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**Instructions: Unfortunately, the manuscript system did not allow for Microsoft Excel files as supplementary files, only Microsoft Word. Therefore we have prepared this file to share our underlying dataset. To work with the data below, it may be easiest to select the table below and copy all values into a spreadsheet program e.g. Excel.**

Trial ID	ROB element	2012 assessment	2012 rationale
M76001	Random sequence generation (selection bias)	Low	
M76001	Allocation concealment (selection bias)	Low	
M76001	Incomplete outcome data (attrition bias), symptoms	Low	
M76001	Incomplete outcome data (attrition bias), complications of influenza	Unclear	Unclear how complications of influenza were defi
M76001	A159: Incomplete outcome data (attrition bias) safety	Low	
M76001	Safety data	Low	
M76001	A159: Selective reporting (reporting bias)	Low	
M76001	A159: Other bias		
M76001	A159: Blinding of participants and personnel (performance bias)		
M76001	All outcomes	Unclear	Capsule size, but no details of colour or taste or ci
M76001	A159: Blinding of outcome assessment (detection bias)	Low	
M76001	All outcomes	Low	
NV16871	Random sequence generation (selection bias)	Low	
NV16871	Allocation concealment (selection bias)	Low	

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NV16871	Incomplete outcome data (attrition bias), symptoms	Low	
NV16871	Incomplete outcome data (attrition bias), complications of influenza	Low	
NV16871	A159: Incomplete outcome data (attrition bias)		
NV16871	Safety data	Low	
NV16871	A159: Selective reporting (reporting bias)		
NV16871	A159: Other bias		
NV16871	A159: Blinding of participants and personnel (performance bias)		
NV16871	All outcomes	Unclear	Placebo colour/taste/contents not clear
NV16871	A159: Blinding of outcome assessment (detection bias)		
NV16871	All outcomes	Low	
WP16263	Random sequence generation (selection bias)	Unclear	Unclear risk Described as randomised; procedure
WV15670	Random sequence generation (selection bias)	Unclear	Described as randomised; procedure generating r "The randomisation numbers were generated by ; inc., Princeton, NJ, USA)." "The investigator telephoned the centre to report
WV15670	Allocation concealment (selection bias)	Low	The randomization number was then supplied by system (IVRS). The investigator entered the rand
WV15670	Incomplete outcome data (attrition bias), symptoms	High	Available data analyzed by ITTI population and no
WV15670	Incomplete outcome data (attrition bias),	High	Possible effect of oseltamivir on antibody product complications in the infected subpopulation non-i

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WV1567 0	complications of influenza Incomplete outcome data (attrition bias), safety data	Low	Based on all participants irrespective of complian
WV1567 0	Selective reporting (reporting bias), other bias	High	Outcomes of primary interest for the ITT populati
WV1567 0	Other bias	Unclear	Placebo contained dehydrocholic acid. Dosage no "In order to maintain blinding, each subject had 2 administered from each bottle twice per day at a the first (day 1) visit
WV1567 0	Blinding of participants and personnel (performance bias), all outcomes	Low	Each bottle was labelled with the subject number placebo. Those subjects receiving 75 mg bid recei matching capsule containing placebo from the oth received one capsule containing 75 mg active dru "No open key to the randomisation code was avai
WV1567 0	Blinding of outcome assessment (detection bias), all outcomes	Low	Roche Headquarters. In the event of a medical er necessary to properly manage the subject, by con
WV1567	Random sequence	Unclear	The blinding was not required to be broken for an Described as randomised; procedure generating

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1	generation (selection bias)		randomisations schedule not available
1	Allocation concealment (selection bias)	Low	“Randomisation was conducted by a central randomiser. The investigator /study coordinator telephoned the subjects initials, date of birth and smoking history. The randomisation number was entered in the appropriate place on the subject’s Case Report Form by the investigator.”
1	Incomplete outcome data (attrition bias), symptoms	Low	Data from study participants without influenza were available for symptom relief
1	Incomplete outcome data (attrition bias), complications of influenza	High	Possible effect of oseltamivir on antibody production makes the assessment of influenza status and associated complications in the infected subpopulation non-comparable between the treatment groups
1	Incomplete outcome data (attrition bias), safety data	Low	Based on all participants irrespective of compliance with treatment or infection status
1	Selective reporting (reporting bias), other bias	Low	Outcomes of primary interest for the ITT population available in the CONSORT reconstruction
1	Other bias	High	Placebo contained dehydrocholic acid Matching placebo used “In order to maintain the double blind nature of the study, subjects received 2 capsules twice daily for all treatments.” “The identification number was added by the investigator at the time of randomisations” “No open key to the code was available at the Study Center...” “The identification number was added by the investigator at the time of randomisations.” “No open key to the code was available at the Study Center, to the Monitors, Statisticians or at Gilead/Roche Headquarters”
1	Blinding of participants and personnel (performance bias), all outcomes	Low	
1	Blinding of outcome assessment (detection bias), all outcomes	Low	
3	Random sequence generation (selection bias)	Unclear	Described as randomised; procedure generating random numbers
3	Allocation concealment (selection bias)	Unclear	Inadequate information available to ascertain concealment

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7			
WV1567			
3	Incomplete outcome		
WV1569	data (attrition bias),		
7	symptoms	Low	Not applicable to the study design (prophylaxis)
WV1567	Incomplete outcome		
3	data (attrition bias),		
WV1569	complications of		
7	influenza	High	Possible effect of oseltamivir on antibody product complications in the infected subpopulation non-i
WV1567	A159: Incomplete		
3	outcome data (attrition		
WV1569	bias)		
7	Safety data	Low	Based on all randomised participants
WV1567			
3	A159: Selective		
WV1569	reporting (reporting		
7	bias)	Low	Outcomes of primary interest for the ITT populati
WV1567			
3			
WV1569			
7	A159: Other bias	Unclear	Placebo contained dehydrocholic acid. Dosage no
WV1567	A159: Blinding of		
3	participants and		
WV1569	personnel (performance		
7	bias)	Unclear	Capsule size, but no details of colour or taste or c
WV1567	All outcomes		
3	A159: Blinding of		
WV1569	outcome assessment		
7	(detection bias)	Unclear	Inadequate information available to ascertain wh
WV1570	All outcomes		
7	Random sequence	Unclear	
WV1570	generation (selection		
7	bias)	Unclear	Described as randomised; procedure generating r
WV1570	Allocation concealment		
7	(selection bias)	Low	"Randomization was performed by a central rand the subject's date of birth, vaccination status and randomisation centre."
WV1570	Incomplete outcome		
7	data (attrition bias),		
WV1570	symptoms	High	Available data analyzed by ITTI population and no
WV1570	Incomplete outcome		
7	data (attrition bias),		
WV1570	complications of		
7	influenza	High	Possible effect of oseltamivir on antibody product complications in the infected subpopulation non-i

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WV1570 7	A159: Incomplete outcome data (attrition bias) Safety data	Low	Based on all randomised participants
WV1570 7	A159: Selective reporting (reporting bias)	High	Outcomes of primary interest for the ITT population
WV1570 7	A159: Other bias A159: Blinding of participants and personnel (performance bias)	Unclear	Placebo contained dehydrocholic acid. Dosage not stated
WV1570 7	All outcomes A159: Blinding of outcome assessment (detection bias)	Low	Presentation of placebo described as identical
WV1570 7	All outcomes Random sequence generation (selection bias)	Unclear	Inadequate information available to ascertain whether randomization was truly random
WV1570 8	Allocation concealment (selection bias)	Unclear	Randomization numbers generated by Roche, but details not provided
WV1570 8	Incomplete outcome data (attrition bias), symptoms	Low	Insufficient details given
WV1570 8	Incomplete outcome data (attrition bias), complications of influenza	Low	Outcomes available on all patients who complete study
WV1570 8	A159: Incomplete outcome data (attrition bias) Safety data	Low	Outcome data on all patients provided.
WV1570 8	A159: Selective reporting (reporting bias)	Low	Outcome data reported.
WV1570 8	A159: Other bias A159: Blinding of participants and personnel (performance bias)	Unclear	Placebo contents and colour and similarity to active treatment not stated. Could not analyze for primary outcome of efficacy
WV1570 8	All outcomes	Low	

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WV1570 8	A159: Blinding of outcome assessment (detection bias) All outcomes Random sequence	Low	Outcome assessors were blind
WV1573 0	generation (selection bias)	Unclear	Described as randomised; procedure generating randomisations schedule not available "Randomization was performed by a central randomisations service. The investigator telephoned the centre to report the subject's data number was then supplied by the randomisations
WV1573 0	Allocation concealment (selection bias)	Low	
WV1573 0	Incomplete outcome data (attrition bias), symptoms	High	Available data analysed by ITTI population and not ITT Possible effect of oseltamivir on antibody production makes the assessment of influenza status and associated complications in the infected subpopulation non-comparable between the treatment groups
WV1573 0	Incomplete outcome data (attrition bias), complications of influenza	High	
WV1573 0	Incomplete outcome data (attrition bias), safety data	Low	Based on all randomised participants
WV1573 0	Selective reporting (reporting bias), other bias	High High	Outcomes of primary interest for the ITT population not made available to the review auth
WV1573 0	Other bias		Placebo capsule contained dehydrocholic acid
WV1573 0	Blinding of participants and personnel (performance bias), all outcomes	Low	Matching placebo. "No open key to the code was available at the study centre, to the monitors, statistician or at Roche Headquarters. In the event of a medical emergency the blinding was to be broken if considered absolutely mandatory to properly manage the patient
WV1573 0	Blinding of outcome assessment (detection bias), all outcomes	Low	
WV1575 8	Random sequence generation (selection bias)	Unclear	Described as randomised; procedure generating randomisations schedule not available
WV1575 8	Allocation concealment (selection bias)	Low	"Randomization was conducted by a central randomisations service, ICTI (Interactive

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			Clinical Technologies Inc., Princeton, NJ). The investigator telephoned the centre to report the subject's date of birth, sex, a centre in the form of a message on an interactive response system (IVRS). The investigator entered the randomisations number in the appropriate place on the case report form. The subject randomisations numbers were allocated sequentially within a stratum in the order in which subjects were enrolled."
WV1575 8	Incomplete outcome data (attrition bias), symptoms	Low	Data available for both influenza infected and non-infected study populations
WV1575 8	Incomplete outcome data (attrition bias), complications of influenza	High	Possible effect of oseltamivir on antibody production makes the assessment of influenza status and associated complications in the infected subpopulation non-comparable be
WV1575 8	Incomplete outcome data (attrition bias), safety data	Low	Based on all randomized patients
WV1575 8	Selective reporting (reporting bias), other bias	Low	Outcomes of primary interest to the review for ITT population available in the CONSORT-based extraction reconstruction
WV1575 8	Other bias	Unclear	Unable to ascertain placebo capsule contents
WV1575 8	Blinding of participants and personnel (performance bias), all outcomes	Low	"No open key to the code was available at the study centre..."
WV1575 8	Blinding of outcome assessment (detection bias), all outcomes	Low	"No open key to the code was available (...) to the Roche monitors, statisticians or at Roche Headquarters."
WV1575 9	Random sequence generation (selection bias)	Unclear	Described as randomised; procedure generating randomisations schedule not available
WV1587 1	Allocation concealment (selection bias)	Low	The subject randomizations numbers will be generated by Roche or its designee and incorp Randomization will be conducted by a central randomization service by telephone.
WV1575 9	Incomplete outcome data (attrition bias), symptoms	Unclear	Insufficient information was available to ascertain populations for analysis and judge risk of bias



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WV1575 9 WV1587 1 WV1575 9 WV1587 1 WV1575 9 WV1587 1 WV1575 9 WV1587 1 WV1575 9 WV1587 1 WV1575 9 WV1587 1 WV1579 9 WV1579 9 WV1579 9 WV1579 9	<p>Incomplete outcome data (attrition bias), complications of influenza</p> <p>Incomplete outcome data (attrition bias), safety data</p> <p>Selective reporting (reporting bias), other bias</p> <p>Other bias</p> <p>Blinding of participants and personnel (performance bias), all outcomes</p> <p>Blinding of outcome assessment (detection bias), all outcomes</p> <p>Random sequence generation (selection bias)</p> <p>Allocation concealment (selection bias)</p> <p>Incomplete outcome data (attrition bias), symptoms</p> <p>Incomplete outcome data (attrition bias), complications of influenza</p> <p>Incomplete outcome data (attrition bias), safety data</p> <p>Selective reporting (reporting bias), other bias</p>	<p>Unclear</p> <p>Unclear</p> <p>High</p> <p>High</p> <p>Low</p> <p>Unclear</p> <p>Unclear</p> <p>Low</p> <p>High</p> <p>Low</p> <p>High</p>	<p>Insufficient information was available to ascertain populations for analysis and judge risk of bias</p> <p>Insufficient information was available to ascertain populations for analysis and judge risk of bias</p> <p>No outcome data were provided in the study CONSORT-based extraction reconstruction</p> <p>Placebo capsule contained dehydrocholic acid</p> <p>Matching placebo</p> <p>Inadequate information available to ascertain whether outcome assessors were aware of treatment group assignment</p> <p>Described as randomised; procedure generating randomisations schedule not available</p> <p>Inadequate information available to ascertain concealment of allocation</p> <p>Not applicable to the study design (prophylaxis) Possible effect of oseltamivir on antibody production makes the assessment of influenza status and associated complications in the infected subpopulation non-comparable between the treatment groups</p> <p>Based on all randomised participants</p> <p>Outcome data for ITT population were not available to the review authors</p>
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WV1579 9	Other bias Blinding of participants and personnel	Unclear	No information available on placebo contents
WV1579 9	(performance bias), all outcomes	Unclear	Inadequate information available to ascertain presentation of placebo capsules
WV1579 9	Blinding of outcome assessment (detection bias), all outcomes	Unclear	Inadequate information available to ascertain whether outcome assessors were aware of treatment group assignment
WV1581 2	Random sequence generation (selection bias)	Unclear	Described as randomised; procedure generating r "The randomisation numbers were generated by : inc., Princeton, NJ, USA)."
WV1581 2	Allocation concealment (selection bias)	Low	"The investigator telephoned the centre to report The randomization number was then supplied by system (IVRS). The investigator entered the rand
WV1581 2	Incomplete outcome data (attrition bias), symptoms	High	Available data analyzed by ITTI population and no
WV1581 2	Incomplete outcome data (attrition bias), complications of influenza	High	Possible effect of oseltamivir on antibody product complications in the infected subpopulation non-i
WV1581 2	Incomplete outcome data (attrition bias), safety data	Low	Based on all participants irrespective of complian
WV1581 2	Selective reporting (reporting bias), other bias	High	Outcomes of primary interest for the ITT populati
WV1581 2	Other bias Blinding of participants and personnel	Unclear	Placebo contained dehydrocholic acid. Dosage no
WV1581 2	(performance bias), all outcomes	Low	Matching placebo described
WV1581	Blinding of outcome	Unclear	Inadequate information available to ascertain wh

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2 WV1587 2 WV1581 9 WV1587 6	assessment (detection bias), all outcomes		of treatment group assignment
WV1597 8 WV1581 9 WV1587 6	Random sequence generation (selection bias)	Unclear	Described as randomised; procedure generating randomisations schedule not available "Randomization was conducted by a central randomisations service via telephone. The investigator or study coordinator telephoned vaccination status and history of COAD, and the telephone number was then supplied by the centre. The randomisation was conducted in the appropriate place on the subject's Case Report Form.
WV1597 8 WV1581 9 WV1587 6	Allocation concealment (selection bias)	Low	
WV1597 8 WV1581 9 WV1587 6	Incomplete outcome data (attrition bias), symptoms	Low	Available data analysed for both by ITTI and ITT populations
WV1597 8 WV1581 9 WV1587 6	Incomplete outcome data (attrition bias), complications of influenza	High	Possible effect of oseltamivir on antibody production makes the assessment of influenza status and associated complications in the infected subpopulation non-comparable between the treatment groups
WV1597 8 WV1581 9 WV1587 6	Incomplete outcome data (attrition bias), safety data	Low	Based on all randomised participants
WV1597 8 WV1581 9 WV1587 6	Selective reporting (reporting bias), other bias	Low	Outcomes of primary interest to the review are available in the CONSORT-based extraction reconstruction
WV1597 6	Other bias	High	Placebo capsule contained dehydrocholic acid

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8			
WV1581			
9			
WV1587	Blinding of participants and personnel		
6			
WV1597	(performance bias), all outcomes	Low	Matching placebo described
8			
WV1581			
9			
WV1587			
6	Blinding of outcome assessment (detection bias), all outcomes	Low	“No open key to the code was available at the study centres, to the monitors, statisticians or at Roche headquarters. In the event of a medic mandatory to properly manage the subject, by co the randomisations centre.”
WV1597	Random sequence generation (selection bias)		
8			
WV1582		Unclear	Described as randomised; procedure generating r
5			
WV1582	Allocation concealment (selection bias)	Unclear	Inadequate information available to ascertain con
5			
WV1582	Incomplete outcome data (attrition bias), symptoms	Low	Not applicable to the study design (prophylaxis)
5			
WV1582	Incomplete outcome data (attrition bias), complications of influenza	High	Possible effect of oseltamivir on antibody product complications in the infected subpopulation non-i
5			
WV1582	Incomplete outcome data (attrition bias), safety data	Low	Based on all randomised participants
5			
WV1582	Selective reporting (reporting bias)	High	Outcome data relating to complications were not
5			
WV1582	Other bias	Unclear	Placebo contained dehydrocholic acid. Dosage no
5			
WV1582	Blinding of participants and personnel		
5			
WV1582	(performance bias), all outcomes	Unclear	
5			
WV1582	Blinding of outcome assessment (detection bias), all outcomes	Unclear	
5			