CLINICAL PRACTICE GUIDELINE

Preterm labour

This document should be read in conjunction with the Disclaimer

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Preterm labour (suspected) - MFAU: Quick reference guide

This Quick Reference Guide must be used in conjunction with this whole guideline: Preterm Labour. Medical and midwifery staff should be familiar with the contents of the full guideline.

*Between 23-25 weeks CTG monitoring should be discussed with the senior registrar or consultant.

1. Woman presents to MFAU with suspected preterm labour

2. Confirm Gestational Age

3. Perform
   - Baseline Maternal Vital Signs
   - An Abdominal palpation
   - Auscultate the fetal heart
   - Obtain a MSU

4. Is the woman contracting >1:10?
   - Yes
     - Commence a CTG*
   - No
     - Perform a speculum examination
       - Note cervical dilation
       - Obtain appropriate swabs

5. Is the woman requesting Analgesia?
   - Yes
     - Arrange URGENT review by an Obstetric Registrar or above
   - No

6. Is there evidence of cervical dilation?
   - Yes
   - The Registrar may discharge the woman if the Fetal Fibronectin is negative
   - No
     - Arrange Review by an Obstetric Registrar or above
Nifedipine tocolytic therapy- LBS: Quick reference guide

Contra-indications to the use of nifedipine
- Cardiogenic shock
- Cardiac disease\textsuperscript{2}
- Hypotension\textsuperscript{2}
- Intrauterine infection\textsuperscript{2}
- Intrauterine death\textsuperscript{2}
- Antepartum haemorrhage
- Pre-eclampsia
- Use with betamimetics, such as Salbutamol
- Contra-indications to any suppression of labour
- Concurrent use of rifampicin\textsuperscript{1}

Precautions
- Congestive heart failure – use of nifedipine may precipitate or exacerbate heart failure\textsuperscript{3}
- Magnesium sulphate (MgSO\textsubscript{4}) therapy – nifedipine may increase the risk of hypotension.\textsuperscript{2}
- Enzyme inducing epileptic drugs e.g. carbamazepine, phenytoin, phenobarbitone may increase the metabolism of nifedipine.\textsuperscript{1}

Prior to commencement of nifedipine regimen
1. Insert an intravenous cannula to provide access.
2. Perform baseline blood tests for:
   - urea and electrolytes (U&E’s) and creatinine
   - liver function tests (LFT’s)
Regimen

Initial dose
1. Administer 20mg Nifedipine orally (not slow release).\(^1\)
2. After 30 minutes, if the contractions persist, give another 20mg Nifedipine orally.\(^1\)
3. After another 30 minutes, if the contractions persist, give another 20mg Nifedipine orally.\(^1\)
4. After 3 hours if the contractions still persist, Nifedipine 20mg every 8 hours. The maximum daily dose should not exceed 120mg.\(^1\)

Maintenance dose
A maintenance dose of 20mg three times per day for 48 hours may be given when indicated.

Note
- The maximum dose of Nifedipine is 120mg per day.\(^1\)
- Onset of tocolysis is at 30 to 60 minutes and institution of a second line of tocolysis should not be considered in the first 2 hours. If contractions do not abate after 2 hours a second line tocolysis may be considered by the obstetric consultant on call.
- The first 2 doses should be crushed or chewed to increase the rate of absorption.\(^2\)

Maternal and fetal observations
- Pulse, blood pressure and respirations:
  - half hourly while the woman is contracting
- Continuous electronic fetal heart rate monitoring until the contractions cease.
- Maternal hypotension – treat with intravenous fluids in the first instance.
- Cardiovascular examination including auscultation of lung bases every 8 hours for first 24 hours of therapy.

Side-effects
- Hypotension\(^1,3\)
- Dyspnoea\(^1\)
- Cardiac failure
- Headache\(^1,3\)
- Nausea\(^1,3\)
- Dizziness\(^1,3\)
- Increase in liver enzymes\(^3\)
- Tachycardia
- Facial flushing\(^3\)
- Reflex tachycardia\(^3\)
Preterm labour

Aims
- To diagnose preterm labour i.e. labour at less than 37 completed weeks
- To establish a cause, if possible, of preterm labour, this may allow treatment of the primary cause of the preterm labour e.g. urinary tract infection.
- To assess the maternal and fetal condition in the situation of preterm labour.
- To establish effective suppression of labour (unless contra-indicated) prior to 34 weeks gestation without undue delay.

Key points
1. There is good evidence\(^1\) that tocolysis alone does not improve neonatal outcome. However, tocolysis should be considered if the few days gained can be used for corticosteroid therapy.
2. In the case of preterm labour occurring at a site without appropriate nursery facilities, the time gained with tocolytic therapy can be used for transport.
3. The evidence suggests there is no benefit to the fetus in the following situations:
   - Using tocolytic therapy longer than 48 hours.
   - Prolonged or repeated tocolytic therapy after corticosteroids have been given and are current.
   - Employing tocolysis at a gestation greater than 34 weeks.
4. It may not be appropriate to suppress labour in situations such as:
   - Birth required immediately or as soon as possible because of maternal or fetal condition.
   - Labour is too far advanced to attempt suppression.
   - The fetus is sufficiently mature that the risks of suppression therapy outweighs benefits to the fetus (>34 weeks gestation).
5. The Consultant Obstetrician and / or Senior Registrar on call must be notified of all such admissions, and participate in the decisions regarding treatment. Their involvement in the management plan shall be documented in the woman’s notes.
6. Labour is diagnosed on the basis of regular contractions (at least 1 per 10 minutes) which are associated with effacement and / or dilatation of the cervix.
7. The absence of fetal fibronectin (fFN) in the cervical secretions is a very useful negative predictor of imminent birth (negative predictive value for birth within 7 days 97-98%).
8. Like fFN, a cervical length is a good negative predictor, but not a good positive predictor i.e. greater than or equal to 30mm is highly reassuring.
9. In cases of threatened preterm labour, a threshold of 30 mm has been consistently reported to exclude preterm labour, but there is no threshold of
cervical length that establishes the diagnosis. In women with contractions and cervical length less than 30 mm, additional testing (such as fetal fibronectin) may help predict the patient's risk of preterm delivery within the next several days.  

10. It is important to ascertain maternal and fetal well-being before instituting tocolytic and corticosteroid therapy. Fetal factors such as chorioamnionitis, antepartum haemorrhage and intrauterine growth restriction may make delay unwise.

11. The best results in postponing birth are obtained in women who have intact membranes and who are less than 5 cm dilated. However, ruptured membranes or excess dilatation are not absolute contraindications to treatment.

12. Any CMP client that presents at <37 weeks gestation with a history of regular uterine contractions or suspects TPL must be referred immediately to their supporting hospital for obstetric review.

This clinical guideline applies to women when the outcome for the fetus may be improved by delaying birth.

Admission and investigation

On admission, a thorough assessment of the woman shall include:

- **History** - particularly relating to rupture of the membranes, contractions and antepartum haemorrhage. Gestational age must be confirmed by menstrual history and any available previous ultrasound data.
- **Examination** - noting particularly temperature, uterine tone and tenderness, amniotic fluid volume and fetal size and presentation.
- **Vaginal examination** - a speculum examination shall be performed with full aseptic technique and not touching the cervix with the speculum. Cervical swabs shall be taken for immediate bacteriological assessment. If the cervix is closed and there is no blood or amniotic fluid to be seen in the vagina, a fetal fibronectin (fFN) test shall be performed (see below). Digital examination shall be avoided unless there is a significant possibility of a cord presentation or prolapse, or the cervix cannot be adequately visualised.
  
  - **For women < 33 completed weeks gestation (i.e. 32+6 weeks and below), an experienced practitioner (senior midwife or Registrar or above) should perform the speculum examination [NEW 2018-RCA recommendation].**
- **Urine microbiology** - if a mid-stream urine is unsatisfactory a catheter specimen of urine shall be obtained for microscopy and culture.
- **Ultrasound** - this may be necessary to assess presentation, gestation, fetal weight, fetal normality, and the possibility/advisability of amniocentesis. Additionally, a trans-vaginal ultrasound is the best way of detecting early changes in the internal cervical os in women who continue to contract.
• Electronic fetal heart monitoring (EFM) - shall be performed to assess fetal wellbeing in the case of a viable fetus. Ongoing, continuous electronic fetal monitoring (EFM) at gestations of less than 25 weeks is a decision which shall be taken in discussion with a consultant.

• Amniocentesis - this investigation may be appropriate to assess the presence or absence of intra-amniotic sepsis, or to assess fetal lung maturity. The use of this investigation shall be made only by a consultant obstetrician.

• The on-call Paediatric registrar shall be notified of the woman’s presence in the Birth Suite.

Ruptured membranes
• If there is evidence of ruptured membranes, continue as per the guidelines for premature rupture of membranes.

Maternal fever
• Any maternal temperature of 37.5°C or more MUST lead to formal review of the woman and review of the treatment plan with the Consultant on call.

Fetal fibronectin (fFN) test
• Fetal fibronectin is a screening test used to assess the risk of preterm delivery within the next 7 days. “Point of Care” fFN testing should be utilised in the assessment of preterm labour.

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<th>Indications</th>
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<td>• Symptomatic preterm labour between 24 and 36 weeks gestation</td>
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<td>• Intact membranes and</td>
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<td>• Cervical dilatation less than 3cm</td>
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<thead>
<tr>
<th>Contraindications</th>
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<tbody>
<tr>
<td>• Ruptured membranes</td>
</tr>
<tr>
<td>• Cervical cerclage insitu</td>
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<tr>
<td>• Cervical dilation more than 3cm</td>
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<tr>
<td>• Presence of soaps, gels, lubricants or disinfectants</td>
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<thead>
<tr>
<th>Relative contraindications</th>
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<tbody>
<tr>
<td>• Visual evidence of moderate or gross bleeding</td>
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<tr>
<td>• Within 24 hours of coitus</td>
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<tr>
<td>• A negative fFN result of less than 10ng/mL is still valid if:</td>
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<tr>
<td>➢ A woman reports having intercourse in the previous 24 hours</td>
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<tr>
<td>➢ In the presence of moderate or gross vaginal bleeding</td>
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<tr>
<th>Procedure</th>
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<tr>
<td>• Performed using a sterile speculum examination prior to any examination or manipulation of the cervix or vagina</td>
</tr>
<tr>
<td>• Use only sterile water as a lubricant</td>
</tr>
<tr>
<td>• Obtain sample for testing from the posterior fornix of the vagina</td>
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<tr>
<td>• As per test instructions</td>
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</table>
| fFN < 50ng/mL (negative) | • Low risk of birth within 7 – 14 days.  
• False negative result may occur due to  
  ➢ Use of lubricant with speculum examination.  
  ➢ Intravaginal disinfectants |
|--------------------------|----------------------------------------------------------------------------------|
| fFN ≥ 50ng/mL (positive) | • False positive may occur as a result of recent  
  ➢ Coitus  
  ➢ Digital vaginal examination  
  ➢ Transvaginal ultrasound  
  ➢ Bleeding |

See also section on fetal fibronectin results in section “Management of preterm labour” on next page.

**Transvaginal ultrasound**

Transvaginal ultrasound of the cervical length (TVCL) is an additional screening test that can aid in assessing the risk of preterm delivery. TVCL must be performed by a credentialed clinician. Lack of local capability to perform this test is not a reason for transfer.

**Interpreting TVCL results**

A cervical length less than 15mm is associated with an increased risk of spontaneous preterm birth.

Due to the distances required for transfer from WA regional centres, a TVCL 'cut off' of 20mm is appropriate.
Management of preterm labour

Tocolysis and steroids are the main strategies to manage preterm labour.

Fetal fibronectin results

**Negative fFN and no evidence of cervical change / TVCL > 20mm**

There is a low risk of delivery within the next 7 days therefore:

- If contractions are infrequent / irregular: offer discharge home with follow up as an outpatient within 7 days. Women discharged home should be advised that if they have increased pain or more frequent tightenings / contractions they must contact the hospital again. When this occurs the woman should be advised to attend MFAU for re assessment.
- If contractions are regular and painful: admit for observation, offer analgesia and reassess in 2 hours
- If contractions are persistent and painful: consider steroids and tocolysis.

**Positive fFN and / or evidence of cervical change / TVCL < 20mm**

There is an increased risk of delivery within the next 7 days therefore:

- Admit and offer analgesia
- Administer steroids and commence tocolysis ( if no contraindications)
- Continuous fetal monitoring with a CTG.
- In established labour IV antibiotics should be given

Administration of magnesium sulphate infusion

- For information regarding the administration of antenatal magnesium sulphate prior to preterm birth for neuro protection of the fetus post birth see Clinical Guideline: Preterm Birth: Magnesium Sulphate for Neuroprotection of the Fetus

Administration of corticosteroids

- For prophylaxis against neonatal respiratory distress syndrome, a single course of betamethasone is given to women with a gestation between 23 and 36+6 weeks.
- The course is administered as two intramuscular injections of 11.4 mg of betamethasone given 24 hours apart.² ³
- No repeat courses are given. In the event of maternal diabetes, and difficulty controlling the blood glucose, level, the on call Diabetes Physician shall be consulted regarding blood glucose level (BGL) control.
Administration of antibiotic therapy

- If progressive labour occurs group B Streptococcus antibiotic prophylaxis shall be prescribed as per Clinical Guideline Group B Streptococcus
- If evidence of urinary tract sepsis is seen on urine microscopy antibiotics shall be prescribed. See Clinical guideline Antibiotic Treatment for Urinary Tract Infection
- If there is clinical chorioamnionitis or generalised sepsis associated with preterm labour, blood cultures, a urine specimen and vaginal swabs shall be taken and broad spectrum intravenous antibiotics shall be commenced. See Pharmacy and Medication guidelines

Tocolytic therapy

- The decision to suppress labour with tocolytic medication shall only be made by a registrar, senior registrar or the consultant on call.
- Any decision to change tocolytic medication shall only be made by the consultant.

1. First line: Nifedipine

- Unless contra-indicated, the first line tocolytic to be used shall be Nifedipine.
- Contraindications to the use of nifedipine include:
  - Contraindications to any suppression of labour including antepartum haemorrhage, pre-eclampsia, chorioamnionitis and fetal distress
  - Cardiac disease including cardiac conduction defects and left ventricular failure
  - Hypotension
  - Concomitant use of betamimetics such as Salbutamol
  - Caution should be taken with simultaneous administration of Magnesium Sulphate (MgSO4). This is not an absolute contraindication but care must be taken since hypotension may result.

Dosage

- Give an initial dose of 20mg of Nifedipine orally (not slow release Nifedipine)
- After 30 minutes, if contractions persist, give another 20mg oral dose
- After a further 30 minutes, if still contracting, follow up with a further 20mg orally
- If BP is stable, a maintenance dose of 20mg three times a day for 48-72 hours may be given where indicated.

  Note: The maximum dose of Nifedipine is 120mg / day.

Precautions

- IV shall be inserted and baseline electrolytes, urea and creatinine and LFT levels measured
• Half hourly maternal pulse, BP and respiratory rate until the contractions cease. Maternal hypotension should be treated with IV fluids in the first instance.
• Continuous electronic fetal heart rate monitoring until contractions have settled.
• Cardiovascular examination including auscultation of lung bases every 8 hours for first 24 hours of therapy.

Action
Onset of tocolysis is at 30-60 minutes and institution of second line tocolysis should not be considered in the first 2 hours. If contractions do not abate after this time a second line tocolytic may be considered by the consultant on call.

Side effects
• Facial flushing
• Headache
• Nausea
• Tachycardia
• Dizziness
• Hypotension – this is unusual in normotensive patients
• Cardiac failure
• Increase in liver enzymes

Contraindications
• Cardiac disease including cardiac conduction defects and left ventricular failure.
• Hypotension.

Other tocolytics which may be used in the event of the failure of Nifedipine tocolysis

2. Salbutamol
• Salbutamol may be used as a second line tocolytic, in the absence of contraindications.
• It must not be used in addition to Nifedipine, as the two drugs have potentially synergistic actions.
• Salbutamol is contra-indicated in the presence of
  ➢ Maternal or fetal cardiac disease
  ➢ Insulin dependent diabetes
  ➢ Thyroid disease
• Salbutamol should be used with care, as it is associated with maternal tachycardia, hypotension, tremor, pulmonary oedema, hyperglycaemia and hypokalaemia.

Dosage
• If Salbutamol is to be used for tocolysis, 5mg (5mL ampoule Ventolin Obstetric Injection) is added to 100mL of normal saline to produce a 50 mcg/mL solution.
• An IV infusion pump must be used for administration.
• Following the establishment of intravenous access, the salbutamol infusion is commenced at 12 mL/hour (10 mcg/minute) and increased by 4 mL/hour (3.3 mcg/minute) every 30 minutes until:
  • Contraction cease
  • Maternal pulse rate reaches 120 beats/minute or
  • The infusion rate reaches a maximum of 36 mL/hour (30 mcg/minute).

**Precautions**

• Baseline electrolytes, urea and creatinine before commencement of infusion; repeat as necessary if abnormal.
• Baseline maternal blood sugar level; repeat 4 hourly if abnormal
• Cardiovascular examination including auscultation of lung bases every 8 hours
• No additional intravenous fluids to avoid fluid overload
• Half hourly maternal pulse, BP and respiratory rate until the maintenance dose is reached
• Reduce the infusion if the maternal pulse >120 bpm
• **CEASE** the infusion and request medical review immediately if there is chest pain, dyspnoea or the respiratory rate >30/min
• Baseline electronic fetal heart rate monitoring
• Do not exceed 48 hours of salbutamol therapy. Only in exceptional circumstances should the treatment be continued for more than 24 hours.

**Side effects**

• Tachycardia
• Tremor
• Nausea
• Dizziness
• Hypotension
• Pulmonary oedema and cardiac failure
• Hypokalaemia

**Contraindications**

• Maternal or fetal cardiac disease
• Thyroid disease
• Insulin dependent diabetes
3. Glyceryl Trinitrate (GTN)

Glyceryl Trinitrate (GTN) is a nitric oxide donor and causes smooth muscle relaxation via the metabolite nitric oxide (NO) which acts as a 2nd messenger to increase Ca\(^{2+}\) uptake. Nitric oxide promotes uterine quiescence in pregnancy; current evidence does not support the routine administration of nitric oxide donors in the treatment of threatened preterm labour\(^5\). Peak action occurs 1-2 hours after application. It acts as a vasodilator. GTN patches provide continuous plasma nitrate concentration up to 24 hours.

**Dosage**
- Apply a 5-10mg transdermal GTN patch to abdominal skin, and repeat the dose in 1 hour if the contractions persist (maximum dose 20mg in 24 hours).

**Side effects**
- Headache
- Facial flushing
- Hypotension and Tachycardia

4. Indomethacin

- Indomethacin use may be indicated in association with the insertion of a cervical suture at pre-viable gestations.
- Indomethacin, short term, may be considered when there is a failure or a contraindication to other tocolytics. Theoretical risks of fetal pulmonary hypertension and reduced renal function are debatable in short-term use but clear in the event of extended use \(^6,7\).

**Dosage**
- Indomethacin is administered as a 100 mg rectal suppository followed by a 25 mg oral dose every 4 hours for 48 hours.
- If regular uterine contractions persist 1-2 hours after the initial 100 mg suppository an additional 100 mg suppository is administered before beginning oral therapy.

**Side effects**
- Prolonged use of indomethacin, especially in the presence of a relatively mature fetus, may lead to narrowing or occlusion of the fetal ductus arteriosus and/or reduction in fetal renal function.
- As there are no evidence-based indications to continue tocolysis for greater than 48 hours, or at gestations greater than 34 weeks, these potential side effects do not contraindicate the use of indomethacin if other regimens are not safely available (e.g. in the absence of the ability to adequately monitor the possibility of side effects of other regimens and in transport situations).

**Contraindication**
- Peptic ulceration.
Preterm labour- Subsequent care of the woman on the antenatal ward

* Full set of observations includes Blood Pressure, Temperature, Respiration and SA02

**Maternal assessment**

- **On Admission**
  - Full set of observations*, Urinalysis, Vaginal loss, uterine activity/ tenderness, pain or rigidity

- **First 24 hours**
  - 4 hourly- Full set of observations*, Vaginal loss, uterine activity/ tenderness, pain or rigidity

- **2nd day**
  - 4 hourly Vaginal loss, uterine activity/ tenderness, pain or rigidity
  - 8 hourly – Full set of observations*
  - 8 hourly Full set of observations* and Bowel activity

- **3rd day**
  - 4 hourly Vaginal loss, uterine activity/ tenderness, pain or rigidity

- **Avoid digital vaginal examination: women with ongoing painful regular contractions should be transferred to LBS for review**

- **Overnight**
  - Observe and perform observations only as required between 22:00 and 06:00

**Fetal assessment**

- **On Admission**
  - 4 hourly FHR and Fetal Movement

- **2nd day and thereafter**
  - 4 hourly FM
  - Twice Daily- FHR
  - Report abnormalities promptly

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*Abbreviations: B.P. = Blood Pressure, T = Temperature, R = Respiration, S.A.02 = SaO2, LBS = Labour Delivery Special Care Baby Unit*
Preterm labour

Immediately if bleeding/contracting (regular, frequent, painful) or c/o pain/tenderness, requesting analgesia, > 30 weeks gestation

Procedures to be considered

- Tocolytic Therapy
- Corticosteroids
- Maternal laboratory investigations:
  - Ultrasound to exclude Low Lying Placenta
  - Swabs: HVS/ LVS for M.C&S, Fetal Fibronectin Test.

Education

Plan of care, tests and procedures
- Preterm birth
- Special Care Nursery
- Breastfeeding

Activity

- Rest in bed with toilet privileges- 48 hours
- Day 3 and thereafter - Gentle ambulation

Documentation

- MR 285 - Observation Chart
- MR 810 - Medication Chart
- MR 250 - Progress Notes
- MR 410 - Neonatal History Sheet
- Baby notes prepared
- Perinatal Database record (STORK) updated

Referrals to be considered

- Neonatologist
- Aboriginal Liaison Officer
- Activities Coordinator
- Parent Educator
- Physiotherapist
- Social Worker
- Psychological Medicine
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<th>Keywords:</th>
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