Management of abnormal cytology in pregnancy

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Colposcopy in pregnancy and in the postpartum period

Primary aim of colposcopy for pregnant women is to exclude invasive disease. Otherwise defer biopsy and treatment until delivery.

Women who have LG cytology and in whom colposcopy excludes HG disease, then repeat colposcopy/ cytology test 3-4 months after delivery.

Women with HG disease in whom colposcopy has excluded suspicion of invasive disease, review 3 monthly with final assessment 3-4 months following delivery. Decision should be made on whether treatment required.

Risk of progression of CIN3 low in pregnancy (up to 0.4%; Paraskevaidis et al, 2002).
However...

If pregnant woman requires colposcopy/ cytology after treatment (or follow-up of untreated CIN1), assessment may be delayed until after delivery.

Unless there is obstetric contraindication, assessment should not be delayed if 1\textsuperscript{st} appointment for follow-up cytology/ colposcopy due following treatment for cGIN. ‘Test of cure’ appointment should not be delayed after treatment for CIN2-3 with involved/ uncertain margin status.
Colposcopy in pregnancy proceeds in the same way as in non-pregnant woman. Colposcopic assessment of pregnant cervix more difficult than in non-pregnant cervix because:

- *cervix larger, oedematous/ more vascular*

- *ordinary cervical speculum may make access to cervix difficult, in which case larger speculum should be used*

- *cervix usually covered by mucus, which is difficult to remove*

- *in primiparous patient in particular there may be immature metaplasia, which can be confusing*

- *and... decidual changes of cervical epithelium can mimic cancerous epithelium.*
In addition, vascular changes associated with abnormality may be more pronounced leaving inexperienced colposcopist to conclude that severity of lesion may be more than it actually is.

A pregnant woman with abnormal cervical smear should be assessed by experienced colposcopist.

If malignancy suggested by cytologic/colposcopic assessment then colposcopically directed biopsy/biopsies should be taken, but in absence of these features biopsy should be postponed until after pregnancy.
PHE Standards

Cervical screening during pregnancy

A woman referred with abnormal cytology should undergo colposcopy in late 1st/ early 2nd trimester unless clinical contraindication, however, for LG changes triaged to colposcopy with +ve HPV test, assessment may be delayed until after delivery.

If previous colposcopy abnormal and in interim the woman becomes pregnant, then repeat colposcopy should not be delayed.
May wish to perform colposcopy only @ follow-up appointment scheduled during pregnancy.

If repeat cytology due, and woman has missed/ defaulted appointment prior to pregnancy, cytology/ colposcopy during pregnancy can be considered.

But maybe

• 5.4% progression rate to cancer in pregnancy (Siegler et al, 2017) if have biopsy proven CIN2/3 in early pregnancy. No research published regarding morbidity in pregnancy relating to 1st trimester and that is when to assess with guided biopsy if possible.

• Biopsy appears safe in 1st 15/40 pregnancy.

Siegler et al, 2017
Management algorithm following initial colposcopy...

If ≤CIN1 suspected, repeat examination 3 months following delivery.

If CIN2/3 suspected, repeat colposcopy @ end of 2nd trimester. If pregnancy already advanced beyond that point, repeat 3 months following delivery.

If invasive disease suspected clinically/colposcopically, biopsy adequate to make diagnosis essential. Cone, wedge, and diathermy loop biopsies all associated haemorrhage and such biopsies should be taken only where appropriate facilities to deal with haemorrhage available.

Punch biopsy suggesting CIN only cannot reliably exclude invasion.
**Evidence**: case series of biopsies taken by diathermy loop in pregnancy have shown risk of haemorrhage approx. 25% (Robinson et al, 1997). Case series of women with LG disease confirm safety of deferring further follow-up until postpartum period (International Collaboration of Epidemiological Studies of Cervical Cancer, 2007; Wetta et al, 2009).

Rate of CIN2+ following LSIL/ ASC-US 3.7% in pregnancy (Dunn et al, 2001)
Colposcopic differentiation between decidual ectopy and carcinoma

1. Location: decidual ectopy well demarcated, situated within typical squamous/ glandular epithelium, carcinoma situated mainly within atypical TZ

2. Form: exophytic growth of decidual ectopy usually multifocal, in carcinoma growth unifocal.

3. Vasculature: in decidual ectopy characterized by smaller vessels/ smaller inter-vessel spaces as compared to carcinoma. In decidual ectopy, vessels do not disappear in acetic acid test.
Nodular decidual ectopy
Pregnancy 16th week

6 weeks after delivery
Vasculature in decidual ectopy compared to carcinoma characterized by smaller vessels/smaller inter-vessel spaces
In decidual ectopy, vessels do not disappear in acetic acid test
Pregnancy 13th week
Pregnancy 22nd week
Pregnancy 36\textsuperscript{th} week
Colposcopy – suspected invasion
9 weeks after delivery
Colposcopy – same case: total remission
Cervical condyloma in pregnancy
Colposcopy follow-up after pregnancy

If colposcopy performed during pregnancy, post-partum assessment of women with abnormal cytology/biopsy-proven CIN essential. Excision biopsy in pregnancy cannot be considered therapeutic.

In immediate puerperium, cervix may also be difficult to assess. Prior to 1st postnatal ovulation, particularly if breastfeeding, cervix may appear atrophic which makes both cytology/colposcopy more difficult.

If this a problem for diagnosis, then short course of vaginal oestrogen helpful. When no suspicion of malignancy, wait until oestrogenic state returned to normal before undertaking colposcopy and/or treatment.
Evidence: low regression rates for pre-invasive cervical disease during pregnancy/ following delivery from retrospective uncontrolled studies/ regression not related to mode of delivery (Ackermann et al, 2006; Everson et al, 2002). Retrospective study of pregnant women treated by cone biopsy for HG CIN/ microinvasion reported high rates of disease persistence (Lapolla et al, 1988).

Ackermann et al. 2006; Everson et al. 2002; Lapolla et al. 1988.
Case

• 27yr
• Ref cytology HG dyskaryosis (favour severe) or LIEHG
• No previous colposcopy
• P4
• Moderate smoker
• Seen colp 12/40 pregnant
• Seen colp 29/40 pregnant
• Del 21.01.10 - 3.94kg female LSCS
• Seen in colp 4 months later
• Loop 20 x 15 x 10mm
Quit smoking!
Algorithm
Colposcopy for pregnant women

Woman (or her GP) contacts colposcopy clinic to inform of pregnancy

Previously attended clinic

Awaiting treatment for CIN 2+
Should attend clinic at 28 weeks of pregnancy for assessment**

Due colposcopic follow-up after treatment for CGIN or for untreated suspected HG abnormality*
Should attend clinic at 28 weeks of pregnancy for assessment

Low grade smears/biopsies/colposcopy only
Colposcopist to review notes – may see at 3/12 postpartum or discharge for follow-up cytology

Direct referral and never attended
Encourage to attend as soon as possible

*if has been treated for CIN can be discharged for smear with HPV ToC at 3 months post partum
**if over 28 weeks, should attend as soon as possible

implemented 29.09.2014
Algorithm

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Encourage to attend now

Direct referral and never attended

What if cytology severe ? invasion or colp suspicious of microinvasion?

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implemented 29.09.2014
Algorithm 9
Colposcopy for pregnant women

Woman (or her GP) contacts colposcopy clinic to inform of pregnancy

Previously attended clinic

Awaiting treatment for CIN 2+

Should attend clinic at 28 weeks of pregnancy for assessment**

Done colposcopic assessment after treatment

Should attend clinic at 28 weeks of pregnancy for assessment

Low grade smears/biopsies/
colposcopy only

Colposcopist to review notes – may see at 3/12 postpartum or discharge for follow-up cytology

Direct referral and never attended

Encourage to attend

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implemented 29.09.2014
• Biopsy acceptable for HSIL.

• In absence of invasive disease/advanced pregnancy, additional colposcopic/cytologic examinations acceptable in pregnant women.

• Colposcopy repeated for CIN 2-3 at intervals no more frequent than every 12 weeks. Repeat biopsy recommended only if appearance of lesion worsens/ if cytology suggests invasive cancer. Deferring re-evaluation until at least 6 weeks postpartum acceptable.

• Diagnostic excisional procedure recommended only if invasion suspected. Unless invasive cancer identified, treatment is unacceptable.

• Re-evaluation with cytology/colposcopy recommended no sooner than 6 weeks postpartum (Massad et al, 2013).
Conclusion

• Colposcopy is difficult to perform in pregnancy
• It is uncomfortable in pregnancy. Natural history of CIN means that delay is possible for the majority of cases. Be careful to identify cancer in pregnancy.
• Biopsies can be taken in pregnancy.
• Assessment by an experienced colposcopist is important.