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Transperineal biopsy in people with suspected prostate cancer - a systematic review and economic evaluation Addendum 2

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1 Meta-analysis scenarios

The DAR Addendum of 10 Jan 2022 included additional network meta-analysis (NMA) scenarios with the trial by Hara and colleagues (2008) ¹ either excluded, or re-labelled as a comparison between GATP and LATRUS, rather than LATP versus LATRUS as assumed in the original DAR analyses. This stemmed from expert advice that the methods of anaesthesia in the Hara study (spinal anaesthesia in the transperineal arm and caudal block in the transrectal arm) were more appropriately aligned with general anaesthesia rather than local anaesthesia. See section 1.1 of the Addendum for further explanation and results of these NMA scenarios.

It has since been noted that another trial (Takenaka et al. 2008)² classified in the original NMAs as a comparison between LATP and LATRUS had used the same methods of anaesthesia as in the Hara trial. We have therefore conducted further NMA analysis excluding or reclassifying both the Hara and Takenaka trials. We show the results of these additional NMA scenarios below.

1.1 Decision question 1

Hara and Takenaka both favour LATRUS over LATP. Thus, removing Takenaka as well as Hara (Figure 1 below) increases the relative risk of cancer detection for LATP and also for GATP (which is only connected to LATRUS via LATP). Whereas relabelling Takenaka and Hara as GATP versus LATRUS (Figure 2) reduces the relative risk for GATP but increases the relative risk for LATP.



Figure 1 Network meta-analysis forest plot of cancer detection rates for LATP-any vs LATRUS vs GATP grid and stepping device, excluding Hara et al 2008 and Takenaka et al 2008 (decision question 1)



Figure 2 Network meta-analysis forest plot of cancer detection rates for LATP-any vs LATRUS vs GATP grid and stepping device, relabelling Hara et al 2008 and Takenaka et al as GATP vs LATRUS (decision question 1)

1.2 Decision question 2

The transperineal arms in both Hara and Takenaka were conducted without a freehand device. Thus, removing these studies does not affect the results for LATP-freehand (which is only connected to LATRUS). Excluding Hara and Takenaka (Figure 3) increases the relative risk for LATP-other (without freehand device) and GATP; whereas relabelling these studies increases the relative risk for LATP-other but reduces the relative risk for GATP.

Comparison: other vs 'LATRUS'							
Treatment	(Random Ef	ffects Model)	RR	95%-CI			
GATP LATP-freehand LATP-other LATRUS	 	** *	1.01 1.40 1.05 1.00	[0.67; 1.51] [0.96; 2.04] [0.83; 1.34]			
	0.5	1 2	2				

Figure 3 Network meta-analysis forest plot of cancer detection rates for LATPfreehand vs LATP-other vs LATRUS vs GATP, excluding Hara et al 2008 and Takenaka et al 2008 (decision question 2)



Figure 4 Network meta-analysis forest plot of cancer detection rates for LATPfreehand vs LATP-other vs LATRUS vs GATP, relabelling Hara et al 2008 and Takenaka et al 2008 as GATP vs LATRUS (decision question 2)

2 Additional economic scenarios

In this section we show results for the above NMA scenarios applied to our revised base case (section 4 of the DAR Addendum of 10 Jan 2022). The additional scenarios have been added to the NMA scenario analyses reported in Tables 33 to 36 in the previous Addendum.

2.1 Decision question 1

When compared with the revised base case with only Hara excluded, the scenarios with both Hara and Takenaka excluded or relabelled reduce the ICERs for LATP (Table 1 and Table 2). With both studies excluded (scenario 3), the ICERs are below £20,000 per QALY in all modelled subgroups. With both studies relabelled (scenario 4), the ICERs are below £20,000 per QALY in subgroups A-C and £20,000 per QALY in subgroup D.

Bionsy mothod	DD a	Total		Incremental		ICERs		
Biopsy method		Cost	QALYs	Cost	QALYs	£/QALY		
Revised EAG base case: NMA excluding Hara								
LATRUS	1.00	£19,878	9.2989					
LATP-any	1.05	£19,937	9.3026	£58	0.0037	£15,669		
GATP	1.01	£20,420	9.3012	£483	-0.0014	Dominated		
NMA scenario	1: Hara c	lassified as	LATP-any ve	rsus LATRU	S			
LATRUS	1.00	£19,878	9.2989					
LATP-any	1.01	£19,944	9.3012	£66	0.0023	£28,322		
GATP	0.96	£20,430	9.2994	£486	-0.0018	Dominated		
NMA scenario	2: Hara c	lassified as	GATP versus	S LATRUS				
LATRUS	1.00	£19,878	9.2989					
LATP-any	1.03	£19,941	9.3019	£62	0.0030	£20,472		
GATP	0.92	£20,439	9.2978	£499	-0.0041	Dominated		
NMA scenario 3: Hara and Takenaka excluded								
LATRUS	1.00	£19,878	9.2989					
LATP-any	1.15	£19,919	9.3058	£40	0.0069	£5,859		
GATP	1.09	£20,405	9.3039	£486	-0.0019	Dominated		
NMA scenario 4: Hara and Takenaka classified as GATP versus LATRUS								
LATRUS	1.00	£19,878	9.2989					
LATP-any	1.09	£19,929	9.3039	£51	0.0050	£10,096		
GATP	0.92	£20,439	9.2978	£510	-0.0061	Dominated		
^a Relative risk for cancer detection compared with LATRUS								

 Table 1 NMA scenarios for decision question 1, subgroup A (deterministic)

Rionsy mothod	RR ^a	ICERs (£ per QALY gained)						
blopsy method		Subgroup A	Subgroup B	Subgroup C	Subgroup D			
Revised EAG base case: NMA excluding Hara								
LATRUS	1.00							
LATP-any	1.05	£15,669	£21,551	£21,095	£25,514			
GATP	1.01	Dominated	Dominated	Dominated	Dominated			
NMA scenario 1	I: Hara c	lassified as LA	FP versus LATR	US				
LATRUS	1.00							
LATP-any	1.01	£28,322	£30,256	£30,188	£31,261			
GATP	0.96	Dominated	Dominated	Dominated	Dominated			
NMA scenario 2	2: Hara c	lassified as GA	TP versus LATF	RUS				
LATRUS	1.00							
LATP-any	1.03	£20,472	£25,271	£24,939	£28,143			
GATP	0.92	Dominated	Dominated	Dominated	Dominated			
NMA scenario 3: Hara and Takenaka excluded								
LATRUS	1.00							
LATP-any	1.15	£5,859	£11,610	£11,111	£16,792			
GATP	1.09	Dominated	Dominated	Dominated	Dominated			
NMA scenario 4: Hara and Takenaka classified as GATP versus LATRUS								
LATRUS	1.00							
LATP-any	1.09	£10,096	£16,369	£15,840	£21,322			
GATP	0.92	Dominated	Dominated	Dominated	Dominated			
^a Relative risk for cancer detection compared with LATRUS								

Table 2 NMA scenarios for decision question 1, subgroup comparison (deterministic)

2.2 Decision question 2

The two new NMA scenarios do not affect the cost-effectiveness results for decision question 2 (Table 3 and Table 4 below). The ICER for LATP-freehand compared with LATRUS is constant across the NMA scenarios because the relative risk for this comparison does not depend on the Hara or Takenaka trial. The other comparators are dominated for all subgroups and NMA scenarios.

Biopsy method	RR ª	Total		Incremental		ICERs	
		Cost	QALYs	Cost	QALYs	£/QALY	
Revised EAG base case: NMA excluding Hara							
LATRUS	1.00	£19,878	9.2989				
LATP-freehand	1.40	£19,888	9.3122	£10	0.0133	£743	
LATP-other	0.98	£19,966	9.3001	£77	-0.0120	Dominated	
GATP	0.93	£20,437	9.2982	£471	-0.0019	Dominated	
NMA scenario 1:	Hara cla	assified as L	ATP-other v	ersus LATR	US		
LATRUS	1.00	£19,878	9.2989				
LATP-freehand	1.40	£19,888	9.3122	£10	0.0133	£743	
LATP-other	0.94	£19,974	9.2986	£86	-0.0135	Dominated	
GATP	0.90	£20,444	9.2970	£470	-0.0016	Dominated	
NMA scenario 2:	Hara cla	assified as G	SATP versus	LATRUS			
LATRUS	1.00	£19,878	9.2989				
LATP-freehand	1.40	£19,888	9.3122	£10	0.0133	£743	
LATP-other	0.97	£19,968	9.2998	£80	-0.0124	Dominated	
GATP	0.89	£20,446	9.2966	£478	-0.0032	Dominated	
NMA scenario 3:	Hara an	d Takenaka	excluded				
LATRUS	1.00	£19,878	9.2989				
LATP-freehand	1.40	£19,888	9.3122	£10	0.0133	£743	
LATP-other	1.05	£19,952	9.3026	£63	-0.0096	Dominated	
GATP	1.01	£20,420	9.3012	£468	-0.0014	Dominated	
NMA scenario 4: Hara and Takenaka classified as GATP versus LATRUS							
LATRUS	1.00	£19,878	9.2989				
LATP-freehand	1.40	£19,888	9.3122	£10	0.0133	£743	
LATP-other	1.01	£19,960	9.3012	£71	-0.0109	Dominated	
GATP	0.90	£20,444	9.2970	£484	-0.0042	Dominated	
^a Relative risk for cancer detection compared with LATRUS							

 Table 3 NMA scenarios for decision question 2, subgroup A (deterministic)

Biopsy method	RR ª	ICERs (£ per QALY gained)						
		Subgroup A	Subgroup B	Subgroup C	Subgroup D			
Revised EAG base case: NMA excluding Hara								
LATRUS	1.00							
LATP-freehand	1.40	£743	£4,595	£9,284	£10,640			
LATP-other	0.98	Dominated	Dominated	Dominated	Dominated			
GATP	0.93	Dominated	Dominated	Dominated	Dominated			
NMA scenario 1:	Hara cla	assified as LAT	P versus LATR	JS				
LATRUS	1.00							
LATP-freehand	1.40	£743	£4,595	£9,284	£10,640			
LATP-other	0.94	Dominated	Dominated	Dominated	Dominated			
GATP	0.90	Dominated	Dominated	Dominated	Dominated			
NMA scenario 2:	Hara re	classified as GA	ATP versus LAT	RUS				
LATRUS	1.00							
LATP-freehand	1.40	£743	£4,595	£9,284	£10,640			
LATP-other	0.97	Dominated	Dominated	Dominated	Dominated			
GATP	0.89	Dominated	Dominated	Dominated	Dominated			
NMA scenario 3:	Hara an	d Takenaka exc	cluded					
LATRUS	1.00							
LATP-freehand	1.40	£743	£4,595	£9,284	£10,640			
LATP-other	1.05	Dominated	Dominated	Dominated	Dominated			
GATP	1.01	Dominated	Dominated	Dominated	Dominated			
NMA scenario 4: Hara and Takenaka reclassified as GATP versus LATRUS								
LATRUS	1.00							
LATP-freehand	1.40	£743	£4,595	£9,284	£10,640			
LATP-other	1.01	Dominated	Dominated	Dominated	Dominated			
GATP	0.90	Dominated	Dominated	Dominated	Dominated			
^a Relative risk for cancer detection compared with LATRUS								

Table 4 NMA scenarios for decision question 2, subgroup comparison (deterministic)

References

- 1. Hara R, Jo Y, Fujii T, et al. Optimal approach for prostate cancer detection as initial biopsy: prospective randomized study comparing transperineal versus transrectal systematic 12core biopsy. *Urology* 2008;71(2):191-5. doi: 10.1016/j.urology.2007.09.029
- 2. Takenaka A, Hara R, Ishimura T, et al. A prospective randomized comparison of diagnostic efficacy between transperineal and transrectal 12-core prostate biopsy. *Prostate Cancer and Prostatic Diseases* 2008;11(2):134-38. doi: 10.1038/sj.pcan.4500985