

Amendment Number	Substantial/ Non-substantial	Details (docs named in AM approval letters)	Version	Date	Ethical Opinion
AM01	Substantial	Details of Hospital Admission and Colorectal Surgery,	2	July 2010	Favourable
		Baseline CRF	2	July 2010	
		Ineligible and Declined Form	3	August 2010	
		Short Summary	2	August 2010	
		Questionnaire: Baseline Patient Reported Outcome	2	July 2010	
		Protocol	3	August 2010	
		6-Weeks CRF	2	July 2010	
AM02	Substantial	Letter to surgeons	1	December 2010	Favourable
		1 week covering letter	1	December 2010	
		Details of Hospital Admission and Colorectal Surgery	3	December 2010	
		6 Weeks CRF	3	December 2010	
		Baseline CRF	3	December 2010	
		Advertisement	1	December 2010	
		Questionnaire – 1 year	2	December 2010	
		Questionnaire – 2 year	2	December 2010	
		Questionnaire – 5 year	2	December 2010	
		Questionnaire – 6 week	2	December 2010	
		Questionnaire – Baseline	3	December 2010	
		Protocol	4	16 December 2010	
		Questionnaire – Surgeons	1	December 2010	
AM03	Non-substantial	1, 2 and 5 year covering letter	1.1	January 2011	Not required
		1, 2 and 5 year reminder letter	1.1	January 2011	
		6 week covering letter	1.1	January 2011	
		6 week first reminder letter	1.1	January 2011	
		3 week covering letter	1.1	January 2011	
		Participant Cost Questionnaire letter sent at 18 months	1.1	January 2011	

		post haemorrhoid operation			
AM04			Error by REC		
AM05	Non-substantial	Protocol	4.1	24 March 2011	Not required
AM06	Non-substantial	6-week CRF	3.1	May 2011	Not required
		Baseline CRF	3.1	May 2011	
AM07	Substantial	Protocol, v5, Patient Information Leaflet	2.1	19 January 2012	Not approved
		Informed Consent Form	2.1	September 2011	
		Information for back of Consent form	1	September 2011	
		Baseline CRF	3.2	January 2012	
		Hospital Admission and Colorectal Surgery CRF	3.1	January 2012	
		Six Weeks CRF	3.2	January 2012	
		One Week Questionnaire	1.1	January 2012	
		Six Week Questionnaire	2.1	January 2012	
		Best Contact first letter	2.1	October 2011	
		Best Contact subsequent letter	2.1	October 2011	
		GP letter	1.1	October 2011	
AM7-1	Substantial	6 weeks CRF, v 3.2,	3.2	26 March 2012	favourable
		Questionnaire: Participant Costs Questionnaire	1.1	26 March 2012	
		Protocol	5.1	16 March 2012	
AM08			Error in numbering by REC		
AM09	Non-substantial	GP Information Sheet	1.1	October 2011	Not required
		GP Letter	1.1	October 2011	
		Project Summary	1	September 2011	
		Details of Hospital Admission and Colorectal Surgery	3.1	January 2012	

		Questionnaire: 1 Week	1.1	January 2012	
		Questionnaire: 6 Weeks	2.1	January 2012	
		Participant Information Leaflet	2.1	December 2011	
		Best Contact Letter with First Form	1.1	October 2011	
		Baseline CRF	3.2	January 2012	
		Best Contact Subsequent Letter	2.1	October 2011	
		Participant Consent Form	2.1	01 September 2011	
AM10	Non-substantial	Ineligible or declined form	3.1	01 June 2012	Not required
		Serious Adverse Event/Death Report Form	1.1	01 June 2012	
		Details of Hospital admission and colorectal surgery	3.2	01 June 2012	
		6 weeks CRF	3.3	01 June 2012	
		Baseline CRF	3.3	01 June 2012	
		Questionnaire 5 year	2.1	01 June 2012	
		Questionnaire: 6 week	2.2	01 June 2012	
		Questionnaire – Baseline	3.1	01 June 2012	
		Questionnaire 2 Year	2.1	01 June 2012	
		Questionnaire 1 Year	2.1	01 June 2012	
AM11	Non-substantial	1 2 5 year covering letter	1.3	August 2012	Not required
		Questionnaire 1 year	2.2	31 Aug 2012	
		Questionnaire 2 year	2.2	31 Aug 2012	
		Questionnaire 5 year	2.2	31 Aug 2012	
AM12	Substantial	Participant poster (Patient recruitment poster)	1	12 Sep 2012	favourable
AM13	Non-substantial	Questionnaire: Surgeon	1.1	28.9.12	Not required
AM14	Non-substantial	Questionnaire:2 years	2.3	October 2012	Not required

AM15	Non-substantial	Ineligible or Declined form	3.2	Jan 2013	Not required
AM16	Substantial	Protocol 1 and 2 year covering letter for IONTI	5.2 1	25.3.13 25 March 2013	favourable
AM17	Non-substantial	Pt review letter (1 and 2 yr IONTI covering letter)	1.1	29.04.2013	Not required
AM18	Non-substantial	Advertisement (International trials day poster)	5	19.04.2013	Not required
AM19	Substantial	Reminder letter - 1,2 &5 years Covering letter - 1,2 &5 years Participant Consent form Protocol Baseline CRF IONTI - 1 and 2 year covering letter SAE form Participant agreement to contact Participant Keeping in contact Sheet Participant Information Sheet: Post Operative Leaflet	1.3 1.4 2.2 5.3 3.4 1.2 1.2 1 1 1	28.6.13 28.6.13 May 2013 20.05.2013 May 2013 28.6.13 May 2013 4.7.13 28.6.13 28.6.13	favourable
AM 20	Non-substantial	6 week covering letter 6 week first reminder letter 3 week covering letter	1.2 1.2 1.2	Oct 2011 Oct 2011 Oct 2011	Not required
AM 21	Non-substantial	1 week covering letter Participant Cost Questionnaire covering letter	1.1 1.1	Oct 2011 Jan 2011	Not required

AM 22	Substantial	Protocol	5.4	26 Sep 2013	favourable
		Questionnaire : Reminder	1	26 Sep 2013	
		One, Two Year covering letter for IONTI	1.3	26 Sep 2013	
		18 months reminder letter	1	26 Sep 2013	
		1,2,5 Year Reminder 1 and 2 letter	1	26 Sep 2013	
		1,2 and 5 Year covering letter	1.5	26 Sep 2013	
AM 23	Non-substantial	Extension of recruitment from 30 April 2014 to 31 July 2014			Not required
AM 24	Substantial	Newsletter	1a	21 March 2014	favourable
		Newsletter	1b	21 March 2014	
AM 25	Non-substantial	1 and 2 Year covering letter for IONTI	1.4	31 March 2014	Not required
AM 26	Substantial	IONTI-Qual interviews		20 May 2014	Not approved
AM 27	Substantial	IONTI-2		8 Sep 2014	favourable
AM 28		REC numbering error			
AM 29(REC 28)	Non-substantial	GANT chart in Protocol updated to reflect recruitment extension	5.5	17 Dec 2014	Not required
		Addition of opt out sentence to receive voucher for IONTI 2 in 1,2,&5 year cover letter And reminder letter and inclusion of 18 month reminders	1.7,1.6, 1.1		

AM30(REC 29)	Non-substantial	Modified letter for 2Y reminders due to database error (86 reminders were not issued)	1	15 January 2015	Not required
AM30-1 (REC 30)	Non-substantial	Amendment of participant newsletters Amendment of trial manager name on 1week, 3week, 6week, GP letters and participant cost questionnaire covering letters	2a & 2b	5 March 2015	Not required
AM31(REC 32)	Non-substantial	Amendment of participant newsletters	2.1a & 2.1b	4 th September 2015	
AM32 (REC 31)	Substantial	DCE covering letter DCE questionnaire block 1 DCE questionnaire block 2 DCE questionnaire block 3 DCE questionnaire block 4	V1 V1 V1 V1 V1	August 2015	Favourable 15 September 2015
AM33 (REC 34)	Non-substantial	Amendment of participant newsletters	3.a & 3.b	3 December 2015	Not required
AM34 (REC 35)	Non-substantial	Protocol Amendment regarding DCE questionnaire (section 6.3 & 6.4) The number to be sent has been amended from n100 to n400 and the questionnaire is now dispatched at 27 months not 24 months to allow collection of the cost questionnaire.	V5.6	18 February 2016	Not required 23 February 2016
No AM35					
AM36 (REC 36)	Substantial	5 year follow-up questionnaire and covering letters Non-validated questionnaire - Participant Questionnaire eTHos Five year covering letter 5 year reminder 1 and 2 letter	V2.3 V1 V1	June 2016 30 May 2016 30 May 2016	Favourable 21 June 2016
AM37 (REC 37)	Non-substantial	Amendment to 5 year follow-up reminder letter Minor amendment to wording of the 5 year reminder 1 & 2	 V1	 30 June 2016	Not required 30 June 2016