# National Institute for Health and Care Excellence

Final

# Intrapartum care

# **GRADE** tables for review M: Uterotonics for the prevention of postpartum haemorrhage

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Final



FINAL

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# **GRADE** tables

# F1 – GRADE tables for postpartum haemorrhage ≥1000mL (pairwise analysis)

Table 1: Carboprost versus Misoprostol ≤600mcg

			Quality asse	ssment			No of patients Effect			Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Carboprost	Misoprostol ≤600mcg	Relative (95% Cl)	Absolute	Quality	Importanc
PPH >1000	PH >1000 mL - Vaginal birth											
1 (Nellore 2006)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/60 (0%)	0/60 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>3</sup>	VERY LOW	CRITICAL
01		DDU	the sufficient last successful									

CI: confidence interval; PPH: postpartum haemorrhage

1 Unclear risk of bias in randomisation; allocation concealment; blinding; incomplete outcome data; selective reporting.

2 Sample size <200

3 Calculated from risk difference

#### Table 2: Ergometrine versus Misoprostol ≤600mcg

			Quality assessm	ent			No of patients Effect					
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine	Misoprostol ≤600mcg	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000 mL - Vagina	al birth											
4 (Chhabra 2008; Humera 2016; Jago 2007; Vimala 2004)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/464 (0%)	0/566 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	LOW	CRITICAL

CI: confidence interval; PPH: postpartum haemorrhage

1 Unclear risk of bias for blinding; incomplete outcome data; selective reporting.

#### Table 3: Ergometrine versus Oxytocin >5 iu to ≤ 10 iu

			Quality as	sessment			No of patients Effect			Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine	Oxytocin >5 iu to ≤ 10 iu	Relative (95% CI)	Absolute	Quality	Importance
PPH >100	0 mL - Vagina	l birth										
1 (Orji 2008)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/303 (0%)	0/297 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	CRITICAL
Cli confin	lanaa intanyal	. DDU	atratum haama	rrhaga								

*CI: confidence interval; PPH: postpartum haemorrhage* 

1 Unclear risk of bias for randomisation; blinding; incomplete outcome data

2 Calculated from risk difference

#### Table 4: Misoprostol + Oxytocin versus Oxytocin >10 iu

			Quality asses	ssment			No of pat	ients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin > 10iu	Relative (95% CI)	Absolute	Quality Importar	Importance
PPH >1000 r	nL - Caesarea	n birth										
1 (Adanikin randomised serious <sup>1</sup> no serious inconsistency indirectness       no serious <sup>2</sup> none       0/109 (0/109)(0/109 (0/109 (0/109 (0/109 (0/109 (0/109 (0/109 (0/10												CRITICAL
CI: confider	nce interval; F	PH: post	partum haemorrha	age								

1 Unclear risk of bias for selective reporting 2 Sample size 200-400

3 Calculated from risk difference

#### Table 5: Misoprostol + Oxytocin versus Oxytocin >5 iu to ≤ 10 iu

			Quality assess	ment			No of pa	atients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000 mL - Caesarean birth												

			Quality assess	sment			No of patients Effect					
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
1 (Elsedeek 2012)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/200 (0%)	0/200 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	CRITICAL

CI: confidence interval; PPH: postpartum haemorrhage

1 Sample size 200-400 2 Calculated from risk difference

#### Table 6: Misoprostol >600 mcg to ≤800 mcg versus Oxytocin >5 iu to ≤ 10 iu

			Quality ass	sessment			No of patients Effect			Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol >600 mcg to ≤800 mcg	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000	mL - Vaginal	birth										
1 (Parsons 2006)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/225 (0%)	0/225 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	CRITICAL

CI: confidence interval; PPH: postpartum haemorrhage 1 Unclear risk of bias for randomisation, blinding and selective reporting 2 Calculated from risk difference

#### Table 7: Misoprostol >600 mcg to ≤800 mcg versus Oxytocin >1 iu to ≤ 5 iu

			Quality ass	essment			No of patients Effect					
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol >600 mcg to ≤800 mcg	Oxytocin >1 iu to ≤ 5 iu	Relative (95% Cl)	Absolute	Quality	Importance
PPH >100	0 mL - Vagina	al birth										
1 (Nasr 2009)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/257 (0%)	0/257 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>1</sup>	HIGH	CRITICAL

CI: confidence interval; PPH: postpartum haemorrhage

1 Calculated from risk difference

#### Table 8: Misoprostol ≤600mcg versus Oxytocin >5 iu to ≤ 10 iu

		Qua	ality assessment			No of patients Effect			Effect			
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol ≤600	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000 mL - Vaginal birt	h											
8 (Afolabi 2010; Bellad 2012; Bhatti 2014; Gupta 2006; Oboro 2003; Sadiq 2011; Tewatia 2014; Walley 2000)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/1980 (0%)	0/1986 (0%)	Not estimable	0 fewer per 1000 (from 0 fewer to 0 more) <sup>2</sup>	MODERATE	CRITICAL

1 Unclear risk of bias for allocation concealment, blinding and selective reporting.

2 Calculated from risk difference

#### Table 9: Ergometrine + Oxytocin versus Oxytocin >10 iu

			Quality assess	sment			No of pati	ents		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine + Oxytocin	Oxytocin >10iu	Relative (95% CI)	Absolute	Quality	Importance
PPH >1000	mL - vaginal b	oirth										
1 (Nuamsiri 2016)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/162 (0%)	0/161 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	CRITICAL

*CI: confidence interval; PPH: postpartum haemorrhage* 1 Sample size 200-400

#### Table 10: Oxytocin >10 iu versus Carbetocin

			Quality asses	sment			No of p	atients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >10iu	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000 r	nL - Caesarear	n birth										
1 (Boucher randomised serious <sup>1</sup> no serious no serious no serious very none 0/28 0/29 Not 0 fewer per 1000 (from 70 1998) rindirectness serious <sup>2</sup>								VERY LOW	CRITICAL			
Cl: confider	an interval: D	DU: naata	ortum hoomorrhod									

*CI: confidence interval; PPH: postpartum haemorrhage* 

1 Unclear risk of bias for randomisation, allocation concealment and selective reporting

2 Sample size <200

3 Calculated from risk difference

#### Table 11: Oxytocin >5 iu to ≤ 10 iu versus Carbetocin

			Quality assess	ment			No of p	atients		Effect			
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin 5- 10	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance	
PPH >1000 mL - Vaginal birth													
1 (Fenix 2012)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	0/30 (0%)	0/30 (0%)	Not estimable	0 fewer per 1000 (from 60 fewer to 60 more) <sup>2</sup>	LOW	CRITICAL	
Cl. and field	into m coli	DDI I											

*CI: confidence interval; PPH: postpartum haemorrhage* 1 Sample size <200

2 Calculated from risk difference

#### Table 12: Oxytocin >1 iu to ≤ 5 iu versus Carbetocin

		C	Quality assessme	ent			No of pat	tients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu	Carbetocin	Relative (95% CI)	Absolute	Quality	Importance
PPH >1000 mL												

		(	Quality assessme	ent			No of pat	tients		Effect			
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu	Carbetocin	Relative (95% CI)	Absolute	Quality	Importance	
2 (Amornpetchakul 2018; Rosseland 2013)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/200 (0%)	0/201 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more)	HIGH	CRITICAL	
PPH >1000 mL - Vaginal Birth													
1 (Amornpetchakul 2018)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/174 (0%)	0/176 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	CRITICAL	
PPH >1000 mL - Caes	arean Birth												
PPH >1000 mL - Caesarean Birth         1 (Rosseland 2013)       randomised trials       no serious no serious indirectness       very serious <sup>3</sup> none       0/26 (0%)       0/25 (0%)       Not estimable (from 70 fewer to LO 70 more) <sup>2</sup>										LOW	CRITICAL		
CI: confidence interv 1 Sample size 200-4 2 Calculated from ris 3 Sample size <200	val; PPH: po 100 sk difference	stpartum h	aemorrhage										

#### Table 13: Oxytocin >1 iu to $\leq$ 5 iu versus Placebo

			Quality assessm	ent			No of patient	s		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu versus Placebo	Control	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000mL												
2 (Jerbi 2007; Rosseland 2013)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/91 (0%)	0/90 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>3</sup>	VERY LOW	CRITICAL
PPH >1000mL - \	/aginal Birth											
1 (Jerbi 2007)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/65 (0%)	0/65 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>3</sup>	VERY LOW	CRITICAL
PPH >1000mL - C	aesarean Bir	th										
1 (Rosseland 2013)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/26 (0%)	0/25 (0%)	Not estimable	0 fewer per 1000 (from 70 fewer to 70 more) <sup>3</sup>	LOW	CRITICAL

CI: confidence interval; PPH: postpartum haemorrhage 1 Unclear risk of bias for randomisation, allocation concealment, blinding, incomplete outcome data and selective reporting.

2 Sample size <200

3 Calculated from risk difference

#### Table 14: Carbetocin versus Placebo

			Quality assessm	ient			No of pa	tients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Carbetocin	Placebo	Relative (95% Cl)	Absolute	Quality	Importance
PPH >1000mL	- Caesarean b	irth										
1 (Rosseland 2013)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	0/25 (0%)	0/25 (0%)	Not estimable	0 fewer per 1000 (from 70 fewer to 70 more) <sup>2</sup>	LOW	CRITICAL

CI: confidence interval; PPH: postpartum haemorrhage 1 Sample size <200 2 Calculated from risk difference

# F2 – GRADE tables for severe maternal morbidity – intensive care admission

			Quality asses	sment			No of pat	ients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >10 iu	Relative (95% Cl)	Absolute	Quality	Importance
Severe maternal morbidity - intensive care admissions - Caesarean birth												
1 (Ugwu 2014)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	0/60 (0%)	0/60 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>2</sup>	LOW	IMPORTANT
CI: confide 1 Sample	ence interval size <200											

#### Table 15: Misoprostol + Oxytocin versus Oxytocin >10 iu

2 Calculated from risk difference

#### Table 16: Misoprostol + Oxytocin versus Oxytocin >5 iu to ≤ 10 iu

			Quality asses	sment			No of pa	atients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
Severe mat	ernal morbid	ity - intensiv	ve care admission	s - Caesarean bi	rth							
1 (El Tahan 2012)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/179 (0%)	0/187 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	

CI: confidence interval

1 Sample size 200-400

#### Table 17: Misoprostol >800 mcg to ≤1000 mcg versus Oxytocin >5 iu to ≤ 10 iu

			Quality asses	ssment			No of patie	nts		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprosotol >800 mcg to ≤1000 mcg	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
Severe mate	ernal morbidit	y - intens	ive care admissio	ns - Vaginal birt	h							
1 (Shrestha 2011)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	serious <sup>2</sup>	none	0/100 (0%)	0/100 (0%)	Not estimable	0 fewer per 1000 (from 20 fewer to 20 more) <sup>3</sup>	LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, and selective reporting

2 Sample size 200-400

3 Calculated from risk difference

#### Table 18: Misoprostol >600 mcg to ≤800mcg versus Oxytocin 10 iu

			Quality assess	ment			No of patier	nts		Effect				
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol >600mcg to ≤800 mcg	Oxytocin 10iu	Relative (95% Cl)	Absolute	Quality	Importance		
Severe mater	vere maternal morbidity - intensive care admissions - Caesarean birth													
1 (Chaudhuri 2010)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious¹	none	0/96 (0%)	0/94 (0%)	Not estimable	0 fewer per 1000 (from 20 fewer to 20 more) <sup>2</sup>	LOW	IMPORTANT		

CI: confidence interval

1 Sample size <200

#### Table 19: Misoprostol >600 mcg to $\leq$ 800 mcg versus Oxytocin >1 iu to $\leq$ 5 iu

			Quality asso	essment			No of pati	ents		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol >600mcg to ≤800 mcg	Oxytocin >1 iu to ≤ 5 iu	Relative (95% CI)	Absolute	Quality	Importance
Severe mate	ernal morbidit	y - intens	ive care admissio	ons - Vaginal bir	th							
2 (Amin 2014; Nasr 2009)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/357 (0%)	0/357 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for randomisation, allocation concealment, blinding, incomplete outcome data. 2 Calculated from risk difference

#### Table 20: Misoprostol ≤600 mcg versus Oxytocin >5 iu to ≤ 10 iu

		c	Quality assessme	ent		No of p	oatients		Effect				
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol 600	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance	
Severe maternal morbidity - intensive care admissions - Vaginal birth													
4 (Afolabi 2010; Kundodyiwa 2001; Musa 2015; Tewatia 2014)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/493 (0%)	0/506 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	CRITICAL	

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, selective reporting.

#### Table 21: Misoprostol ≤600 mcg versus Carbetocin

			Quality asse	ssment			No of pat	ients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol ≤600mcg	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance
Severe ma	ternal morbid	ity - intens	sive care admissio	n - Vaginal birth								
1 (Ibrahim 2017)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	2/30 (6.7%)	0/30 (0%)	POR 7.65 (0.47 to 125.22)	70 more per 1000 (from 40 fewer to 170 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval; POR: Peto odds ratio

1 Unclear risk of bias for blinding, allocation concealment, incomplete outcome data.

2 95% CI crosses 2 MIDs

3 Calculated from risk difference

#### Table 22: Ergometrine + Oxytocin versus Carbetocin

			Quality assessr	nent			No of patients			Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine + Oxytocin	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance
Severe materna	al morbidity -	intensive ca	are admissions - \	/aginal birth								
2 (Nirmala 2009; Samimi 2013)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/160 (0%)	0/160 (0%)	Not estimable	0 fewer per 1000 (from 20 fewer to 20 more) <sup>2</sup>	MODERATE	IMPORTANT

CI: confidence interval

1 Sample size 200-400

#### Table 23: Oxytocin >5 iu to ≤ 10 iu versus Placebo

			Quality asses	sment			No of patie	nts		Effect			
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >5 iu to ≤ 10 iu	oxytocin >5 iu to ≤ 10 iu Placebo		Absolute	Quality	Importance	
Severe mater	vere maternal morbidity - intensive care admissions - Vaginal birth												
1 (Abdel- Aleem 2010)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/1291 (0%)	0/659 (0%)	Not estimable	0 per 1000 (from 0 fewer to 0 more) <sup>1</sup>	HIGH	IMPORTANT	
CI: confidence	ce interval												

1 Calculated from risk difference

#### Table 24: Carbetocin versus Oxytocin >1 iu to $\leq$ 5 iu

			Quality assess	sment			No of	patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Carbetocin	Oxytocin >1 iu to ≤ 5 iu	Relative (95% Cl)	Absolute	Quality	Importance
Severe mat	evere maternal morbidity - intensive care admissions - Caesarean birth											
1 (Attilakos 2010)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	1/188 (0.53%)	0/189 0%	POR 7.43 (0.15 to 374.38)	10 more per 1000 (from 10 fewer to 20 more) <sup>2</sup>	LOW	IMPORTANT
Cl. as official	n a a interrute le		dele vetie									

CI: confidence interval; POR: Peto odds ratio

1 95% CI crosses 2 MIDs

### **F3 – GRADE** tables for need for additional uterotonics

#### Table 25: Misoprostol + Oxytocin versus Oxytocin >10 IU

			Quality asses	sment		No of patients			Effect			
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >10 iu	Relative (95% Cl)	Absolute	Quality	Importance
Need for a	dditional utero	otonics - Caes	sarean birth									
1 (Lapaire 2006)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	0/28 (0%)	0/25 (0%)	Not estimable	0 fewer per 1000 (from 70 fewer to 70 more) <sup>2</sup>	LOW	IMPORTANT
CI: confidence interval 1 Sample size <200 2 Calculated from risk difference												

#### Table 26: Oxytocin >1 iu to ≤ 5 iu versus Carbetocin

			Quality asse	ssment			No of pat	ients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance
Need for a	leed for additional uterotonics - Caesarean birth											
1 (Moertl 2011)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/28 (0%)	0/28 (0%)	Not estimable	0 fewer per 1000 (from 70 fewer to 70 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment.

2 Sample size <200

## F4 – GRADE tables for need for blood transfusion

#### Table 27: Carboprost versus Ergometrine

			Quality asse	ssment			No of	patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Carboprost	Ergometrine	Relative (95% Cl)	Absolute	Quality	Importance
Need for b	lood transfusi	ion - Vagin	al birth									
1 (Supe 2016)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious²	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete data, and selective reporting

2 Sample size <200

3 Calculated from risk difference

#### Table 28: Carboprost versus Misoprostol >600mcg to ≤800mcg

			Quality asso	essment			No	o of patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Carboprost	Misoprostol >600mcg -to ≤800 mcg	Relative (95% CI)	Absolute	Quality	Importance
Need for I	blood transfus	sion - Vagi	inal birth									
1 (Supe 2016)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete data, and selective reporting

2 Sample size <200

3 Calculated from risk difference

#### Table 29: Carboprost versus Placebo

Quality assessment No of patients	Effect	Quality Importance
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No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Carboprost	Placebo	Relative (95% CI)	Absolute		
Need for blood transfusion - Vaginal birth												
1 (Supe 2016)	randomised trials	very serious¹	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete data, and selective reporting

2 Sample size <200

3 Calculated from risk difference

#### Table 30: Ergometrine versus Misoprostol >600mcg to ≤800mcg

			Quality asse	essment			No	of patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine	Misoprostol >600 mcg to ≤800 mcg	Relative (95% CI)	Absolute	Quality	Importance
Need for I	blood transfus	sion - Vag	inal birth									
1 (Supe 2016)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete data, and selective reporting

2 Sample size <200 3 Calculated from risk difference

#### Table 31: Ergometrine versus Misoprostol ≤600 mcg

			Quality assessm	nent			No of	patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine	Misoprostol ≤600 mcg	Relative (95% CI)	Absolute	Quality	Importance
Need for blood transf	fusion - Vagi	nal birth										
4 (Chhabra 2008; Humera 2016; Otoide 2020; Vimala 2004)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/360 (0%)	0/460 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete outcome data and selective reporting

2 Calculated from risk difference

#### Table 32: Ergometrine versus Placebo

			Quality asse	ssment			No of pat	ients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine	Placebo	Relative (95% Cl)	Absolute	Quality	Importance
Need for b	lood transfusi	on - Vagina	al birth									
1 (Supe 2016)	randomised trials	very serious¹	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete data, and selective reporting

2 Sample size <200

#### Table 33: Misoprostol + Oxytocin versus Oxytocin >10 IU

			Quality assoc	smont			No of pati	ionto		Effect		
			Quality asses	Sillent			NO OI pati	ents		Enect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >10iu	Relative (95% CI)	Absolute	Quality	Importance
Need for bl	ood transfusi	on - Caesarea	an birth									
1 (Lapaire 2006)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	0/28 (0%)	0/25 (0%)	Not estimable	0 fewer per 1000 (from 70 fewer to 70 more) <sup>2</sup>	LOW	IMPORTANT
CI: confide	nce interval											

1 Sample size <200

2 Calculated from risk difference

#### Table 34: Misoprostol + Oxytocin versus Oxytocin >5 iu to ≤ 10 iu

			Quality assess	sment			No of pa	atients		Effect	Quality	Importance
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol + Oxytocin	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute		
Need for blo	ood transfusi	on - Caesare	an birth									
1 (Elsedeek 2012)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/200 (0%)	0/200 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	IMPORTANT
CI: confider	nce interval											

CI: confidence interval

1 Sample size 200-400

#### Table 35: Misoprostol >600 mcg to ≤800mcg vs Placebo

			Quality asse	essment			No of patient	S		Effect	Quality	Importance
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol >600 mcg to ≤800	Placebo	Relative (95% Cl)	Absolute		
Need for b	lood transfus	ion - Vagir	nal birth									
1 (Supe 2016)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete data, and selective reporting

2 Sample size <200

3 Calculated from risk difference

#### Table 36: Misoprostol ≤600 mcg versus Ergometrine + Oxytocin

			Quality assess	ment			No of j	patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol ≤600 mcg	Ergoemtrine + Oxytocin	Relative (95% CI)	Absolute	Quality	Importance
Need for blood tra	nsfusion - Va	ginal birt	h									
2 (Bamigboye, Merrell 1998; Harriott 2009)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/301 (0%)	0/303 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, selective reporting

#### Table 37: Misoprostol ≤600 mcg versus Oxytocin >5 iu to ≤ 10 iu

		Qua	ality assessment	t			No of pa	itients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol ≤600 mcg	Oxytocin >5 iu to ≤ 10 iu	Relative (95% CI)	Absolute	Quality	Importance
Need for blood transfusion												
7 (Afolabi 2010; Fazel 2013; Gupta 2006; Lumbiganon 1999; Oboro 2003; Sadiq 2011; Tewatia 2014)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/1844 (0%)	0/1633 (0%)	Not estimable	0 fewer per 1000 (from 0 fewer to 0 more) <sup>2</sup>	MODERATE	IMPORTANT
Need for blood transfusion	- Vaginal bir	th										
6 (Afolabi 2010; Gupta 2006; Lumbiganon 1999; Oboro 2003; Sadiq 2011; Tewatia 2014	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/1794 (0%)	0/1583 (0%)	Not estimable	0 fewer per 1000 (from 0 fewer to 0 more) <sup>2</sup>	MODERATE	IMPORTANT
Need for blood transfusion	- Caesarean	birth										
1 (Fazel 2013)	randomised trials	very serious <sup>3</sup>	no serious inconsistency	no serious indirectness	very serious <sup>4</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>2</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, selective reporting

2 Calculated from risk difference

3 Unclear risk of bias for allocation concealment, blinding, incomplete outcome data, selective reporting

4 Sample size <200

#### Table 38: Misoprostol $\leq$ 600 mcg versus Oxytocin >1 iu to $\leq$ 5 iu

			Quality asses	sment			No of pa	atients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol ≤600 mcg	Oxytocin >1 iu to ≤ 5 iu	Relative (95% CI)	Absolute	Quality	Importance
Need for blood	transfusion	- Vaginal I	birth									
2 (Baskett 2007; Karkanis 2002)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	no serious imprecision	none	0/421 (0%)	0/424 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	IMPORTANT
CI: confidence	interval											

1 Unclear risk of bias for blinding, selective reporting

2 Calculated from risk difference

#### Table 39: Misoprostol ≤600 mcg versus Placebo

			Quality asse	ssment			No of patie	nts		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Misoprostol ≤600 mcg	Placebo	Relative (95% Cl)	Absolute	Quality	
Need for b	lood transfusi	on - Vagin	al birth									
1 (Zgaya 2020)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	serious <sup>2</sup>	none	0/111 (0%)	0/100 (0%)	Not estimable	0 fewer per 1000 (from 20 fewer to 20 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for allocation concealment, blinding, incomplete outcome data, selective reporting

2 Sample size 200-400

#### Table 40: Ergometrine + Oxytocin versus Oxytocin >10 iu

			Quality asses	sment			No of pati	ents		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Ergometrine + Oxytocin	Oxytocin >10 iu	Relative (95% CI)	Absolute	Quality	Importance
Need for <b>b</b>	olood transfus	sion - Caesare	an birth									
1 (Balki 2008)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	very serious <sup>1</sup>	none	0/24 (0%)	0/24 (0%)	Not estimable	0 fewer per 1000 (from 80 fewer to 80 more) <sup>2</sup>	LOW	IMPORTANT
CI: confid	ence interval											

1 Sample size <200

2 Calculated from risk difference

#### Table 41: Oxytocin >10 iu versus Oxytocin >5 iu to $\leq$ 10 iu

			Quality asse	ssment			No o	f patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >10 iu	Oxytocin >5 iu to ≤ 10 iu	Relative (95% Cl)	Absolute	Quality	Importance
Need for b	lood transfusi	on - Caes	arean birth									
1 (Fahmy 2015)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for blinding, incomplete outcome data, selective reporting

2 Sample size <200

#### Table 42: Oxytocin >10 iu versus Carbetocin

		Q	uality assessmen	t			No of p	atients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >10 iu	Carbetocin	Relative (95% CI)	Absolute	Quality	Importance
Need for blood transfus	ion											
3 (Boucher 1998; Fahmy 2015; Taheripanah 2017)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	serious <sup>2</sup>	none	0/188 (0%)	0/189 (0%)	Not estimable	0 fewer per 1000 (from 20 fewer to 20 more) <sup>3</sup>	VERY LOW	IMPORTANT
Need for blood transfus	ion - Vaginal I	birth										
1 (Fahmy 2015)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious⁴	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT
Need for blood transfus	ion - Caesare	an birth										
2 (Boucher 1998; Taheripanah 2017)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	serious <sup>2</sup>	none	0/138 (0%)	0/139 (0%)	Not estimable	0 fewer per 1000 (from 20 fewer to 20 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for randomisation, allocation concealment, blinding, selective reporting 2 Sample size 200-400

3 Calculated from risk difference

4 Sample size <200

#### Table 43: Oxytocin >5 iu to ≤ 10 iu versus Carbetocin

			Quality accoss	mont			No of pat	tionte		Effect		
			Quality assess				NO OI Pai	lients		Lilect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >5 iu to ≤ 10 iu	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance
Need for blood	transfusion											
2 (Fahmy 2015; Fenix 2012)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/80 (0%)	0/80 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>3</sup>	VERY LOW	CRITICAL
Need for blood	transfusion -	Vaginal b	irth									
1 (Fenix 2012)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious²	none	0/30 (0%)	0/30 (0%)	Not estimable	0 fewer per 1000 (from 60 fewer to 60 more) <sup>3</sup>	VERY LOW	CRITICAL
Need for blood	transfusion -	Caesarea	n birth									
1 (Fahmy 2015)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/50 (0%)	0/50 (0%)	Not estimable	0 fewer per 1000 (from 40 fewer to 40 more) <sup>3</sup>	VERY LOW	IMPORTANT
CI: confidence	interval											

1 Unclear risk of bias for blinding, incomplete outcome data, selective reporting 2 Sample size <200 3 Calculated from risk difference

#### Table 44: Oxytocin >1 iu to $\leq$ 5 iu versus Oxytocin <1 iu

			Quality asses	sment			No of pat	ients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu	Oxytocin <1 iu	Relative (95% CI)	Absolute	Quality	Importance
Need for blo	od transfusior	n - Caesaro	ean birth									
1 (Butwick 2010)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious²	none	0/30 (0%)	0/29 (0%)	Not estimable	0 fewer per 1000 (from 60 fewer to 60 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear bias for blinding, selective reporting 2 Sample size <200 3 Calculated from risk difference

#### Table 45: Oxytocin >1 iu to ≤ 5 iu versus Carbetocin

Quality assessment								No of patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu	Carbetocin	Relative (95% Cl)	Absolute	Quality	Importance
Need for blood transfusion - Vaginal birth												
1 (Amornpetchakul 2018)	randomised trials	no serious risk of bias	no serious inconsistency	no serious indirectness	serious <sup>1</sup>	none	0/174 (0%)	0/176 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>2</sup>	MODERATE	IMPORTANT
Cl: confidence interval												

1 Sample size 200-400 2 Calculated from risk difference

#### Table 46: Oxytocin >1 iu to $\leq$ 5 iu versus Placebo

Quality assessment								No of patients		Effect		
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin >1 iu to ≤ 5 iu	Placebo	Relative (95% CI)	Absolute	Quality	Importance
Need for blood transfusion												
2 (Butwick 2010; Jerbi 2007)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious²	none	0/95 (0%)	0/80 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>3</sup>	VERY LOW	IMPORTANT
Need for blood transfusion - Vaginal birth												
1 (Jerbi 2007)	randomised trials	very serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious <sup>2</sup>	none	0/65 (0%)	0/65 (0%)	Not estimable	0 fewer per 1000 (from 30 fewer to 30 more) <sup>3</sup>	VERY LOW	IMPORTANT
Need for blood transfusion - Caesarean birth												
1 (Butwick 2010)	randomised trials	serious <sup>4</sup>	no serious inconsistency	no serious indirectness	very serious²	none	0/30 (0%)	0/15 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>3</sup>	VERY LOW	

CI: confidence interval

1 Unclear risk of bias for randomisation, allocation concealment, blinding, selective reporting 2 Sample size <200

3 Calculated from risk difference

4 Unclear risk of bias for blinding, selective reporting

#### Table 47: Oxytocin <1 iu versus Placebo</th>

			Quality asses	ssment	No of patients			Effect				
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Oxytocin <1 iu	Placebo	Relative (95% Cl)	Absolute	Quality	Importance
Need for blood transfusion - Caesarean birth												
1 (Butwick 2010)	randomised trials	serious <sup>1</sup>	no serious inconsistency	no serious indirectness	very serious²	none	0/29 (0%)	0/15 (0%)	Not estimable	0 fewer per 1000 (from 10 fewer to 10 more) <sup>3</sup>	VERY LOW	IMPORTANT

CI: confidence interval

1 Unclear risk of bias for blinding, selective reporting
 2 Sample size <200</li>
 3 Calculated from risk difference

FINAL