CONTACT Protocol updates

Version change	Section number	Original detail	Amended detail	Rationale for amendment (if applicable)	Date of change (REC approval date)
v1.0 to v2.0	10.4.1 Expectedness Assessment	"SAEs which are reported to Keele CTU as related to trial treatment, will be formally assessed by the CI (or their delegate) for expectedness *within 1 day of receipt*."	"SAEs which are reported to Keele CTU as related to trial treatment, will be formally assessed by the CI (or their delegate) for expectedness ~~*within 1 day of receipt*~~."	Amended to comply with Keele CTU Standard Operating Procedures.	31/07/2013
	9.2 End of trial intervention	N/A	Updated title of Case Report Form.	To reflect agreed change to titles of clinical Case Report Form.	
	Throughout	N/A	Removal of references to web-based Case Report Forms.	The trial did not proceed with web-based Case Report Forms	
v2.0 to v3.0	Section 16 Confidentiality	"Information will be held securely on paper and electronically by Keele University and managed by Keele CTU."	"Information will be held securely on paper and ***managed*** electronically by Keele University ***through*** Keele CTU."	Clarification of arrangements to ensure confidentiality of information provided.	25/09/2013
		N/A, additional text added	"The trial data will be held on a database hosted on a secure server by the Primary Care Clinical Research and Trials Unit (PC-CRTU) at University of Birmingham. Provision of appropriate client server links/permissions will be given to authorised members of the trial team at Keele CTU."		
	Section 6.7 Randomisation	N/A, additional text added	"A back-up randomisation system will be in place should there be a failure of the web-based system."	Information provided confirming use of back-up randomisation procedure.	
V3.0 to v4.0	Section 6.6 – Table 1 and Section 8.1 Table 2	N/A, additional text added	Table 1: additional data item added to tables to be collected on day 1 (within the 7-day diary) as follows: "Time elapsed between symptom onset and starting trial medication"	On the advice of the Trial Steering Committee, an additional question was added to day 1 of the 7-day diary to assess the delay between symptom onset and starting	14/08/2014

				trial medication. 122 participants had been recruited prior to implementation of the amendment.	
	Section 6.7 Randomisation	N/A, additional text added	"If, during office hours, the randomisation system is online but the site network is down, the site will be instructed to call the Keele Clinical Trials Unit (CTU) and the CTU will perform the randomisation on the site's behalf. Authorised staff at the CTU will access the randomisation tool to perform randomisation and inform the healthcare professional of the allocation. If the site is unable to access the randomisation system outside of office hours or the site network is down and Keele CTU cannot access the database, the healthcare professional will call Aberdeen Health Services Research Unit (HSRU) who will provide an emergency 24/7 telephone randomisation service. Aberdeen HSRU will perform the randomisation on their behalf. The healthcare professional will be provided with a random allocation of treatment. To ensure that only patients registered with approved practices are randomised, Aberdeen HSRU will be provided with a list of participant ID numbers that can be used."	Additional information provided further clarifying back-up randomisation procedure.	
V4.0 to v5.0	Section 8.0 Follow-up and outcome assessment	"The week 4 questionnaire will also be available either as a paper or web-based questionnaire according to the participant's preference expressed at study entry."	"The week 4 questionnaire will also be available either as a paper or web-based questionnaire according to the participant's preference expressed ~~at study entry~~."	Some participants returned a paper 7-day diary (included in the baseline study pack) postally and did not enter any follow-up data in the web-based diary despite choosing web-based follow-up at study entry. Amendment requested to allow such participants to be sent a paper 4-week questionnaire postally rather than e-mailed about a web-based 4-week questionnaire.	03/03/2015

V5.0 to v6.0	Section 16 Confidentiality	N/A, additional text added	"At the end of the study, all trial data will be moved from the secure server at the University of Birmingham to a secure server at Keele University under a data migration plan. All data will be stored securely and in line with the Data Protection Act 1998 at all times. At this time, all participants will be notified of this planned move and will be given the opportunity to raise any queries or concerns that have regarding the move of this data. Following the move of the data, the data will go through a series of data cleaning stages, prior to the final dataset being locked down for analysis prior to the data being archived."	The whole trial database needed to be moved from a secure server at the University of Birmingham to a secure server at Keele University at the end of the study for analysis and archiving. The Participant Information Leaflet stated that the data will be stored at the University of Birmingham. This protocol amendment was required to described updated arrangements for data storage and notification of participants	21/04/2016

CI, Chief Investigator; CTU; Clinical Trials Unit; ID, identification; N/A, not applicable; REC, Research Ethics Committee; SAE, Serious Adverse Event