



ICMJE Form for Disclosure of Potential Conflicts of Interest

Instructions

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4. Intellectual Property.

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

5. Relationships not covered above.

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



ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Gordon	2. Surname (Last Name) Smith	3. Date 04-November-2019
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Samuel Klein
5. Manuscript Title Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease		
6. Manuscript Identifying Number (if you know it) 134165-JCI-CMED-1		

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Are there any relevant conflicts of interest? Yes No

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Dr. Smith has nothing to disclose.

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Section 1. Identifying Information

1. Given Name (First Name)
Mahalakshmi

2. Surname (Last Name)
Shankaran

3. Date
04-November-2019

4. Are you the corresponding author? Yes No

Corresponding Author's Name
Samuel Klein

5. Manuscript Title
Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease

6. Manuscript Identifying Number (if you know it)
134165-JCI-CMED-1

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Dr. Shankaran has nothing to disclose.

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1. Given Name (First Name) Mihoko

2. Surname (Last Name) Yoshino

3. Date 04-November-2019

4. Are you the corresponding author? Yes No Corresponding Author's Name Samuel Klein

5. Manuscript Title Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease

6. Manuscript Identifying Number (if you know it) 134165-JCI-CMED-1

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Dr. Yoshino has nothing to disclose.

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1. Given Name (First Name) George	2. Surname (Last Name) Schweitzer	3. Date 04-November-2019
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Samuel Klein
5. Manuscript Title Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease		
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Royalties: Funds are coming in to you or your institution due to your patent



ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Joseph	2. Surname (Last Name) Beals	3. Date 04-November-2019
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Samuel Klein
5. Manuscript Title Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease		
6. Manuscript Identifying Number (if you know it) 134165-JCI-CMED-1		

Section 2. The Work Under Consideration for Publication

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Are there any relevant conflicts of interest? Yes No

ADD

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Are there any relevant conflicts of interest? Yes No

ADD

Section 4. Intellectual Property -- Patents & Copyrights

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



ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 5. Relationships not covered above

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Dr. Beals has nothing to disclose.

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name)
Adewole

2. Surname (Last Name)
Okunade

3. Date
04-November-2019

4. Are you the corresponding author? Yes No

Corresponding Author's Name
Samuel Klein

5. Manuscript Title
Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease

6. Manuscript Identifying Number (if you know it)
134165-JCI-CMED-1

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Bruce	2. Surname (Last Name) Patterson	3. Date 04-November-2019
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Samuel Klein
5. Manuscript Title Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease		
6. Manuscript Identifying Number (if you know it) 134165-JCI-CMED-1		

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Edna	2. Surname (Last Name) Nyangau	3. Date 04-November-2019
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Samuel Klein
5. Manuscript Title Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease		
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Are there any relevant conflicts of interest? Yes No

ADD

Section 4. 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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ICMJE Form for Disclosure of Potential Conflicts of Interest

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1. Given Name (First Name) Claude	2. Surname (Last Name) Sirlin	3. Date 04-November-2019
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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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Royalties: Funds are coming in to you or your institution due to your patent



ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Saswata 2. Surname (Last Name) Talukdar 3. Date 04-November-2019

4. Are you the corresponding author? Yes No Corresponding Author's Name
Samuel Klein

5. Manuscript Title
Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease

6. Manuscript Identifying Number (if you know it)
134165-JCI-CMED-1

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.)?

Are there any relevant conflicts of interest? 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.

Name of Institution/Company	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments	
Merck & Co., Inc	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
						ADD

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 No

ADD

Section 4. Intellectual Property -- Patents & Copyrights

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



ICMJE Form for Disclosure of Potential Conflicts of Interest

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.

Generate Disclosure Statement

Dr. Talukdar reports personal fees from Merck & Co., Inc, during the conduct of the study; .

Evaluation and Feedback

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

ICMJE Form for Disclosure of 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 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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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name)
Marc

2. Surname (Last Name)
Hellerstein

3. Date
04-November-2019

4. Are you the corresponding author? Yes No

5. Manuscript Title
Insulin resistance drives hepatic de novo lipogenesis in nonalcoholic fatty liver disease

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Are there any relevant conflicts of interest? Yes No

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Are there any relevant conflicts of interest? Yes No

If yes, please fill out the appropriate information below.

Name of Entity	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments	
Gilead Sciences Inc.	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
Pfizer Inc	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
Synergenics LLC	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
						ADD

Section 4. Intellectual Property -- Patents & Copyrights

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



ICMJE Form for Disclosure of Potential Conflicts of Interest

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Dr. Hellerstein reports grants from Gilead Sciences Inc., grants from Pfizer Inc, grants from Synergenics LLC, outside the submitted work; .

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Samuel 2. Surname (Last Name) Klein 3. Date 04-November-2019

4. Are you the corresponding author? Yes No

5. Manuscript Title
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Are there any relevant conflicts of interest? Yes No

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Name of Entity	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments	
Merck Sharp & Dohme Corp	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
Janssen Pharmaceuticals	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
Pfizer	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
NovoNordisk	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
Merck Sharp & Dohme Corp	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>		X
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Dr. Klein reports grants from Merck Sharp & Dohme Corp, grants from Janssen Pharmaceuticals, personal fees from Pfizer, personal fees from Novo Nordisk, personal fees from Merck Sharp & Dohme Corp, outside the submitted work; .

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TREND Statement Checklist

Paper Section/Topic	Item No.	Descriptor	Reported?	
			✓	Pg #
TITLE and ABSTRACT				
Title and Abstract	1	• Information on how units were allocated to interventions	✓	2
		• Structured abstract recommended	✓	2
		• Information on target population or study sample	✓	2
INTRODUCTION				
Background	2	• Scientific background and explanation of rationale	✓	4-6
		• Theories used in designing behavioral interventions		N/A
METHODS				
Participants	3	• Eligibility criteria for participants, including criteria at different levels in recruitment/sampling plan (e.g., cities, clinics, subjects)	✓	12
		• Method of recruitment (e.g., referral, self-selection), including the sampling method if a systematic sampling plan was implemented	✓	12
		• Recruitment setting	✓	12
		• Settings and locations where the data were collected	✓	12
Interventions	4	• Details of the interventions intended for each study condition and how and when they were actually administered, specifically including:	✓	13-15
		○ Content: what was given?	✓	13-15
		○ Delivery method: how was the content given?	✓	13-15
		○ Unit of delivery: how were subjects grouped during delivery?	✓	13-15
		○ Deliverer: who delivered the intervention?	✓	14
		○ Setting: where was the intervention delivered?	✓	12
		○ Exposure quantity and duration: how many sessions or episodes or events were intended to be delivered? How long were they intended to last?	✓	14-15
		○ Time span: how long was it intended to take to deliver the intervention to each unit?	✓	14-15
○ Activities to increase compliance or adherence (e.g., incentives)	✓	14		
Objectives	5	• Specific objectives and hypotheses	✓	5-6
Outcomes	6	• Clearly defined primary and secondary outcome measures	✓	5-6
		• Methods used to collect data and any methods used to enhance the quality of measurements	✓	12-16
		• Information on validated instruments such as psychometric and biometric properties		N/A
Sample size	7	• How sample size was determined and, when applicable, explanation of any interim analyses and stopping rules	✓	17
Assignment method	8	• Unit of assignment (the unit being assigned to study condition, e.g., individual, group, community)	✓	12
		• Method used to assign units to study conditions, including details of any restriction (e.g., blocking, stratification, minimization)	✓	12
		• Inclusion of aspects employed to help minimize potential bias induced due to non-randomization (e.g., matching)		N/A
Blinding (masking)	9	• Whether or not participants, those administering the interventions, and those assessing the outcomes were blinded to study condition assignment; if so, statement regarding how the blinding was accomplished and how it was assessed		N/A
Unit of Analysis	10	• Description of the smallest unit that is being analysed to assess intervention effects (e.g., individual, group, or community)	✓	12,14-15
		• If the unit of analysis differs from the unit of assignment, the analytical method used to account for this (e.g., adjusting the standard error estimates by the design effect or using multilevel analysis)		N/A
Statistical methods	11	• Statistical methods used to compare study groups for primary methods outcome(s), including complex methods for correlated data	✓	16-17
		• Statistical methods used for additional analyses, such as subgroup analyses and adjusted analysis		N/A
		• Methods for imputing missing data, if used		N/A
		• Statistical software or programs used	✓	17

TREND Statement Checklist

RESULTS			
Participant flow	12	<ul style="list-style-type: none"> Flow of participants through each stage of the study: enrollment, assignment, allocation and intervention exposure, follow-up, analysis (a diagram is strongly recommended) 	✓ 12, 14-15, Suppl. Figure 2
		<ul style="list-style-type: none"> Enrollment: the numbers of participants screened for eligibility, found to be eligible or not eligible, declined to be enrolled, and enrolled in the study 	✓ 12, 14-15, Suppl. Figure 2
		<ul style="list-style-type: none"> Assignment: the numbers of participants assigned to a study condition 	✓ 12,14-15 Suppl. Figure 2
		<ul style="list-style-type: none"> Allocation and intervention exposure: the number of participants assigned to each study condition and the number of participants who received each intervention 	✓ 14-15, Suppl. Figure 2
		<ul style="list-style-type: none"> Follow-up: the number of participants who completed the follow-up or did not complete the follow-up (i.e., lost to follow-up), by study condition 	✓ 12,14-15 Suppl. Figure 2
		<ul style="list-style-type: none"> Analysis: the number of participants included in or excluded from the main analysis, by study condition 	✓ 12,14-15 Suppl. Figure 2
		<ul style="list-style-type: none"> Description of protocol deviations from study as planned, along with reasons 	N/A
Recruitment	13	<ul style="list-style-type: none"> Dates defining the periods of recruitment and follow-up 	12-15
Baseline data	14	<ul style="list-style-type: none"> Baseline demographic and clinical characteristics of participants in each study condition 	✓ 6, Tables 1 & 2
		<ul style="list-style-type: none"> Baseline characteristics for each study condition relevant to specific disease prevention research 	N/A
		<ul style="list-style-type: none"> Baseline comparisons of those lost to follow-up and those retained, overall and by study condition 	N/A
		<ul style="list-style-type: none"> Comparison between study population at baseline and target population of interest 	N/A
Baseline equivalence	15	<ul style="list-style-type: none"> Data on study group equivalence at baseline and statistical methods used to control for baseline differences 	✓ 6-7
Numbers analyzed	16	<ul style="list-style-type: none"> Number of participants (denominator) included in each analysis for each study condition, particularly when the denominators change for different outcomes; statement of the results in absolute numbers when feasible 	✓ 12, 14-15, Figures 1 & 2, Tables 1 & 2
		<ul style="list-style-type: none"> Indication of whether the analysis strategy was "intention to treat" or, if not, description of how non-compliers were treated in the analyses 	N/A
Outcomes and estimation	17	<ul style="list-style-type: none"> For each primary and secondary outcome, a summary of results for each estimation study condition, and the estimated effect size and a confidence interval to indicate the precision 	✓ 6-8, Tables 1 & 2, Figures 1 & 2
		<ul style="list-style-type: none"> Inclusion of null and negative findings 	N/A
		<ul style="list-style-type: none"> Inclusion of results from testing pre-specified causal pathways through which the intervention was intended to operate, if any 	✓ 7, Figure 1
Ancillary analyses	18	<ul style="list-style-type: none"> Summary of other analyses performed, including subgroup or restricted analyses, indicating which are pre-specified or exploratory 	N/A
Adverse events	19	<ul style="list-style-type: none"> Summary of all important adverse events or unintended effects in each study condition (including summary measures, effect size estimates, and confidence intervals) 	N/A

TREND Statement Checklist

DISCUSSION			
Interpretation	20	• Interpretation of the results, taking into account study hypotheses, sources of potential bias, imprecision of measures, multiplicative analyses, and other limitations or weaknesses of the study	✓ 8-11
		• Discussion of results taking into account the mechanism by which the intervention was intended to work (causal pathways) or alternative mechanisms or explanations	✓ 8-11
		• Discussion of the success of and barriers to implementing the intervention, fidelity of implementation	N/A
		• Discussion of research, programmatic, or policy implications	✓ 8-11
Generalizability	21	• Generalizability (external validity) of the trial findings, taking into account the study population, the characteristics of the intervention, length of follow-up, incentives, compliance rates, specific sites/settings involved in the study, and other contextual issues	✓ 8-11
Overall evidence	22	• General interpretation of the results in the context of current evidence and current theory	✓ 8-11