

Instructions

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4. Intellectual Property.

This section asks about patents and copyrights, whether pending, issued, licensed and/or receiving royalties.

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Section 1.	Identifying Inform	nation		
1. Given Name (F Johan	irst Name)	2. Surname (Last Name) Sundstrom		3. Date 05-November-2014
4. Are you the co	rresponding author?	✓ Yes No		
5. Manuscript Titl Effects of Blood		Mild Hypertension: Systema	tic Review and Meta-Anal	lysis
6. Manuscript Ide M14-0773	ntifying Number (if you k	now it)		
	•			
Section 2.	The Work Under C	onsideration for Public	ation	
any aspect of the statistical analysis	submitted work (includin , etc.)?	g but not limited to grants, dat	. , .	ommercial, private foundation, etc.) for esign, manuscript preparation,
-	levant conflicts of inter	est? 🗌 Yes 🖌 No		

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Are there any relevant conflicts of interest?		Yes	\checkmark	No
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Section 4. **Intellectual Property -- Patents & Copyrights**

Do you have any patents, whether planned, pending or issued, broadly relevant to the work?	Yes	\checkmark	No



Section 5. Relationships not covered above

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On an advisory board for Itrim, a weight loss company.

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Section 6.

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1. Given Name (First Name) Kazem	2. Surname (Last Name) Rahimi	3. Date 11-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Sundstrom
5. Manuscript Title Effects of Blood Pressure Reduction in N	/lild Hypertension: System	atic Review and Meta-Analysis
6. Manuscript Identifying Number (if you kn	low it)	
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The Work Under Co	onsideration for Public	ation
any aspect of the submitted work (including statistical analysis, etc.)?	but not limited to grants, da	a third party (government, commercial, private foundation, etc.) for ta monitoring board, study design, manuscript preparation,
Are there any relevant conflicts of intere	est? Yes 🖌 No	
Section 3. Relevant financial	activities outside the s	ubmitted work.
Place a check in the appropriate boxes i		

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Are there any relevant conflicts of interest?	Yes	🖌 No
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	1 1		



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3. Date 06-December-2014
or's Name
a-Analysis

Section 2. The Work Under Consideration for Publication

Did you or your institution **at any time** receive payment or services from a third party (government, commercial, private foundation, etc.) for any aspect of the submitted work (including but not limited to grants, data monitoring board, study design, manuscript preparation, statistical analysis, etc.)?

🖌 No

Are there any relevant conflicts of interest? Yes

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No

Are there any relevant conflicts of interest? \checkmark Yes

If yes, please fill out the appropriate information below.

Name of Entity	Grant?	Personal Fees	Non-Financial Support?	Other?	Comments	
Servier International	\checkmark	\checkmark			for the ADVANCE-ON post-trial follow-up study	

-		
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20	CUIU	

Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes 🗸 No



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Dr. Chalmers reports grants and personal fees from Servier International, outside the submitted work; .

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Section 1.	Identifying Inform	ation			
1. Given Name (Fii Bruce	rst Name)	2. Surnar Neal	me (Last Name)		3. Date 11-April-2014
4. Are you the corresponding author?		Yes	✓ No	Corresponding Author's Name Johan Sundstrom	
5. Manuscript Title Effects of Blood Pressure Reduction in Mild Hypertension: Systematic Review and Meta-Analysis					
6. Manuscript Ider	ntifying Number (if you kn	ow it)			

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No

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If yes, please fill out the appropriate information below.

Name of Entity	Grant?	Personal Fees	Non-Financial Support <mark>?</mark>	Other?	Comments	
Abbott		\checkmark			Honorarium for meeting presentation	
Novartis		\checkmark			Honorarium for meeting presentation	
Pfizer		\checkmark			Honorarium for meeting presentation	
Roche		\checkmark			Honorarium for meeting presentation	
Servier		\checkmark			Honorarium for meeting presentation	
Janssen	\checkmark				Large-scale outcome trial in diabetes	
Abbvie	\checkmark				Large-scale outcome trial in CKD	
Dr Reddy's Laboratories	\checkmark				Large-scale outcome trial in CVD	



Name of Entity	Grant?	Personal Fees ?	Non-Financial Support?	Other?	Comments
Servier	\checkmark				Large-scale outcome trial in CVD
Merck Schering Plough	\checkmark				Large-scale outcome trial in CVD
Roche	\checkmark				Large-scale outcome trial in CVD

Section 4.

Intellectual Property -- Patents & Copyrights

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Dr. Neal reports grants from National Health and Medical Research Council of Australia, during the conduct of the study; grants from Roche, personal fees from Abbott, personal fees from Novartis, personal fees from Pfizer, personal fees from Roche, personal fees from Servier, grants from Janssen, grants from Abbvie, grants from Dr Reddy's Laboratories, grants from Servier, grants from Merck Schering Plough, outside the submitted work; .



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Dr. Jackson has nothing to disclose.

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1. Given Name (First Name) Hisatomi		2. Surname (Last Nam Arima	e)	3. Date 04-November-2014
4. Are you the correspondin	ig author?	Yes 🖌 No	Corresponding Author's Na	ame
 Manuscript Title Effects of Blood Pressure Manuscript Identifying No. 			tematic Review and Meta-Ana	lysis
	uniber (ir you			

Section 2. The Work Under Consideration for Publication

Did you or your institution **at any time** receive payment or services from a third party (government, commercial, private foundation, etc.) for any aspect of the submitted work (including but not limited to grants, data monitoring board, study design, manuscript preparation, statistical analysis, etc.)?

🖌 No

Are there any relevant conflicts of interest?	Yes
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Section 3. Relevant financial activities outside the submitted work.

Place a check in the appropriate boxes in the table to indicate whether you have financial relationships (regardless of amount of compensation) with entities as described in the instructions. Use one line for each entity; add as many lines as you need by clicking the "Add +" box. You should report relationships that were **present during the 36 months prior to publication**.

Are there any relevant conflicts of interest?	Yes	\checkmark	No
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Section 4. Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work?	Yes	\checkmark	No



Section 5. Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

Yes, the following relationships/conditions/circumstances are present (explain below):

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Section 6. Disclosure Statement

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Dr. Arima has nothing to disclose.

Evaluation and Feedback



Instructions

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Section 1.	Identifying Infor	mation	
1. Given Name (Fi Mark	rst Name)	2. Surname (Last Name) Woodward	3. Date 04-November-2014
4. Are you the cor	responding author?	Yes 🖌 No	Corresponding Author's Name Sundstrum
5. Manuscript Title Effects of Blood		n Mild Hypertension: Syster	natic Review and Meta-Anal
6. Manuscript Ider M14-0773	ntifying Number (if you	know it)	

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✓ No

Are there any relevant conflicts of interest? Yes

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No

Are there any relevant conflicts of interest? \checkmark Yes

If yes, please fill out the appropriate information below.

Name of Entity	Grant?	Personal Fees	Non-Financial Support?	Other?	Comments	
Novartis		\checkmark			Advice on the design of a new trial	

Section 4. Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes 🗸 No



Section 5. Relationships not covered above

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Dr. Woodward reports personal fees from Novartis, outside the submitted work; .

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1. Given Name (First Name) Fiona	2. Surname (Last Name) Turnbull	3. Date 12-November-201
. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundstrom
. Manuscript Title ffects of Blood Pressure Reduction ir	n Mild Hypertension: Syste	matic Review and Meta-Analysis
5. Manuscript Identifying Number (if you W14-0773	know it)	

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🖌 No

Are there any re	levant confl	icts of intere	est?	Yes
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Are there any relevant conflicts of interest? Yes 🗸 No

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Do you have any patents, whether planned, penuing of issued, broadly relevant to the work? res \mathbf{v} no	e any patents, whether planned, pending or issued, broadly relevant to	the work?	Yes	🖌 No
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Royalties: Funds are coming in to you or your institution due to your patent

5.



1. Given Name (I LAWRENCE	First Name)	2. Surname (Last Name AGODOA	3. Date 15-December-2014
4. Are you the co	prresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundstrom
5. Manuscript Tit Effects of Blood		Aild Hypertension: A Sys	tematic Review and Meta-analysis.
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Section 5. Relationship

Relationships not covered above

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1. Given Name (First N Craig	ame)	2. Surnan Andersoi	ne (Last Name) n		3. Date 17-Decen	nber-2014
4. Are you the correspo	onding author?	Yes	V No	Corresponding Autho Johan Sundström		
5. Manuscript Title Effects of Blood Pres	sure Reduction in	Mild Hypert	ension: A Syste	matic Review and Me	eta-analysis	
6. Manuscript Identifyi	ng Number (if you l	know it)				
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Section 1. Identifying Infor	mation	
1. Given Name (First Name) Folkert	2. Surname (Last Name) Asselbergs	3. Date 12-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Sundström
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syste	ematic Review and Meta-analysis.
6. Manuscript Identifying Number (if you	know it)	
Section 2. The Work Under	Consideration for Publi	ication
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Dr. Asselbergs has nothing to disclose.

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identifying info	rmation	I STATE I STATE I STATE
1. Given Name (First Name) Barry	2. Surname (Last Name) Brenner	3. Date 21-January-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundström
5. Manuscript Title Effects of Blood Pressure Reduction in	n Mild Hypertension: A Syste	matic Review and Meta-analysis
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Section 1. Identifying Info	ormation	
1. Given Name (First Name) Morris	2. Surname (Last Name) Brown	3. Date 20-January-2015
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundström
5. Manuscript Title Effects of Blood Pressure Reduction	in Mild Hypertension: A Syste	matic Review and Meta-analysis
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Section 1. Identifying Infor	mation	
1. Given Name (First Name) christopher	2. Surname (Last Name) bulpitt	3. Date 01-July-2015
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Dr Sundstrom
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syste	ematic Review and Meta-analysis
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Section 1. Identifying Inform	mation	and a second
1. Given Name (First Name) Robert	2. Surname (Last Name) Byington	3. Date 23-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Sundstrom
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syste	ematic Review and Meta-analysis
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1. Given Name (First Name) Rory	2. Surname (Last Name) Collins		3. Date 23-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Auth Johan Sundström	oor's Name
5. Manuscript Title Effects of Blood Pressure Reduction ir	n Mild Hypertension: A Syste	matic Review and M	eta-analysis
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See attached list of grants and approach to maintaining independence of research.

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Section 6.

Disclosure Statement

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Dr. Collins reports various grants to Oxford University for CTSU outside the submitted work; and See attached list of grants and approach to maintaining independence of research.

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Grants to Oxford University for any Clinical Trial Service Unit (CTSU) trials or other commercially-funded research over the past 20 years (but excluding CTSU core support from MRC, CR-UK and BHF for these and other studies, as well as other non-commercially funded research)

ACST-2 trial of carotid procedures (2007-ongoing) BUPA: £60K NHS-HTA: £200K ASCEND trial of aspirin and fish oils (2004-ongoing) British Heart Foundation: £2.7M Abbott/Solvay: £2.0M plus drug supply Bayer: £1.7M plus drug supply ATLAS trial of tamoxifen duration (1997-ongoing) US Army: \$1.6M AstraZeneca: £1.0M plus drug supply ATLAS trial of psychosis treatment (2011-ongoing) NHS-HTA: £500K BEST-D pilot trial of vitamin D (2012-ongoing) British Heart Foundation: £150K Tishcon: free drug supply only CCS-2 trial of metoprolol and clopidogrel (1999-2005) AstraZeneca: £1.1M plus drug supply Sanofi: £1.1M plus drug supply China Kadoorie Biobank (2002-ongoing) Kadoorie Trust: £4.7M Medical Research Council: £1.3M Wellcome Trust: £2.5M AstraZeneca: \$300K GlaxoSmithKline: £2.2M Merck: £200K Elinogrel feasibility trial (2010-2011) Novartis: £500K Establishing Fuwai-Oxford research centre (2010-ongoing) Merck: £1.1M Heart Protection Study (1993-2002) British Heart Foundation: £1.2M Medical Research Council £9.6M Merck: £5.5M plus drug supply Roche: £5.5M plus drug supply Heart Protection Study follow-up studies (2003-ongoing) BHF: £350K Merck: £1.2M GlaxoSmithKline: \$400K Liposcience: £50K HPS2-THRIVE trial of niacin/laropiprant (2005-ongoing) Merck: £53M plus drug supply HPS3/TIMI55-REVEAL trial of anacetrapib (2010-ongoing) Merck: £96M plus drug supply Leukaemia trials (2010-ongoing) Cancer Research UK: £200K

Medical Research Council: £2.8M Kay Kendall Leukaemia Fund: £200K Pfizer Innovation Award (2004) Pfizer: £50K to CTSU for unrestricted research PROCARDIS genetic study (1998-2011) European Union: £2.0M AstraZeneca: £1.7M SEARCH trial of simvastatin dose (1997-2010) Merck: £22.7M plus drug supply SHARP trial of simvasatin/ezetimibe (2002-2013) Merck/Schering: £40M plus drug supply STICS trial of rosuvastatin (2011-2014) British Heart Foundation: £225K European Union: £15K AstraZeneca: \$100K 3-C trial of transplant rejection (2009-2017) NHS Blood Transfusion: £250K Pfizer: £500K Novartis: £350K

Patents: Myopathy-related genetic variant; licensed by Oxford University to Boston Heart Diagnostics; all personal income waived in favour of the University and CTSU to support research

<u>NOTE</u>: The CTSU conducts, analyses and interprets its clinical trials and other research independently of industry and other funders, with the datasets held by the CTSU rather than by the funders. In accordance with CTSU's long-term policy (see attached), honoraria, consultancy or other payments have not been received directly or indirectly from industry, either personally by Professor Collins or by the University (except for reimbursement of travel and accommodation costs for taking part in relevant scientific meetings).

Funding and independence of research at the Clinical Trial Service Unit & Epidemiological Studies Unit ("CTSU") of the Nuffield Department of Population Health, University of Oxford

CTSU is a world leader in the conduct of large-scale randomised controlled trials, and combined analyses ("meta-analyses") of detailed data from randomised trials, which provide reliable evidence about the safety and efficacy of treatments. Such trials and meta-analyses – which may be of existing or new treatments – are essential to help guide health-care strategies appropriately.

The conduct of trials involving thousands of participants, often in multiple countries around the world, requires a substantial research effort and can be very costly. CTSU has been successful in receiving large research grants from government (e.g. National Institute for Health Research, Medical Research Council) and charitable sources (e.g. British Heart Foundation), as well as from the pharmaceutical industry. Given the costs involved in running large studies, industry funding helps to ensure that the trials are of sufficient size and scope to assess the safety and efficacy of treatments reliably.

In order to ensure that CTSU's research is conducted independently of all sources of funding, we have had a number of measures in place for many years:

CTSU policy on consultancies, honoraria or other financial benefits

CTSU has had an explicit policy for about 30 years of not accepting any personal payments directly or indirectly from industry. It only seeks reimbursement to the University of Oxford of the costs of travel and accommodation to participate in scientific meetings.¹ This approach ensures that decisions to give lectures or advice are determined by the scientific value of doing so, and not by personal gain.

CTSU staff are also advised that it is generally not appropriate to have shares in tobacco, alcohol, drug or biotechnology companies which might be directly or indirectly affected by CTSU publications or public statements.¹ (Professor Rory Collins holds no shares in any company.)

Obtaining funding for CTSU research

CTSU decides what trials it wants to do for scientific reasons and then tries to persuade industry and non-industry sources to cover the costs. For example, for our Heart Protection Study, it took several years to obtain the funding, with half coming from the UK Medical Research Council (government) and the British Heart Foundation (charity), one quarter from Merck (supplier of simvastatin) and one quarter from Roche (supplier of vitamins E, C and beta-carotene). This trial showed that statin therapy reduced the risk of heart attacks and strokes safely for a wide range of patients at high-risk of such events, but the vitamins produced no benefit; both results were published alongside each other.^{2,3}

In other instances, CTSU has obtained all of the necessary funding from industry for trials intended to address important public health questions that would not otherwise be answered. For example, our most recently completed THRIVE trial (funded by Merck) showed that niacin – which has been used routinely for over 50 years to modify cholesterol – did not reduce the risk of heart attacks and strokes, but did increase the risk of other serious adverse events (such as diabetes, bleeding and infections). As soon as those findings emerged, they were provided to regulatory authorities, presented publicly, and will soon be published prominently with detailed tabulations of adverse events.⁴

All of CTSU's research that receives industry funding is governed by University of Oxford contracts which protect our independence in the way that we design, conduct, analyse, interpret and report the results of the research. Indeed, in the case of all of our clinical trials, CTSU (not the funders) hold the databases and control all of the analyses, with no restrictions on what is reported.

Cholesterol Treatment Trialists' (CTT) Collaborative meta-analyses of statin trials

This collaborative effort was established in the early 1990s.⁵ It is coordinated jointly by the CTSU in Oxford and the National Health and Medical Research Council Clinical Trials Centre (CTC) in Sydney, with the database held securely and analysed in both locations. It is <u>not</u> funded by the pharmaceutical industry; instead, funding has been provided by government (Australian National Health and Medical Research Council, European Community Biomed Programme) and charities (Australian National Heart Foundation, British Heart Foundation, Cancer Research UK).

The aim of the CTT collaboration was to bring together data for each patient in all of the large-scale randomised controlled trials of statin therapy in order to be able to assess – more reliably than was possible with any of the individual trials – the effects of cholesterol-lowering statin therapy on heart attacks, strokes, revascularisation procedures and vascular deaths among different types of patient (for example, men and women; young and old; higher and lower risk).⁵ The CTT collaboration also aimed to assess any effects of statins on other types of death and on site-specific cancers.

The academic investigators who ran the statin trials, and the funders of those trials, agreed to provide the required data on major vascular events, cause-specific mortality and site-specific cancer on the understanding that their data would not be released to third parties without their explicit permission. (Note: Data on all other adverse events – including muscle aching or myopathy – were not sought, and so the CTT collaboration does not currently have access to such data for meta-analyses.⁵)

Obtaining these individual patient data from each of the trials, running careful consistency checks of them, conducting extensive analyses of the collective database (involving about 170,000 patients in 27 trials) and reporting the results in a series of papers has been a substantial collaborative effort.⁶⁻⁸ As a consequence of this effort, the CTT collaboration has provided information that would not have otherwise have emerged about the effects of statins on these outcomes for different types of patient which has contributed considerably to helping doctors to use statins more appropriately.

CTSU patent for statin-related myopathy genetic test

CTSU's SEARCH trial (funded by Merck and governed by an Oxford University contract, as described above) was set up to determine whether more intensive lowering of cholesterol levels with 80mg daily simvastatin might be better than a standard 20mg daily dose. We found that the higher dose caused more cases of "myopathy" (defined as muscle symptoms plus substantial elevation of blood levels of the muscle enzyme creatine kinase), and also identified a genetic variant which predisposed patients to an increased risk of myopathy. This information was reported prominently.^{9,10}

In order to ensure that a test for this genetic variant was made available to patients, the University took out a patent on this finding and licensed it to Boston Heart Diagnostics who provide the test. Consistent with CTSU's approach to personal payments (see above), its staff waived their rights to any income from this discovery, which has instead been made available to support research.

Awards to CTSU for contributions to medical research and public health

From time to time, CTSU scientists are awarded medals or prizes by scientific societies, charitable foundations or industry. In accordance with CTSU's policy (see above), no money associated with awards from industry is accepted by staff. For example, in 2004, Rory Collins was awarded the Pfizer Medal which came with a £50,000 unrestricted research grant to Oxford University. (Professor Collins has not received other research funding from Pfizer, although he did serve unpaid on the Steering Committee for the ASCOT trial of atorvastatin, which was coordinated by Imperial College, London.)

CTSU (1 July 2014)

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- 1. <u>http://www.ctsu.ox.ac.uk/about-ctsu/documents/guidelines</u>
- 2. Heart Protection Study Collaborative Group. MRC/BHF Heart Protection Study of cholesterol lowering with simvastatin in 20,536 high-risk individuals: a randomised placebo-controlled trial. Lancet. 2002; 360: 7-22.
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- 10. Link E, Parish S, Armitage J, Bowman L, Heath S, Matsuda F, et al. SLCO1B1 variants and statin-induced myopathy--a genomewide study. N Engl J Med. 2008; 359: 789-99.



Instructions

The purpose of this form is to provide readers of your manuscript with information about your other interests that could influence how they receive and understand your work. The form is designed to be completed electronically and stored electronically. It contains programming that allows appropriate data display. Each author should submit a separate form and is responsible for the accuracy and completeness of the submitted information. The form is in six parts.

Identifying information.

The work under consideration for publication.

This section asks for information about the work that you have submitted for publication. The time frame for this reporting is that of the work itself, from the initial conception and planning to the present. The requested information is about resources that you received, either directly or indirectly (via your institution), to enable you to complete the work. Checking "No" means that you did the work without receiving any financial support from any third party -- that is, the work was supported by funds from the same institution that pays your salary and that institution did not receive third-party funds with which to pay you. If you or your institution received funds from a third party to support the work, such as a government granting agency, charitable foundation or commercial sponsor, check "Yes".

Relevant financial activities outside the submitted work.

This section asks about your financial relationships with entities in the bio-medical arena that could be perceived to influence, or that give the appearance of potentially influencing, what you wrote in the submitted work. You should disclose interactions with ANY entity that could be considered broadly relevant to the work. For example, if your article is about testing an epidermal growth factor receptor (EGFR) antagonist in lung cancer, you should report all associations with entities pursuing diagnostic or therapeutic strategies in cancer in general, not just in the area of EGFR or lung cancer.

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Intellectual Property.

This section asks about patents and copyrights, whether pending, issued, licensed and/or receiving royalties.

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4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Dr. Sundstrom
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Takeda				\checkmark	Consultancy agreement; honoraria paid to institution
Novartis					Consultancy agreement; honoraria paid to institution
Merck Darmstadt AG	\checkmark				research grant; paid to organisation
BMS				\checkmark	Consultancy agreement; honoraria paid to institution
Hemocue				\checkmark	Consultancy agreemnen; honararia paid to institution
Merck Sharpe & Dohme				\checkmark	Consultancy agreement; honararia paid to institution
Fresenius				\checkmark	Consultancy agreement; honararia paid to Institution
Eli Lilly					Consultancy agreement; honararia paid to Institution

Section 4.

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Yes

V No



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Relevant financial activities outside the submitted work.

This section asks about your financial relationships with entities in the bio-medical arena that could be perceived to influence, or that give the appearance of potentially influencing, what you wrote in the submitted work. You should disclose interactions with ANY entity that could be considered broadly relevant to the work. For example, if your article is about testing an epidermal growth factor receptor (EGFR) antagonist in lung cancer, you should report all associations with entities pursuing diagnostic or therapeutic strategies in cancer in general, not just in the area of EGFR or lung cancer.

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Intellectual Property.

This section asks about patents and copyrights, whether pending, issued, licensed and/or receiving royalties.

Relationships not covered above.

Use this section to report other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work.

Definitions.

Entity: government agency, foundation, commercial sponsor, academic institution, etc.

Grant: A grant from an entity, generally [but not always] paid to your organization

Personal Fees: Monies paid to you for services rendered, generally honoraria, royalties, or fees for consulting , lectures, speakers bureaus, expert testimony, employment, or other affiliations

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Other: Anything not covered under the previous three boxes Pending: The patent has been filed but not issued Issued: The patent has been issued by the agency Licensed: The patent has been licensed to an entity, whether earning royalties or not Penulties: Fund earn agencies in to use a reput institution due to use

Royalties: Funds are coming in to you or your institution due to your patent



and the second se	ntifying Inforr	nation	
1. Given Name (First Nar Raymond	ne)	2. Surname (Last Name) Estacio	3. Date 20-January-2015
4. Are you the correspor	nding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundström
5. Manuscript Title Effects of Blood Pressu	ire Reduction in	Mild Hypertension: A Syste	ematic Review and Meta-analysis
6. Manuscript Identifying M14-0773	g Number (if you k	now it)	
Section 2. The	Work Under C	onsideration for Publi	cation
	ted work (includin	g but not limited to grants, d	n a third party (government, commercial, private foundation, etc.) for ata monitoring board, study design, manuscript preparation,
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Section 5. Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

Yes, the following relationships/conditions/circumstances are present (explain below):

No other relationships/conditions/circumstances that present a potential conflict of interest

At the time of manuscript acceptance, journals will ask authors to confirm and, if necessary, update their disclosure statements. On occasion, journals may ask authors to disclose further information about reported relationships.

Section 6. Disclosure Statement

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Evaluation and Feedback

Please visit http://www.icmje.org/cgi-bin/feedback to provide feedback on your experience with completing this form.



Instructions

The purpose of this form is to provide readers of your manuscript with information about your other interests that could influence how they receive and understand your work. The form is designed to be completed electronically and stored electronically. It contains programming that allows appropriate data display. Each author should submit a separate form and is responsible for the accuracy and completeness of the submitted information. The form is in six parts.

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Royalties: Funds are coming in to you or your institution due to your patent



Section 1. Identifying Info	rmation	
1. Given Name (First Name) Robert	2. Surname (Last Name) Fagard	3. Date 12-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundstrom
5. Manuscript Title Effects of blood pressure reduction in	n mild hypertension: A syster	matic review and meta-analysis
6. Manuscript Identifying Number (if you -	know it)	
Sale Statistication and an and a set		
Section 2. The Work Under	Consideration for Public	cation
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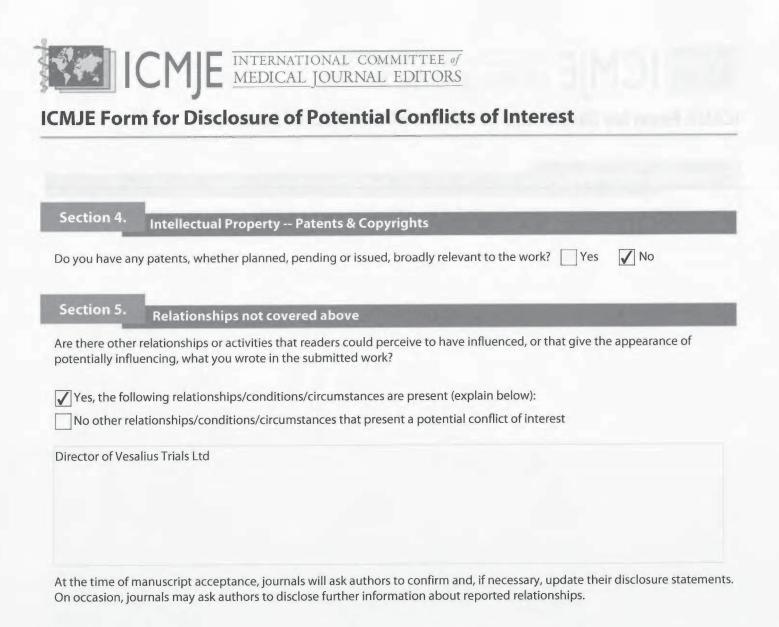
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Royalties: Funds are coming in to you or your institution due to your patent



Section 1. Identifying Inf	ormation	760			The second particular second
1. Given Name (First Name) Kim	2. Surnar Fox	ne (Last Na	me)		3. Date 17-December-2014
4. Are you the corresponding author?	Yes	No No	Correspon Dr Sundst	-	nor's Name
5. Manuscript Title Effects of Blood Pressure Reductior	a in Mild Hyper	tension: A	systematic Revie	w and M	eta-Analysis
6. Manuscript Identifying Number (if y	ou know it)				
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Name of Entity	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments
rvier					Executive Committee CLARIFY, MODIFY, Consultancy to attend EMA, lectures, editorial work (Dialogues and Medicographia) travel expenses
tra Zeneca					Executive Committee Themis Study
JrX		\checkmark			DSMB Committee
ngo			\checkmark		Travel expenses to attend SAB
oadview Ventures					Travel expenses and fees to attend SAB
llAegis		\checkmark			SAB



Section 6. Disclosure Statement

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Professor Fox reports personal fees and non financial support from Servier, professional fees from TaurX and AstraZeneca. He is director of Vesalius Trials Ltd



Evaluation and Feedback

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ICMJE Uniform Disclosure Form for Potential Conflicts of Interest

INSTRUCTIONS:

The purpose of this form is to provide readers of your manuscript with information about your other interests that could influence how they receive and understand your work. The form has five parts.

1. Identifying information.

Each author should submit a separate form. Provide complete information and double-check the manuscript number. If you are NOT the corresponding author please insert his or her name.

2. The work under consideration for publication.

Please provide information about the work that you have submitted for publication. The time frame for this reporting is that of the work itself, from the initial conception and planning to the present. The idea is to provide for the reader information about resources that you received, either directly or indirectly (via your institution), to enable you to complete the work. If you check the "No" box it means that you did the work without receiving any financial support from any third party -- that is, the work was supported by funds from the same institution that pays your salary and that institution did not receive third-party funds to pay you. If you or your institution did receive funds from a third party to support the work, check "Yes" along with the appropriate boxes to indicate the type of support and whether you or your institution received it.

3. Relevant financial activities outside the submitted work.

Please report all sources of revenue relevant to the submitted work that accrued either directly to you or were paid to your institution on your behalf over the 36 months prior to submission of the work. This should include all monies from sources with relevance to the submitted work, not just monies from the entity that sponsored the research. If there is any question, it is usually better to disclose a relationship than not to do so. Please note that your interactions with the work's sponsor outside the submitted work should be listed here. For each category list each entity on a separate line. Use as many lines as necessary to provide complete information. In addition, please disclose relationships that fall outside the 36-month window that readers may want to know about and could reasonably criticize you for not disclosing (for example, long-term financial relationships that are now ended).

The goal of this section is to provide information for our reviewers and readers about your interactions with entities in the biomedical arena that could be perceived to influence, or that give the appearance of potentially influencing, what you wrote in the submitted work. You should disclose interactions with ANY entity that could be considered broadly relevant to the work. For example, if your article is about testing an epidermal growth factor receptor (EGFR) antagonist in lung cancer, you should report all associations with entities pursuing diagnostic or therapeutic strategies in cancer in general, not just in the area of EGFR or lung cancer. For grants you have received for work outside the submitted work, you should disclose support ONLY from entities that could be perceived to benefit financially from the published work, such as drug companies, or foundations supported by entities that could be perceived to have a financial stake in the outcome. Public funding sources, such as the NIH or the MRC, need not be disclosed. For example, if the NIH sponsored a piece of work you have been involved in but drugs were provided by a pharmaceutical company, you need only list the pharmaceutical company.

4. Financial relationships involving your spouse or partner or your children (under 18 years of age).

If monies from the types of relationships listed in Section 3 were paid to your spouse or partner or dependent children, please list the type of activity and source of the money.

5. Nonfinancial associations.

Please report any personal, professional, political, institutional, religious, or other associations that a reasonable reader would want to know about in relation to the submitted work.



ICMJE Uniform Disclosure Form for Potential Conflicts of Interest

Section 1. Identifying Information.

Given Name: (or first)	John	Surname: (or last)	Kostis	Effective Date:	16-December-2014
Are you the	e corresponding a	author? 🗌 Yes 🛛 N	lo	Format exam	ple: 07-August-2008
Correspondin	g author's name	Johan Sundstrom			
Manuscript T	itle: Effects of B	lood Pressure Reductior	in Mild Hypertens	sion: A Systematic Review and Meta-	-analysis"
Manuscript Id	lentifying Num	ber (if you know it):			

Section 2. Information about the support of the work under consideration for publication.

Did you or your institution at any time receive payment or support in kind for any aspect of the submitted work (including but not limited to grants, data monitoring board, study design, manuscript preparation, statistical analysis, etc...)?

X No

Yes, specify nature of compensation

Section 3. Information about relevant financial relationships outside the submitted work.

Place a check in the appropriate boxes in the table to indicate whether you have financial relationships (regardless of amount of compensation) with any entities that have an interest related to the submitted work. Use one line for each entity; add as many lines as you need. Use the comments column to indicate any additional information that you think a reader or editor would want to know about the compensation. Report relationships that were present during the 36 months prior to submission. In addition please disclose relationships that fall outside the 36-month window that readers may want to know about and could reasonably criticize you for not disclosing (for example, long-term financial relationships that are now ended).

If you have more than one relationship, click "Add +" to add a row. Click "Del ×" to delete an extra row.

Type of Relationship (in alphabetical order)	No	Money Paid to You	Money to Your institution	Entity	Comments	
Board membership	\boxtimes					Del ×
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Consultancy		\boxtimes		St. Jude Medical	DSMB	Del ×
Consultancy				Merck	Advisory Board	Del ×
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Employment	\boxtimes					Del ×
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Type of Relationship (in alphabetical order)	No	Money Paid to You	Money to Your institution	Entity	Comments	
Expert testimony						Del ×
						Add +
Gifts				line de		Del ×
						Add +
Grants/grants pending				Novartis		Del ×
Grants/grants pending				Sanofi		Del ×
Grants/grants pending				Pfizer		Del ×
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Honoraria						Del ×
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Payment for manuscript preparation	\boxtimes					Del ×
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Patents (planned, pending or issued)						Del ×
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Royalties	\boxtimes					Del ×
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Payment for development of educational presentations including service on speakers' bureaus				BM5/Pfizer	speaker	Del ×
Payment for development of educational presentations including service on speakers' bureaus				Sanofi-Aventis		Del ×
						Add +
Stock/stock options	\boxtimes					Del ×
						Add +
Fravel/accommodations expenses covered or eimbursed						Del ×
						Add +



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Other (err on the side of full disclosure)						Del ×
						Add +

Section 4. Information about financial relationships involving your spouse or partner or your children (under 18 years of age).

Do your children or your spouse or partner have financial relationships with entities that have an interest in the content of the submitted work?

X No other relationships/conditions/circumstances that present potential conflict of interest

Yes, the following relationships/conditions/circumstances are present (explain below):

Section 5. Information about relevant nonfinancial associations.

Do you have any relevant nonfinancial associations or interests (personal, professional, political, institutional, religious, or other) that a reasonable reader would want to know about in relation to the submitted work?

X No relevant nonfinancial relationships/conditions/circumstances to report.

Yes, the following relevant nonfinancial relationships/conditions/circumstances are present (explain below):

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying I	Information	1.1.2			ALL STATE STATES	
1. Given Name (First Name) Bertram	2. Surna Pitt	me (Last Nai	me)		3. Date 10-December-2014	
4. Are you the corresponding autho	r? Yes	✓ No		Corresponding Author's Name Sundstrom		
5. Manuscript Title Effects of blood pressure reduction	on in mild hyper	tension : a :	systemic review a	nd meta	analysis	
6. Manuscript Identifying Number (ii	f you know it)					
Section 2. The Work Un	der Considera	tion for P	ublication	a ta at	the second	
Place a check in the appropriate k of compensation) with entities as	ncial activities boxes in the table described in the uld report relation f interest?	to indicat instruction onships tha	ns. Use one line fo	ive financ or each ei	cial relationships (regardless of amo ntity; add as many lines as you nee e 36 months prior to publication	
lame of Entity	Grant?	Personal Fees ?	Non-Financial Support?	Other?	Comments	
er					consultant	
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a zeneca		\checkmark			consultant	
psa		\checkmark		hannand		
harmaceuticals		From the second			consultant and stock options	
		\checkmark			consultant and stock options consultant and stock options	
alth peptides		\checkmark				

ICMJE Form for Disclosure of Potential Conflicts of Interest

Name of Entity	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments
novartis					member DMC
oxygen biotherapeutics		\checkmark			member DMC
juventis		\checkmark			member events committee
DaVinci therapeutics		\checkmark			stock options
aura sense		\checkmark			consultant, stock options

Section 4.

Section 5.

Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes No If yes, please fill out the appropriate information below. If you have more than one entity press the "ADD" button to add a row. Excess rows can be removed by pressing the "X" button.

Patent	Pending?	Issued ²	Licensed?	Royalties ?	Licensee?	Comments	
site specific deliver y of eplerenone to the myocardium							
to the myocardium							

Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

Yes, the following relationships/conditions/circumstances are present (explain below):

No other relationships/conditions/circumstances that present a potential conflict of interest

At the time of manuscript acceptance, journals will ask authors to confirm and, if necessary, update their disclosure statements. On occasion, journals may ask authors to disclose further information about reported relationships.



Section 6. Di

Disclosure Statement

Based on the above disclosures, this form will automatically generate a disclosure statement, which will appear in the box below.

Dr. Pitt reports personal fees from pfizer, personal fees from bayer, personal fees from astra zeneca, personal fees from relypsa, personal fees from scpharmaceuticals, personal fees from stealth peptides, personal fees from j&J, personal fees from novartis, personal fees from oxygen biotherapeutics, personal fees from juventis, personal fees from DaVinci therapeutics, personal fees from aura sense, outside the submitted work; In addition, Dr. Pitt has a patent site specific deliver y of eplerenone to the myocardium pending.

Evaluation and Feedback



Instructions

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Royalties: Funds are coming in to you or your institution due to your patent



			the second s	
1. Given Name (First Name) Neil	2. Surname (Last Nam Poulter	ne)	3. Date 21-January-2015	
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Autho Johan Sundström	r's Name	
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A S	Systematic Review and Me	a-analysis	
6. Manuscript Identifying Number (if you M14-0773	know it)			
Section 2. The Work Under	Consideration for Pu	ublication		-
Did you or your institution at any time rec any aspect of the submitted work (includir statistical analysis, etc.)? Are there any relevant conflicts of inte	ng but not limited to grant	ts, data monitoring board, stu		
Section 3. Relevant financia	l activities outside t	he submitted work.		197
Place a check in the appropriate boxes		whether you have financ	al relationshins (regardles	
		s. Use one line for each en	tity; add as many lines as y	you need by
licking the "Add +" box. You should re Are there any relevant conflicts of inte	eport relationships that rest? 🖌 Yes 🗌 N	s. Use one line for each en	tity; add as many lines as y	you need by
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Section 4. Intellectua	l Property Patents & Copyrights
Do you have any patents, whe	ther planned, pending or issued, broadly relevant to the work? 🗌 Yes 📝 No
Section 5	
Section 5. Relationsh	nips not covered above
	or activities that readers could perceive to have influenced, or that give the appearance of ou wrote in the submitted work?
Yes, the following relations	ships/conditions/circumstances are present (explain below):
No other relationships/con	ditions/circumstances that present a potential conflict of interest
Chairman of the BHS Guideline Member of Council of ISH	es and Information Service Working Party

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Dr. Poulter reports grants from National Institute for Health Research Health Technology Assessment, grants from Diabetes UK, grants from British Hypertension Society, grants from Servier, grants from Pfizer, grants from Menorini, personal fees from Servier, personal fees from Menorini, from null, outside the submitted work; and Chairman of the BHS Guidelines and Information Service Working Party Member of Council of ISH.



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Remuzzi

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information							
1. Given Name (First Name)2. Surname (Last Name)3. DateGiuseppeRemuzzi12-December	-2014						
4. Are you the corresponding author? Yes ✓ No Corresponding Author's Name Johan Sundström							
5. Manuscript Title Effects of Blood Pressure Reduction in Mild Hypertension: A Systematic Review and Meta-analysis							
6. Manuscript Identifying Number (if you know it)							
Section 2. The Work Under Consideration for Publication							
Did you or your institution at any time receive payment or services from a third party (government, commercial, private any aspect of the submitted work (including but not limited to grants, data monitoring board, study design, manuscript statistical analysis, etc.)?							
Are there any relevant conflicts of interest? 🗌 Yes 🖌 No							
	1000 State 2						
Section 3. Relevant financial activities outside the submitted work.							
Place a check in the appropriate boxes in the table to indicate whether you have financial relationships (regardless of amount of compensation) with entities as described in the instructions. Use one line for each entity; add as many lines as you need by clicking the "Add +" box. You should report relationships that were present during the 36 months prior to publication .							
Are there any relevant conflicts of interest? \checkmark Yes \square No	•						
If yes, please fill out the appropriate information below.							
Name of Entity Grant? Personal Non-Financial Other? Comments							
AbbVie Consultancy agreemen remuneration is accept							

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Consultancy agreement. No personal remuneration is accepted, compensations are paid to his institution for research and educational activities.

Remuzzi

Alexion Pharmaceuticals

ICMJE Form for Disclosure of Potential Conflicts of Interest

Name of Entity	Grant?	Personal Fees [?]	Non-Financial Support?	Other?	Comments	
Bayer Healthcare				I	Consultancy agreement. No personal remuneration is accepted, compensations are paid to his institution for research and educational activities.	
Reata Pharmaceuticals				\checkmark	Consultancy agreement. No personal remuneration is accepted, compensations are paid to his institution for research and educational activities.	
Novartis Pharma				\checkmark	Consultancy agreement. No personal remuneration is accepted, compensations are paid to his Institution for research and educational activities.	
AstraZeneca				\checkmark	Consultancy agreement. No personal remuneration is accepted, compensations are paid to his institution for research and educational activities.	
Otsuka Pharmaceutical Europe				\checkmark	Consultancy agreement. No personal remuneration is accepted, compensations are paid to his Institution for research and educational activities.	
*, Concert Pharmaceuticals				1	Consultancy agreement. No personal remuneration is accepted, compensations are paid to his institution for research and educational activities.	

Section 4. Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work?

✓ No



Section 5.

Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

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Section 1. Identifying Inform	nation	
1. Given Name (First Name) Anthony	2. Surname (Last Name) Rodgers	3. Date 12/12/2014
4. Are you the corresponding author?	Yes No	
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Section 2. The Work Under C	onsideration for Publication	
	ive payment or services from a third party (governm) but not limited to grants, data monitoring board, si	
Are there any relevant conflicts of intere	est? Yes 🖌 No	ADD
Section 3. Relevant financial	activities outside the submitted work.	
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Section 4. Intellectual Proper	rty Patents & Copyrights	
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Section 5. Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

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Generate Disclosure Statement

Dr. Rodgers has nothing to disclose.

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1. Given Name (First Name) Rury	2. Surname (Last Name) Holman	3. Date 12-January-2015
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundstro¨m,
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syst	ematic Review and Meta-analysis
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and the second		
id you or your institution at any time rec ny aspect of the submitted work (includir ratistical analysis, etc.)?	ig but not limited to grants, d	cation a a third party (government, commercial, private foundation, etc.) for ata monitoring board, study design, manuscript preparation,
bid you or your institution at any time rec ny aspect of the submitted work (includir tatistical analysis, etc.)? are there any relevant conflicts of inte	eive payment or services fron 1g but not limited to grants, d	n a third party (government, commercial, private foundation, etc.) f ata monitoring board, study design, manuscript preparation,
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Section 5. Relationships not covered above

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Dr. Holman has nothing to disclose.

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Are you the corresponding author?	Yes Vo	Corresponding Author's Name Johan Sundström
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Royalties: Funds are coming in to you or your institution due to your patent



1. Given Name (First Name) Stephan	2. Surname (Last Name Lueders	e)	3. Date 29-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's N Sundström	ame
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Sy	vstematic Review and Meta-ar	nalysis
5. Manuscript Identifying Number (if you l	know it)		
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Dr. Lueders has nothing to disclose.

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Royalties: Funds are coming in to you or your institution due to your patent



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1. Given Name (First Name) GIUSEPPE	2. Surname (Last Name) MANCIA		3. Date 11-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Aut SUNDSTRŌM	hor's Name
5. Manuscript Title EFFECTS OF BLOOD PRESSURE REDUC	CTION IN MILD HYPERTENSI	ON: A SYSTEMATIC R	EVIEW AND META-ANALYSIS
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Section 4.

Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes V No

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5.



1. Given Name (First Name) Steven	2. Surname (Last Name) Nissen		3. Date 17-December-2014
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5. Manuscript Title Effects of Blood Pressure			
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patent



1. Given Name (First Name) Toshio	2. Surname (Last Nam Ogihara	e) 3. Date 12-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name
i. Manuscript Title ffects of Blood Pressure Reduction in	Mild Hypertension: A Sy	vstematic Review and Meta-analysis
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Section 5.

Relationships not covered above

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Dr. Ogihara has nothing to disclose.

Evaluation and Feedback



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1. Given Name (First Name) Takayoshi	2. Surname (Last Name) Ohkubo		3. Date 12-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Sundström	Name
5. Manuscript Title Effects of Blood Pressure Reduction	in Mild Hypertension: A Syste	ematic Review and Meta-	analysis
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Section 4. Intellectual Prop Do you have any patents, whether pl			rk? Yes 🖌 No



Section 5. Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

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Dr. Ohkubo reports personal fees from Dainihon-Sumitomo Pharm., personal fees from Daiichi-Sankyo Pharm., outside the submitted work; .

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Cytori

Fujisawa HealthCare Inc.

Research grant

Research grant



Name of Entity	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments
Gilead Sciences Inc.	\checkmark				Research grant
Ikaria Development Subsidiary One LLC	\checkmark				Research grant
InfraReDx	\checkmark				Research grant
inVentive Health Clinical LLC	\checkmark				Research grant
Lilly/Cleveland Clinic DSMB member for a phase 2 efficacy and safety study of Ly2484595					consultant
Medtelligence		\checkmark			consultant
NHLBI DSMB Chair for FREEDOM trial		\checkmark			consultant
NHLBI Study Section for Progenitor Cell Biology Consortium					consultant
NHLBI/NCRR CTSA	\checkmark				Grant
NIH Study Section for Cardiovascular Sciences Small Business Activities		\checkmark			consultant
NIH/NHLBI	\checkmark				Research grants
Park-Davis	\checkmark				Educational grant
Pfizer	\checkmark				Educational grant
Sanofi-Aventis	\checkmark				Educational grant

Section 4.

Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work?

Section 5. Relationships not covered above

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V No



Section 6. Disclosure Statement

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Dr. Pepine reports grants from AHA, grants from Amorcyte/Neostem, grants from Astra/Zeneca, grants from Baxter, grants from Brigham & Women's Hospital, grants from Capricor, grants from Cytori, grants from Fujisawa HealthCare Inc., grants from Gilead Sciences Inc., grants from Ikaria Development Subsidiary One LLC, grants from InfraReDx, grants from inVentive Health Clinical LLC, personal fees from Lilly/Cleveland Clinic DSMB member for a phase 2 efficacy and safety study of Ly2484595, personal fees from Medtelligence, personal fees from NHLBI DSMB Chair for FREEDOM trial, from NHLBI Study Section for Progenitor Cell Biology Consortium, grants from NHLBI/NCRR CTSA, personal fees from NIH Study Section for Cardiovascular Sciences Small Business Activities, grants from NIH/NHLBI, grants from Park-Davis, grants from Pfizer, grants from Sanofi-Aventis, outside the submitted work; .

Evaluation and Feedback



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1. Given Name (First Name) Marc	2. Surname (Last Name) Pfeffer		3. Date 15-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Aut Johan Sundstrom	
5. Manuscript Title Effect of Blood Pressure Reduction in	Mild Hypertension: Syster	nic Review and Meta-	Analysis.
6. Manuscript Identifying Number (if you	ı know it)		
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Name of Entity	Grant?	Personal Fees?	Non-Financial Support [?]	Other?	Comments
Abbott Vascular		\checkmark			Consulting fees.
Celladon					Institutional agreement to the Brigham and Women's Hospital.
Cleveland Clinic		\checkmark			Consulting fees.
Concert		\checkmark			Consulting fees.
Hamilton Health Sciences	\checkmark				Institutional agreement to the Brigham and Women's Hospital. Not considered relevant.
FibroGen					Consulting fees.
Daiichi Sankyo					Consulting fees.
Medtronic					Consulting fees.
Merck		\checkmark			Consulting fees.
Novartis	\checkmark	\checkmark			Institutional agreement to the Brigham and Women's Hospital and Consulting fees.
Roche		\checkmark			Consulting fees.
Sanofi Aventis					Institutional agreement to the Brigham and Women's Hospital. Not considered relevant.
Servier					Consulting fees.
Teva					Consulting fees.
Genzyme		\checkmark			Consulting fees.
Novo Nordisk		\checkmark			Consulting fees.
Salix		\checkmark			Consulting fees.
Sanderling		\checkmark			Consulting fees.
Relypsa		\checkmark			Consulting fees.

Section 4.

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Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Ves No If yes, please fill out the appropriate information below. If you have more than one entity press the "ADD" button to add a row. Excess rows can be removed by pressing the "X" button.

ICMJE INTERNATIONAL COMMITTEE of MEDICAL JOURNAL EDITORS

ICMJE Form for Disclosure of Potential Conflicts of Interest

Patent?	Pending?	Issued?	Licensed?	Royalties?	Licensee?	Comments
Patent?	Pending?	Issued?	Licensed?	Royalties?	Licensee?	Comments
ovartis. The Brigham and Women's ospital has patents for the use of hibitors of the RAS in selected rvivors of MI with Novartis Dr. effer is a co-inventor. His share of e licensing agreement is evocably transferred to charity.			V			Not considered relevant.

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1. Given Name (First Name) Mahboob	2. Surname (Last Name) Rahman	3. Date 22-December-2104
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Sundström
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syste	matic Review and Meta-analysis
6. Manuscript Identifying Number (if you l	know it)	
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1. Given Name (First Name) Piero	2. Surname (Last Name) Ruggenenti	3. Date 02-February-2015
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundström
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Sys	tematic Review and Meta-analysis
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Dr. Ruggenenti has nothing to disclose.

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1. Given Name (First Name)	2. Surname (Last Name)		3. Date
ТАКАО	SARUTA		29-January-2015
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Nar Johan Sundström	me
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ICMJE INTERNATIONAL COMMITTEE of MEDICAL JOURNAL EDITORS

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1. Given Name (First Name) Robert	2. Surname (Last Name) Schrier	3. Date 15-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundstrom
5. Manuscript Title Effects of blood pressure reduction in	mild hypertension: A syste	matic review and meta-analysis
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Royalties: Funds are coming in to you or your institution due to your patent



1. Given Name (First Name)	2. Surname (Last Name)	3. Date
Peter S	Sever	23-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Dr. Sundström
5. Manuscript Title Effects of Blood Pressure Reduction i	n Mild Hypertension: A Syste	ematic Review and Meta-analysis
6. Manuscript ldentifying Number (if you M14-0773	know it)	
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1. Given Name (First Name) Peter	2. Surname (Last Name) Sleight		3. Date 23-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Nam Johan Sundström	ne
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syste	matic Review and Meta-anal	lysis
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1. Given Name (First Name) Jan A.	2. Surname (Last Name) Staessen	3. Date 21-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name J Sundström
i. Manuscript Title Effects of Blood Pressure Reduction ir	Mild Hypertension: A Syste	matic Review and Meta-analysis
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1



1. Given Name (First Name) Koon	2. Surname (Last Name) Teo	3. Date 12-December-2014
4. Are you the corresponding author?	Yes Vo	Corresponding Author's Name Johan Sundstrom
5. Manuscript Title Effects of Blood Pressure Reduction in	n Mild Hypertension: a syste	matic review and meta-analysis
6. Manuscript Identifying Number (if you	know it)	
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1. Given Name (First Name) JIGUANG	2. Surname (Last Name) WANG		3. Date 26-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Na Dr. Sundström	ame
5. Manuscript Title Effects of Blood Pressure Reduction ir	Mild Hypertension: A Sys	ematic Review and Meta-an	alysis
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Other: Anything not covered under the previous three boxes Pending: The patent has been filed but not issued Issued: The patent has been issued by the agency Licensed: The patent has been licensed to an entity, whether earning royalties or not Royalties: Funds are coming in to you or your institution due to your patent

1



. Given Name (First Name) Paul	2. Surname (Last Name Whelton) 3. Date 16-January-2015
. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan Sundström
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1. Given Name (First Name) Lindon	2. Surname (Last Name) Wing	3. Date 01-July-2105
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Johan SUNDSTROM
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1. Given Name (First Name) Yoshiki	2. Surname (Last Name) Yui	3. Date 12-December-2014	
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Dr Sundstrom	
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Sys	tematic Review and Meta-analysis	
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1. Given Name (First Name) Salim	2. Surname (Last Name) Yusuf	3. Date 15-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Sundstrom
5. Manuscript Title Effects of Blood Pressure Reduction	in Mild Hypertension: A Syste	matic Review and Meta-analysis
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Section 5.

Relationships not covered above

Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?

Yes, the following relationships/conditions/circumstances are present (explain below):

No other relationships/conditions/circumstances that present a potential conflict of interest

At the time of manuscript acceptance, journals will ask authors to confirm and, if necessary, update their disclosure statements. On occasion, journals may ask authors to disclose further information about reported relationships.

Section 6.

Disclosure Statement

Based on the above disclosures, this form will automatically generate a disclosure statement, which will appear in the box below.

Dr. Yusuf has nothing to disclose.

Evaluation and Feedback



Instructions

The purpose of this form is to provide readers of your manuscript with information about your other interests that could influence how they receive and understand your work. The form is designed to be completed electronically and stored electronically. It contains programming that allows appropriate data display. Each author should submit a separate form and is responsible for the accuracy and completeness of the submitted information. The form is in six parts.

Identifying information.

The work under consideration for publication.

This section asks for information about the work that you have submitted for publication. The time frame for this reporting is that of the work itself, from the initial conception and planning to the present. The requested information is about resources that you received, either directly or indirectly (via your institution), to enable you to complete the work. Checking "No" means that you did the work without receiving any financial support from any third party -- that is, the work was supported by funds from the same institution that pays your salary and that institution did not receive third-party funds with which to pay you. If you or your institution received funds from a third party to support the work, such as a government granting agency, charitable foundation or commercial sponsor, check "Yes".

Relevant financial activities outside the submitted work.

This section asks about your financial relationships with entities in the bio-medical arena that could be perceived to influence, or that give the appearance of potentially influencing, what you wrote in the submitted work. You should disclose interactions with ANY entity that could be considered broadly relevant to the work. For example, if your article is about testing an epidermal growth factor receptor (EGFR) antagonist in lung cancer, you should report all associations with entities pursuing diagnostic or therapeutic strategies in cancer in general, not just in the area of EGFR or lung cancer.

Report all sources of revenue paid (or promised to be paid) directly to you or your institution on your behalf over the 36 months prior to submission of the work. This should include all monies from sources with relevance to the submitted work, not just monies from the entity that sponsored the research. Please note that your interactions with the work's sponsor that are outside the submitted work should also be listed here. If there is any question, it is usually better to disclose a relationship than not to do so.

For grants you have received for work outside the submitted work, you should disclose support ONLY from entities that could be perceived to be affected financially by the published work, such as drug companies, or foundations supported by entities that could be perceived to have a financial stake in the outcome. Public funding sources, such as government agencies, charitable foundations or academic institutions, need not be disclosed. For example, if a government agency sponsored a study in which you have been involved and drugs were provided by a pharmaceutical company, you need only list the pharmaceutical company.

Intellectual Property.

This section asks about patents and copyrights, whether pending, issued, licensed and/or receiving royalties.

Relationships not covered above.

Use this section to report other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work.

Definitions.

Entity: government agency, foundation, commercial sponsor, academic institution, etc.

Grant: A grant from an entity, generally [but not always] paid to your organization

Personal Fees: Monies paid to you for services rendered, generally honoraria, royalties, or fees for consulting, lectures, speakers bureaus, expert testimony, employment, or other affiliations

Non-Financial Support: Examples include drugs/equipment supplied by the entity, travel paid by the entity, writing assistance, administrative support, etc.



1. Given Name (First Name) Alberto	2. Surname (Last Name) Zanchetti		3. Date 12-December-2014
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Auth Johan Sundstrom	nor's Name
5. Manuscript Title Effects of Blood Pressure Reduction in	Mild Hypertension: A Syste	ematic Review and M	eta-analysis
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Section 4. Intellectual Property -- Patents & Copyrights

Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes Ves



Section 5.

Relationships not covered above

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Section 6.

Disclosure Statement

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Dr. Zanchetti reports personal fees from Menarini International, personal fees from Recordati, personal fees from CRVx, outside the submitted work; .

Evaluation and Feedback