Haemorrhage
- Fetal distress
- Severe IUGR
- IUFD
- Known lethal fetal anomalies.

Administration and Dosage

- Confirm threatened or actual preterm labour;
- Check systolic blood pressure >90 mmHg before administering nifedipine;
- Give stat oral dose nifedipine 20 mg; chew or crush* to aid the speed of absorption.

If uterine contractions persist after 30 minutes:

 Give stat oral dose nifedipine 20 mg; chew or crush* to aid the speed of absorption; the maximum dose of nifedipine in the first hour is 40 mg.

If uterine contractions persist after 3 hours:

- Give stat oral dose nifedipine 20 mg; chew or crush* to aid the speed of absorption.
- Administer oral nifedipine 20 mg every three hours for 48 hours (unless contractions cease or the woman establishes in labour).
- The maximum dose of nifedipine is 160 mg in 24 hours

Do not continue nifedipine beyond 48 hours. A repeat course of nifedipine should only be considered if the woman represents with PTL and the same conditions are met. Nifedipine is NOT to be used prophylactically.

* Nifedipine tablets may be crushed to aid administration. Crushed tablets should be administered within 30-60 seconds of crushing to avoid significant loss of potency.



INFORMAL COPY WHEN PRINTED Page 21 of 33

Prevention, Diagnosis & Management

Observations:

- Maternal baseline BP, TPR, FHR before administering the first dose of nifedipine 20 mg
- Continue hourly BP and maternal pulse for four hours
- Temperature every 4 hours
- The rate of observations should be tapered according to the clinical situation
- Continuous CTG while contracting
- Recommence CTG in the presence of:
 - o Regular abdominal pains or tenderness
 - o Change in amount, colour of liquor
 - Antepartum haemorrhage
- And arrange medical review
- Stop nifedipine if:
 - There is marked hypotension e.g. systolic < 90 mm Hg
 - Significant dyspnoea

Tocolysis (Second Line) - Salbutamol

Betamimetics such as salbutamol have historically been used for tocolysis. A 2014 Cochrane review⁶⁵ found that betamimetics can delay birth effectively, however are associated with **serious** (potentially life threatening) maternal side effects including pulmonary oedema.

However, in situations where nifedipine is unavailable, salbutamol may be used with caution as a second-line tocolytic. It is not to be used in conjunction with nifedipine due to a high degree of synergism.

The indication for IV salbutamol requires that:

- The woman is less than 34+6 weeks pregnant and;
- Requiring transfer to a gestation appropriate maternity service.

Contraindications include:

- Maternal:
 - A systolic blood pressure of <90mmHg
 - Cervical dilatation >3cm, especially in the context of PPROM
 - Hypersensitivity to salbutamol or any excipients within the formulation
 - Cardiac comorbidities including congestive cardiac failure and aortic stenosis
 - Hepatic dysfunction
 - o Preeclampsia/eclampsia
 - Insulin-dependent diabetes
 - o Thyroid disease
 - Multiple pregnancy (increased risk of pulmonary oedema)
- Fetal:
 - o Chorioamnionitis
 - o Placental abruption/Antepartum Haemorrhage
 - Fetal distress
 - o Severe IUGR
 - o IUFD
 - o Known lethal fetal anomalies.
 - o Cardiac anomalies

Adverse Reactions include:

- Tachycardia
- Hypotension
- Tremor
- Pulmonary oedema
- Hyperglycaemia
- Hypokalaemia.



INFORMAL COPY WHEN PRINTED Page 22 of 33

Prevention, Diagnosis & Management

- If salbutamol is to be used for tocolysis, an IV infusion/syringe pump must be used for administration
- > Salbutamol should be used with care, as it is associated with maternal tachycardia, hypotension, tremor, pulmonary oedema, hyperglycaemia and hypokalaemia

Preparation of infusion

> Add 5 mg of salbutamol (5mL ampoule, Ventolin Obstetric Injection®) to 100 mL of 0.9% sodium chloride to prepare a 50 microgram / mL solution

Using a medication added label write "salbutamol 50 micrograms per mL" and attach label to syringe

Administration

- > IV infusion/syringe pump must be used for administration
- > Initial rate: 12 mL / hour (10 micrograms / minute)
- > Increments: increase by 4 mL / hour (3.3 micrograms / minute) every 30 minutes until:
 - > Contractions cease or
 - > Maternal pulse rate reaches 120 beats/minute
- > Maximum: 36 mL / hour (30 micrograms / minute)
- > Maintain rate for 1 hour after contractions have stopped, then gradually reduce by half every 6 hour

Do not exceed 48 hours of salbutamol therapy. Only in exceptional circumstances should the treatment be continued for more than 24 hours.

Practice Points

- > The dose is determined by the woman's tolerance (i.e. clinical indicators) of adverse effects against desired response;
- > Women should be warned about tremors, anxiety, dizziness and headaches:
- > Collect baseline electrolytes, urea, creatinine and maternal blood sugar level before commencement of infusion; repeat 4-hourly if abnormal;
- > Perform half hourly maternal pulse, BP and respiratory rate until the maintenance dose is reached;
- Exercise caution with any additional intravenous fluids to avoid fluid overload;
- > Perform cardiovascular examination including auscultation of lung bases once in the first 24 hours of therapy;
- > **Reduce** the infusion if the maternal pulse >120bpm;
- > Cease the infusion and request medical review immediately if there is chest pain, dyspnoea or the respiratory rate >30/min;
- > Continuous electronic fetal heart rate monitoring >28/40; cease the infusion if the fetal heart rate >180bpm;

Betamimetics can cause a fall in serum potassium (K+). This is related to the movement of K+ intracellularly and is usually limited and self-reversing. No treatment is needed unless ECG changes occur or the serum potassium falls below 2.5 mmol/L

INFORMAL COPY WHEN PRINTED Page 23 of 33

Prevention, Diagnosis & Management

Preterm Labour Management (Flowchart 3)

If a woman is more than 3cm dilated at presentation, or contracting regularly with progressive cervical effacement and dilatation, management of preterm labour must be initiated according to the gestational age of the baby and the setting. In units without neonatal facilities suitable for the gestation, consult with tertiary centre. Consider maternal transfer if birth is not imminent (telephone the Perinatal Advice Line on 137 827) or consult with neonatal retrieval service (also phone 137 827) if birth is anticipated (see *Perinatal Advice and Emergency Transport PPG* available at www.sahealth.sa.gov.au/perinatal).

Antibiotics

There is no role for the use of antibiotics to treat threatened preterm labour as doing so has been associated with an increased risk of cerebral palsy. If a woman, however, is in active preterm labour, IV antibiotics for GBS prophylaxis are recommended to reduce the risk of early onset GBS neonatal sepsis. See *Antibiotics in the Peripartum Period* PPG (available at www.sahealth.sa.gov.au/perinatal) for antibiotic choice.

Commence IV antibiotic prophylaxis for GBS for women in active preterm labour.

If there is a significant clinical suspicion of chorioamnionitis, birth should be expedited under intravenous antibiotic cover to improve maternal and neonatal outcomes. See *Antibiotics in the Peripartum Period* PPG (available at www.sahealth.sa.gov.au/perinatal) for antibiotic choices.

Chorioamnionitis:

- · Signs including:
 - Maternal pyrexia;
 - Maternal tachycardia;
 - Fetal tachycardia;
 - Uterine tenderness;
 - Offensive and/or purulent vaginal discharge;
 - Elevated white cell count:
 - Raised C-Reactive Protein.
- Consult with senior obstetrician, MFM and/or Infectious Diseases physician when deciding whether or not to augment labour. Uncertainty in diagnosis, particularly at very preterm gestations may warrant amniocentesis to confirm the clinical suspicion of chorioamnionitis.
- Complete septic screen should include histological and microbiological examination of the fetal and maternal surfaces of the placenta and membranes after birth (see Histopathology Management of the Placenta PPG available at www.sahealth.sa.gov.au/perinatal).

Magnesium Sulphate

Intravenous magnesium sulphate must be administered to any woman considered at imminent risk of preterm birth or where birth is planned at <30 weeks.

The administration of a bolus of magnesium sulphate in women at imminent risk of preterm birth at less than 30 weeks' gestation has been shown to reduce neonatal deaths and cerebral palsy by as much as 15% and cerebral palsy alone by almost 30% with numbers needed to treat of 42 and 63, respectively.⁶⁷ This is a significant intervention clinicians can implement to reduce the burden of preterm birth and improve outcomes once preterm labour is established. We anticipate further advice from the MAGENTA study⁶⁸ as to whether there is benefit for women and fetuses at risk of preterm birth between 30 and 34 weeks in the near future.

In women at risk of imminent preterm birth <30 weeks magnesium sulphate should be administered regardless of:

- Plurality (number of babies in utero);
- The reason the woman is considered to be at risk of preterm birth;
- Parity (number of previous births the woman has had);
- The anticipated mode of birth, and;
- Whether or not antenatal corticosteroids have been given.



INFORMAL COPY WHEN PRINTED Page 24 of 33

Prevention, Diagnosis & Management

When birth is planned, commence magnesium sulphate as close to **four hours before birth** as possible.

Contraindications and Considerations

- Allergy
- Concomitant use of nifedipine (increased risk of hypotension)
- In situations where urgent birth is necessary because of maternal or fetal compromise, the birth should not be delayed to administer magnesium sulphate.

See Magnesium Sulphate for Neuroprotection of the Fetus in Women at Risk of Preterm Birth PPG available at www.sahealth.sa.gov.au/perinatal for full details.

Mode of Birth

There are few clinical trials and little high-quality evidence to guide mode of delivery for preterm birth. The exact gestation at which the birth is occurring, the presentation of the fetus and the clinical circumstances relating to maternal and fetal wellbeing all need to be taken into account. So too does the setting and plans for neonatal resuscitation and clinical experience of the obstetrician, GP obstetrician and/or midwife. A Cochrane systematic review included only 122 women⁶⁹ whilst a recent retrospective analysis of WHO datasets did not include context and reasons for clinical decision making, making interpretation of the results challenging.⁷⁰

Generally, the available evidence suggests a trend to increased maternal morbidity and decreased neonatal death and morbidity with elective caesarean for very preterm births. See *Perinatal Care at the Threshold of Viability PPG* (available at www.sahealth.sa.gov.au/perinatal) for further information to guide decision making at very preterm gestations.

In each circumstance consider:

- Is there clinical justification for delivery by emergency caesarean section on maternal grounds?
- What is the agreed gestation of the fetus?
- Is aggressive neonatal resuscitation planned and available?
- What is the presentation of the fetus?
- Is the labour advanced and are the membranes intact? Emergency caesarean section can result in severe maternal and fetal injury where a preterm fetus with ruptured membranes is deep in the pelvis with little or no lower segment.
- What is the condition of the fetus? For example, is there growth restriction, or a known congenital anomaly or insult? Chorioamnionitis and suspected fetal acidosis are relative contraindications to delivery by caesarean section.
- What is the parity and age of the mother?
- What is the skill and experience of the most senior available accoucheur?

Involve a Senior Obstetrician and/or Maternal Fetal Medicine sub-specialist in your decision-making where possible.

Principles to bear in mind:

- Keeping the membranes intact the delivery of the extremely preterm fetus en caul will
 minimise fetal trauma, whether at vaginal or caesarean birth.
- Birth using vacuum extraction (ventouse) is contraindicated at less than 34 weeks' gestation.
- Birth using forceps is relatively contraindicated at less than 34 weeks' gestation.
- Preterm breech births are at greater risk of cervical head entrapment; if proceeding with a
 vaginal preterm breech delivery, be prepared to incise the cervix and aim to ensure
 adequate maternal analgesia i.e. epidural (see <u>Table 2</u>).
- A Classical caesarean section can facilitate ease of delivery, minimise fetal trauma and is
 preferable to an extended lower segment incision (J or T incision). This does however have
 implications for future pregnancy management, and a significantly increased risk of
 placenta accreta.



INFORMAL COPY WHEN PRINTED Page 25 of 33

Prevention, Diagnosis & Management

References

- 1. Verburg PE, Dekker GA, Venugopal K, Scheil W, Erwich J, Mol BW, et al. Long-term Trends in Singleton Preterm Birth in South Australia From 1986 to 2014. Obstet Gynecol. 2018;131(1):79-89.
- 2. Pregnancy Outcome in South Australia 2016. Adelaide: Pregnancy Outcome Unit, Prevention and Population Health Branch, SA Health, Government of South Australia; 2018.
- 3. Fraser AM, Brockert JE, Ward RH. Association of young maternal age with adverse reproductive outcomes. N Engl J Med. 1995;332(17):1113-7.
- 4. Fuchs F, Monet B, Ducruet T, Chaillet N, Audibert F. Effect of maternal age on the risk of preterm birth: A large cohort study. PLoS One. 2018;13(1):e0191002.
- 5. Puthussery S, Li L, Tseng PC, Kilby L, Kapadia J, Puthusserry T, et al. Ethnic variations in risk of preterm birth in an ethnically dense socially disadvantaged area in the UK: a retrospective cross-sectional study. BMJ Open. 2019;9(3):e023570.
- 6. Fettweis JM, Serrano MG, Brooks JP, Edwards DJ, Girerd PH, Parikh HI, et al. The vaginal microbiome and preterm birth. Nat Med. 2019;25(6):1012-21.
- 7. Zhang G, Feenstra B, Bacelis J, Liu X, Muglia LM, Juodakis J, et al. Genetic Associations with Gestational Duration and Spontaneous Preterm Birth. N Engl J Med. 2017;377(12):1156-67.
- 8. Kyrgiou M, Athanasiou A, Paraskevaidi M, Mitra A, Kalliala I, Martin-Hirsch P, et al. Adverse obstetric outcomes after local treatment for cervical preinvasive and early invasive disease according to cone depth: systematic review and meta-analysis. BMJ. 2016;354:i3633.
- 9. Fox NS, Roman AS, Stern EM, Gerber RS, Saltzman DH, Rebarber A. Type of congenital uterine anomaly and adverse pregnancy outcomes. J Matern Fetal Neonatal Med. 2014;27(9):949-53.
- 10. Ridout AE, Ibeto LA, Ross GN, Cook JR, Sykes L, David AL, et al. Cervical length and quantitative fetal fibronectin in the prediction of spontaneous preterm birth in asymptomatic women with congenital uterine anomaly. Am J Obstet Gynecol. 2019;221(4):341 e1- e9.
- 11. Shennan AH, Girling JC. Premature Labour. BMJ Best Practice. London2018.
- 12. RANZCOG. Guidance regarding the use of low-dose aspirin in the prevention of pre-eclampsia in high-risk women. 2018.
- 13. Middleton P, Gomersall JC, Gould JF, Shepherd E, Olsen SF, Makrides M. Omega-3 fatty acid addition during pregnancy. Cochrane Database Syst Rev. 2018;11:CD003402.
- 14. Makrides M. omega-3 Fatty Acids in Pregnancy: Time for Action. J Nutr. 2019;149(4):549-50.
- 15. Simmonds LA, Sullivan TR, Skubisz M, Middleton PF, Best KP, Yelland LN, et al. Omega-3 fatty acid supplementation in pregnancy-baseline omega-3 status and early preterm birth: exploratory analysis of a randomised controlled trial. BJOG. 2020;127(8):975-81.
- 16. Ota E, Mori R, Middleton P, Tobe-Gai R, Mahomed K, Miyazaki C, et al. Zinc supplementation for improving pregnancy and infant outcome. Cochrane Database Syst Rev. 2015(2):CD000230.
- 17. Saunders AV, Craig WJ, Baines SK. Zinc and vegetarian diets. Med J Aust. 2013;199(S4):S17-21.
- 18. Palacios C, Kostiuk LK, Pena-Rosas JP. Vitamin D supplementation for women during pregnancy. Cochrane Database Syst Rev. 2019;7:CD008873.
- 19. McManemy J, Cooke E, Amon E, Leet T. Recurrence risk for preterm delivery. Am J Obstet Gynecol. 2007;196(6):576 e1-6; discussion e6-7.
- 20. Norman JE, Marlow N, Messow CM, Shennan A, Bennett PR, Thornton S, et al. Vaginal progesterone prophylaxis for preterm birth (the OPPTIMUM study): a multicentre, randomised, double-blind trial. Lancet. 2016;387(10033):2106-16.
- 21. Dodd JM, Jones L, Flenady V, Cincotta R, Crowther CA. Prenatal administration of progesterone for preventing preterm birth in women considered to be at risk of preterm birth. Cochrane Database Syst Rev. 2013(7):CD004947.
- 22. Likis FE, Edwards DR, Andrews JC, Woodworth AL, Jerome RN, Fonnesbeck CJ, et al. Progestogens for preterm birth prevention: a systematic review and meta-analysis. Obstet Gynecol. 2012;120(4):897-907.



Prevention, Diagnosis & Management

- 23. RANZCOG. Progesterone: Use in the second and third trimester of pregnancy for the prevention of preterm birth. Royal Australian and New Zealand College of Obstetricians and Gynaecologists, Melbourne; 2017.
- 24. Owen J, Yost N, Berghella V, Thom E, Swain M, Dildy GA, 3rd, et al. Mid-trimester endovaginal sonography in women at high risk for spontaneous preterm birth. JAMA. 2001;286(11):1340-8.
- 25. Society for Maternal-Fetal Medicine. Electronic address pso, McIntosh J, Feltovich H, Berghella V, Manuck T. The role of routine cervical length screening in selected high- and low-risk women for preterm birth prevention. Am J Obstet Gynecol. 2016;215(3):B2-7.
- 26. Moreau C, Kaminski M, Ancel PY, Bouyer J, Escande B, Thiriez G, et al. Previous induced abortions and the risk of very preterm delivery: results of the EPIPAGE study. BJOG. 2005;112(4):430-7.
- 27. Levine LD, Sammel MD, Hirshberg A, Elovitz MA, Srinivas SK. Does stage of labor at time of cesarean delivery affect risk of subsequent preterm birth? Am J Obstet Gynecol. 2015;212(3):360 e1-7.
- 28. Watson HA, Carter J, David AL, Seed PT, Shennan AH. Full dilation cesarean section: a risk factor for recurrent second-trimester loss and preterm birth. Acta Obstet Gynecol Scand. 2017;96(9):1100-5.
- 29. Kozuki N, Walker N. Exploring the association between short/long preceding birth intervals and child mortality: using reference birth interval children of the same mother as comparison. BMC Public Health. 2013;13 Suppl 3:S6.
- 30. Shachar BZ, Mayo JA, Lyell DJ, Baer RJ, Jeliffe-Pawlowski LL, Stevenson DK, et al. Interpregnancy interval after live birth or pregnancy termination and estimated risk of preterm birth: a retrospective cohort study. BJOG. 2016;123(12):2009-17.
- 31. Cavoretto P, Candiani M, Giorgione V, Inversetti A, Abu-Saba MM, Tiberio F, et al. Risk of spontaneous preterm birth in singleton pregnancies conceived after IVF/ICSI treatment: meta-analysis of cohort studies. Ultrasound Obstet Gynecol. 2018;51(1):43-53.
- 32. Brocklehurst P, Gordon A, Heatley E, Milan SJ. Antibiotics for treating bacterial vaginosis in pregnancy. Cochrane Database Syst Rev. 2013(1):CD000262.
- 33. Schwendicke F, Karimbux N, Allareddy V, Gluud C. Periodontal treatment for preventing adverse pregnancy outcomes: a meta- and trial sequential analysis. PLoS One. 2015;10(6):e0129060.
- 34. Thinkhamrop J, Hofmeyr GJ, Adetoro O, Lumbiganon P, Ota E. Antibiotic prophylaxis during the second and third trimester to reduce adverse pregnancy outcomes and morbidity. Cochrane Database Syst Rev. 2015;1:CD002250.
- 35. Lamont RF, Nhan-Chang CL, Sobel JD, Workowski K, Conde-Agudelo A, Romero R. Treatment of abnormal vaginal flora in early pregnancy with clindamycin for the prevention of spontaneous preterm birth: a systematic review and metaanalysis. Am J Obstet Gynecol. 2011;205(3):177-90.
- 36. Stinson LF, Payne MS. Infection-mediated preterm birth: Bacterial origins and avenues for intervention. Aust N Z J Obstet Gynaecol. 2019;59(6):781-90.
- 37. Immunisation ATAGo. Australian Immunisation Handbook. In: Health AGDo, editor. Canberra2018.
- 38. Pregnancy Outcome in South Australia 2016. Adelaide: SA Health, Government of South Australia, Pregnancy Outcome Unit PaPHB; 2018.
- 39. Murray SR, Stock SJ, Cowan S, Cooper ES, Norman JE. Spontaneous preterm birth prevention in multiple pregnancy. Obstet Gynaecol. 2018;20(1):57-63.
- 40. Goldenberg RL, Culhane JF, lams JD, Romero R. Epidemiology and causes of preterm birth. Lancet. 2008;371(9606):75-84.
- 41. Ncube CN, Enquobahrie DA, Albert SM, Herrick AL, Burke JG. Association of neighborhood context with offspring risk of preterm birth and low birthweight: A systematic review and meta-analysis of population-based studies. Soc Sci Med. 2016;153:156-64.
- 42. Ruiz RJ, Dwivedi AK, Mallawaarachichi I, Balcazar HG, Stowe RP, Ayers KS, et al. Psychological, cultural and neuroendocrine profiles of risk for preterm birth. BMC Pregnancy Childbirth. 2015;15:204.



Prevention, Diagnosis & Management

- 43. Hollowell J, Oakley L, Kurinczuk JJ, Brocklehurst P, Gray R. The effectiveness of antenatal care programmes to reduce infant mortality and preterm birth in socially disadvantaged and vulnerable women in high-income countries: a systematic review. BMC Pregnancy Childbirth. 2011;11:13.
- 44. Rumbold AR, Bailie RS, Si D, Dowden MC, Kennedy CM, Cox RJ, et al. Delivery of maternal health care in Indigenous primary care services: baseline data for an ongoing quality improvement initiative. BMC Pregnancy Childbirth. 2011;11:16.
- 45. Kildea S, Gao Y, Hickey S, Kruske S, Nelson C, Blackman R, et al. Reducing preterm birth amongst Aboriginal and Torres Strait Islander babies: A prospective cohort study, Brisbane, Australia. EClinicalMedicine. 2019;12:43-51.
- 46. Middleton P, Bubner T, Glover K, Rumbold A, Weetra D, Scheil W, et al. 'Partnerships are crucial': an evaluation of the Aboriginal Family Birthing Program in South Australia. Aust N Z J Public Health. 2017;41(1):21-6.
- 47. Donovan BM, Spracklen CN, Schweizer ML, Ryckman KK, Saftlas AF. Intimate partner violence during pregnancy and the risk for adverse infant outcomes: a systematic review and meta-analysis. BJOG. 2016;123(8):1289-99.
- 48. Sandall J, Soltani H, Gates S, Shennan A, Devane D. Midwife-led continuity models versus other models of care for childbearing women. Cochrane Database Syst Rev. 2016;4:CD004667.
- 49. Hughes K, Sim S, Roman A, Michalak K, Kane S, Sheehan P. Outcomes and predictive tests from a dedicated specialist clinic for women at high risk of preterm labour: A ten year audit. Aust N Z J Obstet Gynaecol. 2017;57(4):405-11.
- 50. Min J, Watson HA, Hezelgrave NL, Seed PT, Shennan AH. Ability of a preterm surveillance clinic to triage risk of preterm birth: a prospective cohort study. Ultrasound Obstet Gynecol. 2016;48(1):38-42.
- 51. Malouf R, Redshaw M. Specialist antenatal clinics for women at high risk of preterm birth: a systematic review of qualitative and quantitative research. BMC Pregnancy Childbirth. 2017;17(1):51.
- 52. Whitworth M, Quenby S, Cockerill RO, Dowswell T. Specialised antenatal clinics for women with a pregnancy at high risk of preterm birth (excluding multiple pregnancy) to improve maternal and infant outcomes. Cochrane Database Syst Rev. 2011(9):CD006760.
- 53. Norberg H, Kowalski J, Marsal K, Norman M. Timing of antenatal corticosteroid administration and survival in extremely preterm infants: a national population-based cohort study. BJOG. 2017;124(10):1567-74.
- 54. Haviv HR, Said J, Mol BW. The place of antenatal corticosteroids in late preterm and early term births. Semin Fetal Neonatal Med. 2019;24(1):37-42.
- 55. Heath VC, Souka AP, Erasmus I, Gibb DM, Nicolaides KH. Cervical length at 23 weeks of gestation: the value of Shirodkar suture for the short cervix. Ultrasound Obstet Gynecol. 1998;12(5):318-22.
- 56. Iams JD, Paraskos J, Landon MB, Teteris JN, Johnson FF. Cervical sonography in preterm labor. Obstet Gynecol. 1994;84(1):40-6.
- 57. Bruijn M, Vis JY, Wilms FF, Oudijk MA, Kwee A, Porath MM, et al. Quantitative fetal fibronectin testing in combination with cervical length measurement in the prediction of spontaneous preterm delivery in symptomatic women. BJOG. 2016;123(12):1965-71.
- 58. Lockwood CJ, Senyei AE, Dische MR, Casal D, Shah KD, Thung SN, et al. Fetal fibronectin in cervical and vaginal secretions as a predictor of preterm delivery. N Engl J Med. 1991;325(10):669-74.
- 59. Abbott DS, Radford SK, Seed PT, Tribe RM, Shennan AH. Evaluation of a quantitative fetal fibronectin test for spontaneous preterm birth in symptomatic women. Am J Obstet Gynecol. 2013;208(2):122 e1-6.
- 60. Dawes LK, Prentice LR, Huang Y, Groom KM. The Biomarkers for Preterm Birth Study-A prospective observational study comparing the impact of vaginal biomarkers on clinical practice when used in women with symptoms of preterm labor. Acta Obstet Gynecol Scand. 2019.
- 61. Gomez R, Romero R, Medina L, Nien JK, Chaiworapongsa T, Carstens M, et al. Cervicovaginal fibronectin improves the prediction of preterm delivery based on sonographic cervical length in patients with preterm uterine contractions and intact membranes. Am J Obstet Gynecol. 2005;192(2):350-9.



Prevention, Diagnosis & Management

- 62. Hermans FJR, Bruijn MMC, Vis JY, Wilms FF, Oudijk MA, Porath MM, et al. Risk stratification with cervical length and fetal fibronectin in women with threatened preterm labor before 34 weeks and not delivering within 7 days. Acta Obstet Gynecol Scand. 2015;94(7):715-21.
- 63. Kuhrt K, Smout E, Hezelgrave N, Seed PT, Carter J, Shennan AH. Development and validation of a tool incorporating cervical length and quantitative fetal fibronectin to predict spontaneous preterm birth in asymptomatic high-risk women. Ultrasound Obstet Gynecol. 2016;47(1):104-9.
- 64. Watson HA, Carter J, Seed PT, Tribe RM, Shennan AH. The QUiPP App: a safe alternative to a treat-all strategy for threatened preterm labor. Ultrasound Obstet Gynecol. 2017;50(3):342-6.
- 65. Neilson JP, West HM, Dowswell T. Betamimetics for inhibiting preterm labour. Cochrane Database of Systematic Reviews 2014 [cited 2020, Oct 13], Issue 2. Art. No.: CD004352. DOI: 10.1002/14651858.CD004352.pub3.
- 66. Hughes RG BP, Steer PJ, Heath P, Stenson BM Prevention of early-onset neonatal group B streptococcal disease. 2017.
- 67. Doyle LW, Crowther CA, Middleton P, Marret S, Rouse D. Magnesium sulphate for women at risk of preterm birth for neuroprotection of the fetus. Cochrane Database Syst Rev. 2009(1):CD004661.
- 68. Crowther CA, Middleton PF, Wilkinson D, Ashwood P, Haslam R, Group MS. Magnesium sulphate at 30 to 34 weeks' gestational age: neuroprotection trial (MAGENTA)--study protocol. BMC Pregnancy Childbirth. 2013;13:91.
- 69. Alfirevic Z, Milan SJ, Livio S. Caesarean section versus vaginal delivery for preterm birth in singletons. Cochrane Database Syst Rev. 2013(9):CD000078.
- 70. Thanh BYL, Lumbiganon P, Pattanittum P, Laopaiboon M, Vogel JP, Oladapo OT, et al. Mode of delivery and pregnancy outcomes in preterm birth: a secondary analysis of the WHO Global and Multi-country Surveys. Sci Rep. 2019;9(1):15556.
- 71. Sylvester HL, H-S. Practicalities of preterm delivery. O&G Magazine. 2019;21(1).

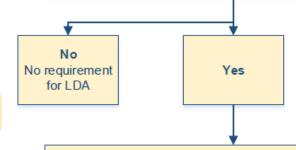


Appendices

Appendix 1. Major Risk Factors for Preeclampsia and Aspirin Prophylaxis¹²

Major Risk Factors Present at First Assessment?

- Preeclampsia in previous pregnancy & birth <37 weeks or HELLP Syndrome
- Predisposing medical conditions
 - Autoimmune
 - Systemic Lupus Erythematosus
 - Scleroderma
 - Anti-Phospholipid Syndrome
 - Chronic hypertension (especially severe)
 - Diabetes type 1 and 2
 - Chronic kidney disease
- Assisted conception with oocyte donation
- Family history of preeclampsia (mother and/or sister)



- Refer for prompt face-to-face specialist consultation
- Prescription for LDA and calcium to be provided by obstetric service (as early in pregnancy as possible but should be prior to 16 weeks)
- If timely access to specialist appointment is a barrier, telephone or telehealth consultation is recommended prior to prescribing

For women with any major risk factor for preeclampsia, metaanalysis suggests a reduction in risk of preeclampsia of 46% (95% CI 30% - 50%) when low dose aspirin (LDA) therapy is initiated in early pregnancy. These women may also benefit from calcium. LDA will reduce the risk from about 1 in 5 to 1 in 10 high risk women.

> Plan and document roles and responsibilities for ongoing antenatal care along with monitoring and timing for follow-up with specialist



INFORMAL COPY WHEN PRINTED

Prevention, Diagnosis & Management

Appendix 2: Stratification of Preterm Birth Risk by fFN Concentration in Symptomatic Women⁵⁹

| fFN Level | N (%) | Delivery ≤ 7 days | Delivery ≤ 14 days | Delivery before 34 wks, 0 days |
|------------------|-----------|-------------------|--------------------|-----------------------------------|
| < 10 ng/mL | 170 (57%) | 1% | 1.8% | 1.5% |
| 10 to 49 ng/mL | 62 (21%) | 0% | 1.6% | 8.2% |
| 50 to 199 ng/mL | 41 (14%) | 0% | 7.7% | 11.5% |
| 200 to 499 ng/mL | 14 (5%) | 14% | 29% | 33% |
| ≥ 500 ng/mL | 13 (4%) | 38% | 46% | 75% |

Appendix 3: Table 2. Management of Cervical Head Entrapment at Vaginal Breech Delivery⁷¹

| Manoeuvres | Description | Risks |
|--|--|---|
| McRoberts position | Flexion of maternal knees so that the anterior aspect of the thighs are pressed against the abdomen | |
| Uterine relaxation | Beta adrenergic agonist (terbutaline 250 µg subcutaneous) Nitroglycerin (50-200 micrograms IV or one metred dose of sublingual spray (400 micrograms) General anaesthesia | Maternal tachycardia Uterine atony |
| Duhrssen's incision | 1–2 fingers placed between the partially dilated cervix and the presenting part, with incisions made along the length of the undilated cervix at 6, 2 and 10 o'clock | Extension of incision to the lower uterine segment or broad ligament Injury to uterine vessels, ureter and bladder Severe haemorrhage Cervical incompetence in subsequent pregnancy |
| Symphysiotomy | Infiltration of the symphysis pubis and overlying skin with local anaesthesia Insertion of firm catheter into the urethra to displace it laterally Incision made over the symphysis to separate it just enough to deliver the head | Pelvic instability, requiring delayed orthopaedic repair |
| Zavanelli manoeuvre and caesarean delivery | Administration of tocolytic and attempt to replace the fetal body into the uterus, followed by caesarean section | Complications of caesarean section Cervical injury and subsequent cervical incompetence |



Prevention, Diagnosis & Management

Acknowledgements

The South Australian Perinatal Practice Guidelines gratefully acknowledge the contribution of clinicians and other stakeholders who participated throughout the guideline development process particularly:

Write Group Lead Dr Monika Skubisz

Write Group Members

Dr Danielle Crosby A/Prof Rosalie Grivell Catherine Leggett Dr Anupam Parange A/Prof Chris Wilkinson

Other major contributors

Prof Maria Makrides
A/Prof Philippa Middleton
Paula Medway
A/Prof Alice Rumbold
Dr John Craven
Dr Christopher Beck
Rachael Yates
Dr Amy Keir
Rebecca Smith

SAPPG Management Group Members

Sonia Angus
Lyn Bastian
Dr Elizabeth Beare
Elizabeth Bennett
Dr Feisal Chenia
John Coomblas
Dr Danielle Crosby
Dr Vanessa Ellison
Jackie Kitschke
Dr Kritesh Kumar
Catherine Leggett
Dr Anupam Parange
Rebecca Smith
A/Prof Chris Wilkinson



INFORMAL COPY WHEN PRINTED

Prevention, Diagnosis & Management

Document Ownership & History

Developed by: SA Maternal, Neonatal & Gynaecology Community of Practice

Contact: <u>HealthCYWHSPerinatalProtocol@sa.gov.au</u>
Endorsed by: Commissioning and Performance, SA Health

Next review due: 25/11/2025

ISBN number: 978-1-76083-340-4

PDS reference: CG141

Policy history: Is this a new policy (V1)? N

Does this policy amend or update and existing policy? Y

If so, which version? **V 7.0**

Does this policy replace another policy with a different title? N

If so, which policy (title)?

| Approval Date | Version | Who approved New/Revised Version | Reason for Change |
|---------------|---------|---|---|
| 1/02/21 | V7.1 | Chair, SA Maternal, Neonatal & Gynaecology Community of Practice | Minor amendment. Recommendation for corticosteroids for elective caesarean section < 37 weeks |
| 25/11/20 | V7 | Deputy CE, Commissioning and Performance Division, SA Department for Health and Wellbeing | Major review |
| 07/09/15 | V6 | SA Health Safety and Quality Strategic Governance Committee | Revised |
| 20/05/14 | V5 | SA Health Safety and Quality Strategic Governance Committee | Revised |
| 22/05/12 | V4 | SA Maternal & Neonatal Clinical Network | Revised |
| 21/03/11 | V3 | SA Maternal & Neonatal Clinical Network | Revised |
| 29/12/08 | V2 | SA Maternal & Neonatal Clinical Network | Revised |
| 10/05/04 | V1 | SA Maternal & Neonatal Clinical Network | Original SA Maternal & Neonatal Clinical Network approved version |

INFORMAL COPY WHEN PRINTED Page 33 of 33