

#### 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 Infor	mation	
1. Given Name (Fi Kazuya		2. Surname (Last Name) Miyagawa	3. Date 08-July-2018
4. Are you the cor	responding author?	Yes 🖌 No	Corresponding Author's Name Spiekerkoetter, Edda
5. Manuscript Titl Fragile Histidine		Modifier Gene in Pulmona	ry Arterial Hypertension
6. Manuscript Ide Blue-201712-25	ntifying Number (if you 53OC.R2	know it)	
Section 2.	The Work Under	Consideration for Publ	ication
	submitted work (includi		n a third party (government, commercial, private foundation, etc.) for lata monitoring board, study design, manuscript preparation,

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 rov	v.
Excess rows can be removed by pressing the "X" button.	

Name of Institution/Company	Grant?	Personal Fees	Non-Financial Support?	Other?	Comments	
Japan Heart Foundation / Bayer Yakuhin Research Grant Abroad	$\checkmark$					
The Uehara Memorial Foundation	$\checkmark$					

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Are there any relevant conflicts of interest?

Yes 🖌 No

#### Section 4.

Intellectual Property -- Patents & Copyrights

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



# Section 5. Relationships not covered above

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Dr. Miyagawa reports grants from Japan Heart Foundation / Bayer Yakuhin Research Grant Abroad, grants from The Uehara Memorial Foundation, during the conduct of the study; .

#### **Evaluation and Feedback**



Section 1.
Identifying Information
1. Given Name (First Name) LISA UHEELER 3. Date JULY 19. 2018
4. Are you the corresponding author?
5. Manuscript Title Fragile Histidine Triad (FHIT), a Novel Modifier Gene in Pulmonary Arterial Hypertension
6. Manuscript Identifying Number (if you know it) Blue-201712-2553OC.R2
Section 2. The Work Under Consideration for Publication
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Are there any relevant conflicts of interest? Yes X No
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1. Given Name (First Kay	Name)	2. Surnar Huebnei	ne (Last Name) r	3. Date 11-July-2018
4. Are you the corre	sponding author?	Yes	✓ No	Corresponding Author's Name Edda Spiekerkoetter
	fying Number (if you		ne in Pulmona	ary Arterial Hypertension

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Are there any relevant conflicts of interest?	Yes	$\checkmark$	No
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🖌 No

Are there any rele	evant conflicts of	f interest?		Yes
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Section 1.					
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1. Given Name (F Deepti	irst Name)	2. Surname (L Sudheendra			3. Date 10-July-2018
4. Are you the co	rresponding author?	✓ Yes	No		
5. Manuscript Titl Fragile Histidine	e e Triad (FHIT), a Novel M	1odifier Gene ir	n Pulmonary Arterial I	Hypertension	
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any aspect of the statistical analysis	submitted work (includin , etc.)?	g but not limited	to grants, data monito		ommercial, private foundation, etc.) for esign, manuscript preparation,
Fragile Histidine 6. Manuscript Ide Blue-201712-25 Section 2. Did you or your in any aspect of the statistical analysis	Triad (FHIT), a Novel M ntifying Number (if you k 53OC.R2 The Work Under C stitution <b>at any time</b> rece submitted work (includin	now it) Consideration eive payment or s g but not limited	for Publication	rty (government, cc	-

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Are there any relevant conflicts of interest? Y	es	$\checkmark$	No
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work? 🗌 Yes 🛛	🖌 N	10
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5. Manuscript Title Fragile Histidine		Modifier Gene in Pulmona	iry Arterial Hypertension
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Dr. Saldivar has nothing to disclose.

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Section 1.			
Section 1.	Identifying Infor	mation	
1. Given Name (F Svenja	irst Name)	2. Surname (Last Name) Dannewitz Prosseda	3. Date 10-July-2018
4. Are you the co	rresponding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
5. Manuscript Titl Fragile Histidine		Modifier Gene in Pulmona	ry Arterial Hypertension
6. Manuscript Ide Blue-201712-25	entifying Number (if you 53OC.R2	know it)	
Section 2.	The Work Under	Consideration for Pub	lication
any aspect of the statistical analysis	submitted work (includi	ng but not limited to grants, o	m a third party (government, commercial, private foundation, etc.) for data monitoring board, study design, manuscript preparation,

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No

Are there any relevant conflicts of interest?		Yes	✓
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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

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?	Licensed?	Royalties?	Licensee?	Comments	
PCT/US2018/033533		$\checkmark$	$\checkmark$		Denovo Biopharma LLC	License number S17-202	



# Section 5. Relationships not covered above

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Dr. Dannewitz Prosseda reports In addition, Dr. Dannewitz Prosseda has a patent PCT/US2018/033533 licensed to Denovo Biopharma LLC.

#### **Evaluation and Feedback**



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Section 1.	Identifying Info	rmation	
1. Given Name (Fi Edda	rst Name)	2. Surname (Last Name) Spiekerkoetter	3. Date 18-December-2017
4. Are you the co	responding author?	✓ Yes No	
5. Manuscript Titl Fragile Histidine		Modifier Gene in Pulmonary Arterial Hypertension	
6. Manuscript Ide	ntifying Number (if you	ı know it)	

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Are there any relevant conflicts of interest?	🖌 Yes	No
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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 <mark>?</mark>	Other?	Comments	
National Heart Lung and Blood Institute	$\checkmark$				K08HL107450-01, R01 HL128734-01A1	
Pulmonary Hypertension Association (PHA)/ American Heart Association (AHA)	$\checkmark$				supplemental K08 award 2011	

#### Section 3.

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Are there any relevant conflicts of interest?

Yes 🖌 No

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

No



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S17-202/USSN 62/508,881	$\checkmark$				Stanford University		

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Dr. Spiekerkoetter reports grants from National Heart Lung and Blood Institute, grants from Pulmonary Hypertension Association (PHA)/ American Heart Association (AHA), during the conduct of the study; In addition, Dr. Spiekerkoetter has a patent S17-202/USSN 62/508,881 pending to Stanford University.

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1. Given Name (First Name) Mario	2. Surname (Last Name) Boehm	3. Date 09-July-2018
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
5. Manuscript Title Fragile Histidine Triad (FHIT), a Novel		ry Arterial Hypertension
6. Manuscript Identifying Number (if you Blue-201712-2553OC.R2	know it)	

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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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	٧o



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Section 1. 1. Given Name (F David	Identifying Infor	mation 2. Surname (Last Name Solow-Cordero	) 3. Date 09-July-2018
·	rresponding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
	e Triad (FHIT), a Novel i ntifying Number (if you		ary Arterial Hypertension
Section 2.	The Work Under	Consideration for Pub	lication

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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?		Yes	√	
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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

No

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Patent?	Pending?	Issued?	Licensed?	Royalties?	Licensee?	Comments	
9,474,745 Use of FK506 for the Treatment of Pulmonary Arterial Hypertension		$\checkmark$	$\checkmark$	$\checkmark$	Vivus, Inc.		



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Dr. Solow-Cordero reports In addition, Dr. Solow-Cordero has a patent 9,474,745 Use of FK506 for the Treatment of Pulmonary Arterial Hypertension with royalties paid to Vivus, Inc..

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1. Given Name (First Name) Fan	2. Surname (Last Name) Zhang	3. Date 09-July-2018
Are you the corresponding author	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
5. Manuscript Title Fragile Histidine Triad (FHIT), a Nc 6. Manuscript Identifying Number (if		ry Arterial Hypertension

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Are there any relevant conflicts of interest?		Yes
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work?	Yes	🖌 No	)



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Section 1. Identifying Info	rmation		
1. Given Name (First Name)2. Surname (LastMicheleDonato		3. Date 15-October-2018	
4. Are you the corresponding author?	Yes 🗸 No	Corresponding Author's Name Edda Spiekerkoetter	
5. Manuscript Title Fragile Histidine Triad (FHIT), a Nove	l Modifier Gene in Pulmona	ary Arterial Hypertension	
6. Manuscript Identifying Number (if yo Blue-201712-2553OC	u know it)		

### Section 2. The Work Under Consideration for Publication

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Are there any relevant conflicts of interest?	$\checkmark$	Yes		No
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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 <mark>?</mark>	Other?	Comments	
NIH	$\checkmark$					

### Section 3. Relevant financial activities outside the submitted work.

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Are there any relevant conflicts of interest? Yes 🗸 No

### 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

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. Donato reports grants from NIH, during the conduct of the study.

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1. Given Name (First Name) Lingli	2. Surname (Last Name) Wang	3. Date 14-October-2018
Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
. Manuscript Title ragile Histidine Triad (FHIT), a Novel	Modifier Gene in Pulmon	ary Arterial Hypertension
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Are there any relevant conflicts of interest?	Yes
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Are there any relevant conflicts of interest?	Yes	$\checkmark$	No
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work?		Yes	$\checkmark$	No
	1 1		•	



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Dr. Wang has nothing to disclose.

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1. Given Name (First Name) Eric	rmation 2. Surname (Last Name) Austin	3. Date 22-October-2018
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter, MD
<b>F</b> 14 <b>F</b> 14		
<ol> <li>Manuscript Title</li> <li>Fragile Histidine Triad (FHIT), a Novel</li> <li>Manuscript Identifying Number (if you</li> <li>Blue-201712-2553OC</li> </ol>		ry Arterial Hypertension

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Are there any relevant conflicts of interest?	)	Yes
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Are there any relevant conflicts of interest?	Yes	$\checkmark$	No
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work?	Yes	٧o



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Dr. Austin has nothing to disclose.

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Section 1.	Identifying Infor	mation	
1. Given Name (Fi Kazuya	rst Name)	2. Surname (Last Name) Kuramoto	3. Date 13-October-2018
4. Are you the cor	responding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
5. Manuscript Title Fragile Histidine		Modifier Gene in Pulmona	ry Arterial Hypertension
6. Manuscript Ider Blue-201712-255	ntifying Number (if you 53OC	know it)	

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Are there any relevant conflicts of interest?		Yes
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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 Fees?	Non-Financial Support?	Other?	Comments	
Nippon-Shinyaku Co. Ltd.,				$\checkmark$	Employer	

# Section 4. Intellectual Property -- Patents & Copyrights Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes Vo



# Section 5. Relationships not covered above

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Dr. Kuramoto reports other from Nippon-Shinyaku Co. Ltd.,, outside the submitted work; .

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1. Given Name (First Name) Xuefei	2. Surname (Last Name) Tian	3. Date 12-October-2018
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name Edda Spiekerkoetter
5. Manuscript Title Fragile Histidine Triad (FHIT), a Novel 6. Manuscript Identifying Number (if you		ry Arterial Hypertension

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Are there any relevant conflicts of interest?		Yes	
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Are there any relevant conflicts of interest?	Yes
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work?	Yes	; 🖌 No	5



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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".

### 3. 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.

### 4. 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.

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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



Section 1.	Identifying Inform	mation			
	identifying infor	nation			
1. Given Name (Fi Purvesh	rst Name)	2. Surnan Khatri	ne (Last Name)		3. Date 13-October-2018
4. Are you the cor	responding author?	Yes	✓ No	Corresponding Author's Nam	ne
				Edda Spiekerkoetter	
5. Manuscript Title Fragile Histidine		Nodifier Ger	ie in Pulmona	ry Arterial Hypertension	
6. Manuscript Ider Blue-201712-255	ntifying Number (if you k 53OC	now it)			
Section 2.	The Work Under C	Considerat	ion for Pub	ication	

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?	$\checkmark$	Yes		No
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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 <mark>?</mark>	Other?	Comments	
National Institutes of Health	$\checkmark$				R01 funding for the work reported in this manuscript.	

### 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

If yes, please fill out the appropriate information below.

Name of Entity	Grant?	Personal Fees?	Non-Financial Support?	Other?	Comments
Inflammatix, Inc.				$\checkmark$	Co-founder; Inflammatix commercializes tests for infectious diseases



Name of Entity	Grant?	Personal Fees <b>?</b>	Non-Financial Support?	Other?	Comments	
Vir Bio	$\checkmark$			$\checkmark$	Funding for developing methods for identifying drug targets and drug repurposing	

# Section 4. 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?	Licensed?	Royalties?	Licensee?	Comments	
ENZASTAURIN AND FRAGILE HISTIDINE TRIAD (FHIT)-INCREASING AGENTS FOR THE TREATMENT OF PULMONARY HYPERTENSION	$\checkmark$						

# 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. Disc

**Disclosure Statement** 

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

Dr. Khatri reports grants from National Institutes of Health, during the conduct of the study; other from Inflammatix, Inc., grants and other from Vir Bio, outside the submitted work; In addition, Dr. Khatri has a patent ENZASTAURIN AND FRAGILE HISTIDINE TRIAD (FHIT)-INCREASING AGENTS FOR THE TREATMENT OF PULMONARY HYPERTENSION pending.

**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.

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Section 1. Identifying Inf	ormation						
1. Given Name (First Name) James	2. Surname (Last Name) Loyd	3. Date 09-July-2018					
4. Are you the corresponding author?	Yes 🖌 No	Corresponding Author's Name					
5. Manuscript Title Fragile Histidine Triad (FHIT), a Novel Modifier Gene in Pulmonary Arterial Hypertension							
6. Manuscript Identifying Number (if y Blue-201712-2553OC.R2	ou know it)						
Section 2. The Work Under	er Consideration for Pub	lication					
	iding but not limited to grants,	m a third party (government, commercial, private foundation, etc.) for data monitoring board, study design, manuscript preparation,					

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Are there any relevant conflicts of interest?	Yes	$\checkmark$	No
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work?	<b></b>	Yes	✓ No	,
	1 1		•	



# Section 5. Relationships not covered above

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Dr. Loyd has nothing to disclose.

#### **Evaluation and Feedback**