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Evaluation for the Genetic Association between Store-Operated Calcium Influx Pathway (STIM1 and ORAI1) and Human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Biology 2020, 9(11), 388; <https://doi.org/10.3390/biology9110388>

by Lahu Muhammad Irham^{1,2}, Wan-Hsuan Chou^{1,3}, Yu-Shiuan Wang^{4,5}, Wirawan Adikusuma^{1,6}, Henry Sung-Ching Wong^{1,3}, Dyah Aryan Periwitasari², Wan-Chen Huang⁴, Ben-Kuen Chen^{7,8,*}, Hwai-I Yang^{9,10,*} and Wei-Chiao Chang^{1,3,11,12,*}

Reviewer 1: Kuanwei Sheng

Reviewer 2: Anonymous

Biology 2020, 9(11), 388; <https://doi.org/10.3390/biology9110388>

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(This article belongs to the Section Genetics and Genomics)

Round 1

Reviewer 1 Report

The manuscript improved significantly compared to the last version. The addition of cell culture experiments supplements the sequencing results nicely. I therefore agree to publish the manuscript.

Reviewer 2 Report

Authors have addressed all questions and concerns.

This manuscript is a resubmission of an earlier submission. The following is a list of the peer review reports and author responses from that submission.

Round 1


Reviewer 1 Report

Irham et al conducted large-scale profiling for STIM1 and ORAI1 SNP in large HBV patient sets. Though the statistical significance are low for the SNP probably due to not enough HCC patients, the data are still quite valuable to the community and have potential clinical benefits. I would happily accept the paper if the author can address my following questions.

1. I am quite interested in whether there are gender differences in SNP profile. Namely, since you have a large dataset, could you evaluate whether there is any SNP signature that is more dominant in males or females? And after the gender grouping, if there are any SNPs that showed significant association?
2. For all the SNPs you identified, could you please show some examples that some of the SNPs can cause functional alterations to the proteins (mutations in critical domains that may cause a structural change of the protein), especially in the SNPs that shows trends to be associated with the HCC?

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shows trends to be associated with the HCC?

Reviewer 2 Report

The manuscript 'Evaluation of genetic susceptibility between store-operated calcium influx pathway (STIM1 and ORAI1) and human hepatocellular carcinoma in patients with chronic hepatitis B infection' aims to identify polymorphisms associated with human hepatocellular carcinoma in 3631 Taiwanese patients. Overall, the manuscript presents data that are important to the field, yet a few points need to be clarified. The specific points that resulted in this conclusion are listed below.

The title is not reflective of the research presented in the manuscript. Genetic susceptibility is used in the title however the research is investigating association of SNP with human hepatocellular carcinoma not genetic susceptibility. Genetic susceptibility is also known as genetic predisposition, which cannot be properly evaluated in the current study. Title needs to be revised to reflect the presented study.

Line 28: Remove 'of from 'in total of 3631'.

Line 32-33. The authors state that 'our study revealed that calcium signaling is essential for hepatitis B virus replication'. This is not a valid conclusion of the study presented. While there is published research supporting this conclusion, the current study does not. The current study only evaluates SNP in STIM1 and ORAI1 and calcium signaling is not measured. Remove sentence from paragraph.

Line 130: Specify which Qiagen kit was used.

Line 279-281: The authors state that 'We acknowledge a lack of calcium concentration data as a limitation of this study, which prevented us from understanding its correlation with HCC progression in CHB patients, even though a previous study revealed that the lowest Ca²⁺ level could be correlated with a decrease in HBV replication'. This is beyond the scope of the current study as the presented research does not evaluate calcium signaling. This sentence and similar statements in the discussion need to be removed.

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Font size varies throughout the entire manuscript. Keep font size consistent.

Table 2: Font of column titles is not consistent (minor allelic frequencies). Keep font consistent.

Line 137: Change 'an' to 'a'.

Line 139: Define OR. OR is not defined until line 162.

Line 233: Remove 'of from 'in total of 3631'.

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Author Response to Reviewer Comment

Reviewer 1.

Comments and Suggestions for Authors

Irham et al conducted large-scale profiling for *STIM1* and *ORAI1* SNP in large HBV patient sets. Though the statistical significance are low for the SNP probably due to not enough HCC patients, the data are still quite valuable to the community and have potential clinical benefits. I would happily accept the paper if the author can address my following questions.

Answer: We sincerely thank the reviewer for the time taken to review our work.

Q1: Reviewer #1. I am quite interested in whether there are gender differences in SNP profile. Namely, since you have a large dataset, could you evaluate whether there is any SNP signature that is more dominant in males or females? And after the gender grouping, if there are any SNPs that showed significant association?

A1: We sincerely thank the reviewer for their suggestion. It is a very important point. As suggested by reviewer, we confirmed all the SNPs (40 SNPs) with stratified analyses in association between the SNPs and gender (male and female). In the present analyses showed that male gender is more dominant (2196 subjects) compared to female gender (1435 subjects) as showed in the following **Table**. In addition, we found that two SNPs (rs2959081 and rs7116520) of *STIM1* and one SNP of rs6486795 *ORAI1* gene showed an association for gender. The trend of these two *STIM1* SNPs (rs2959081 and rs7116520) showed an association in female gender, while the *ORAI1* SNP rs6486795 showed an association in male gender. Interestingly, through this analysis, we still identified the trend of two SNPs (rs7116520 and rs6486795) are consistent to be associated not only in HCC but also in case of gender. Furthermore, herein, we identified the new SNP of *ORAI1* rs2959081 showed the trend of significant association in male gender. However, after correction for multiple testing, none of the SNPs reached a significant level.

Associations of *STIM1* and *ORAI1* genes with gender (Male and Female)

SNP	Gene	Effect allele	Male Gender N=2196 (60.48%)		Female Gender N=1435 (39.52%)	
			(Genotype) <i>p</i> value ^a [q value]	(Dominant) <i>p</i> value ^a [q value]	(Genotype) <i>p</i> value ^a [q value]	(Dominant) <i>p</i> value ^a [q value]
rs4243966	<i>STIM1</i>	C	0.879 [0.963]	0.993 [0.993]	0.349 [0.971]	0.321 [0.975]
rs7943201	<i>STIM1</i>	A	0.776 [0.963]	0.485 [0.993]	0.417 [0.971]	0.323 [0.975]
rs11030122	<i>STIM1</i>	G	0.817 [0.963]	0.821 [0.993]	0.927 [0.971]	0.727 [0.975]
rs7951076	<i>STIM1</i>	A	0.633 [0.963]	0.362 [0.993]	0.919 [0.971]	0.719 [0.975]
rs7952083	<i>STIM1</i>	C	0.775 [0.963]	0.806 [0.993]	0.512 [0.971]	0.403 [0.975]
rs7120828	<i>STIM1</i>	T	0.372 [0.963]	0.203 [0.993]	0.277 [0.971]	0.130 [0.975]
rs10458894	<i>STIM1</i>	T	0.370 [0.963]	0.193 [0.993]	0.263 [0.971]	0.547 [0.975]
rs7120683	<i>STIM1</i>	G	0.278 [0.963]	0.110 [0.993]	0.366 [0.971]	0.160 [0.975]
rs10835262	<i>STIM1</i>	A	0.913 [0.963]	0.908 [0.993]	0.920 [0.971]	0.828 [0.975]
rs4622250	<i>STIM1</i>	T	0.512 [0.963]	0.926 [0.993]	0.970 [0.971]	0.873 [0.975]
rs75197750	<i>STIM1</i>	T	0.196 [0.963]	0.131 [0.993]	0.668 [0.971]	0.380 [0.975]
rs10500589	<i>STIM1</i>	C	0.603 [0.963]	0.319 [0.993]	0.966 [0.971]	0.883 [0.975]
rs11030209	<i>STIM1</i>	C	0.642 [0.963]	0.926 [0.993]	0.905 [0.971]	0.761 [0.975]
rs10835270	<i>STIM1</i>	T	0.394 [0.963]	0.180 [0.993]	0.794 [0.971]	0.837 [0.975]
rs11030210	<i>STIM1</i>	T	0.729 [0.963]	0.945 [0.993]	0.381 [0.971]	0.479 [0.975]
rs7929653	<i>STIM1</i>	A	0.693 [0.963]	0.963 [0.993]	0.923 [0.971]	0.928 [0.975]
rs6578418	<i>STIM1</i>	G	0.240 [0.963]	0.095 [0.993]	0.328 [0.971]	0.249 [0.975]
rs10835272	<i>STIM1</i>	A	0.375 [0.963]	0.838 [0.993]	0.598 [0.971]	0.826 [0.975]

rs10742189	<i>STIM1</i>	C	0.808 [0.963]	0.755 [0.993]	0.404 [0.971]	0.232 [0.975]
rs7129444	<i>STIM1</i>	C	0.824 [0.963]	0.563 [0.993]	0.736 [0.971]	0.843 [0.975]
rs7118422	<i>STIM1</i>	T	0.926 [0.963]	0.929 [0.993]	0.715 [0.971]	0.419 [0.975]
rs4910863	<i>STIM1</i>	C	0.887 [0.963]	0.835 [0.993]	0.208 [0.971]	0.076 [0.975]
rs11030264	<i>STIM1</i>	G	0.757 [0.963]	0.476 [0.993]	0.183 [0.971]	0.068 [0.975]
rs7924984	<i>STIM1</i>	A	0.389 [0.963]	0.506 [0.993]	0.240 [0.971]	0.319 [0.975]
rs2412338	<i>STIM1</i>	T	0.396 [0.963]	0.501 [0.993]	0.163 [0.971]	0.358 [0.975]
rs10835402	<i>STIM1</i>	C	0.963 [0.963]	0.790 [0.993]	0.109 [0.971]	0.170 [0.975]
rs11030472	<i>STIM1</i>	G	0.147 [0.963]	0.078 [0.993]	0.456 [0.971]	0.214 [0.975]
rs11030478	<i>STIM1</i>	A	0.688 [0.963]	0.934 [0.993]	0.920 [0.971]	0.824 [0.975]
rs11030486	<i>STIM1</i>	T	0.698 [0.963]	0.874 [0.993]	0.926 [0.971]	0.823 [0.975]
rs12284835	<i>STIM1</i>	A	0.071 [0.963]	0.109 [0.993]	0.822 [0.971]	0.636 [0.975]
rs727152	<i>STIM1</i>	G	0.519 [0.963]	0.315 [0.993]	0.725 [0.971]	0.951 [0.975]
rs2959081	<i>STIM1</i>	C	0.608 [0.963]	0.543 [0.993]	0.021 [0.971]	0.005 [0.975]
rs11030639	<i>STIM1</i>	G	0.770 [0.963]	0.666 [0.993]	0.054 [0.971]	0.195 [0.975]
rs7116520	<i>STIM1</i>	G	0.410 [0.963]	0.209 [0.993]	0.007 [0.971]	0.001 [0.975]
rs1442725	<i>STIM1</i>	T	0.304 [0.963]	0.373 [0.993]	0.706 [0.971]	0.467 [0.975]
rs11030841	<i>STIM1</i>	A	0.611 [0.963]	0.486 [0.993]	0.381 [0.971]	0.885 [0.975]
rs4910882	<i>STIM1</i>	A	0.637 [0.963]	0.501 [0.993]	0.362 [0.971]	0.975 [0.975]
rs6486795	<i>ORAI1</i>	C	0.078 [0.963]	0.037 [0.993]	0.971 [0.971]	0.850 [0.975]
rs74936888	<i>ORAI1</i>	C	0.304 [0.963]	0.124 [0.993]	0.419 [0.971]	0.418 [0.975]
rs3741595	<i>ORAI1</i>	C	0.184 [0.963]	0.081 [0.993]	0.731 [0.971]	0.637 [0.975]

^a Adjusted by alanine transaminase and age. Significant *p* values are in **bold***. The *q* value is the false discovery rate (FDR) estimation for multi-testing.

Q2: Reviewer #1. For all the SNPs you identified, could you please show some examples that some of the SNPs can cause functional alterations to the proteins (mutations in critical domains that may cause a structural change of the protein), especially in the SNPs that shows trends to be associated with the HCC?

A2: We sincerely thank the reviewer for the time taken to review our work and the important suggestions are given. According to reviewer suggestion, we try to confirm the effect of the alteration/ deleteriousness of variants in protein by using some of the functional annotation tools such as the scale-invariant feature transform (SIFT), **Polymorphism Phenotyping v2 (PlyPhen-2)**, ClinVar and Combined Annotation Dependent Depletion (CADD) databases. As shown in the following **Table**, all four SNPs (*STIMI* rs6578418, rs11030472, rs7116520, and *ORAI1* rs6486795) are intron variants and we could not find the information of these four SNPs (the SNPs that shows trends to be associated with the HCC) effect of the change on protein function in three databases (SIFT, PlyPhen-2, and ClinVar). However, we still identified the information of four SNPs from CADD. The scores of CADD are displayed in four variants with score less than 30 which mean those four variants more likely to be benign rather than deleterious.

Information about the variants consequence from various databases							
SNP	Position (hg38) (bp)	Effect Allele	Consequence Type	PolyPhen2	SIFT	ClinVar	CADD
rs6578418	Chr11:3905002	G	Intron variant	NA	NA	NA	4.363(Benign)
rs11030472	Chr11:3978105	G	Intron variant	NA	NA	NA	8.781(Benign)
rs7116520	Chr11:4032147	G	Intron variant	NA	NA	NA	0.541(Benign)
rs6486795	Chr12:121638011	C	Intron variant	NA	NA	NA	0.481(Benign)

NA: Not Available

Additionally, we try to elaborate the relationship between the *STIMI* rs6578418, rs11030472, rs7116520 and *ORAI1* rs6486795 and gene expression. Through this way, we utilized the publicly available databases GTEx portal ([http:// www.gtexportal.org/home/](http://www.gtexportal.org/home/)) to confirmed the tissue expression quantitative trait loci. For two *STIMI* SNPs rs6578418 and rs11030472, we could not find the expression data on the GTEx portal database due to limited available tissue (**Table S3**). While the trend of *STIMI* rs7116520 showed in a variety of tissues including heart-left ventricle and nerve tibial lung tissue with GG genotype had higher expression compared to those with AG and AA genotype (**Figure S3**). Furthermore, *ORAI1* rs6486795 with CC genotype showing a highly expressed compared to those with TC and TT genotype in variety of tissues (*e.g.* whole blood, esophagus, thyroid, heart, stomach, pancreas, muscle, colon). This paragraph has been added in the result part of this study [Page 13, lines 298-308] and we also added the SNP annotation data query in the method section [Page 5, lines 138-141]. Furthermore, the Table (**Table S3**) and Figure (**Figure S3**) were added in the supplementary materials [Page 15, lines, 409-410 and page 16, lines 417-418].

Table S3. Expression Quantitative trait loci (eQTL) results of The SNP from Genotype-tissue expression (GTEx).

SNP ID	Gencode ID (ENSG00000-)	Gene symbol	p value	Effect size	Tissue	Actions
rs7116520	167323.11	<i>STIMI</i>	1.1e-9	-0.18	Nerve - Tibial	GG>AG>AA
	167323.11	<i>STIMI</i>	2.0e-8	-0.22	Heart-Left Ventricle	GG>AG>AA
rs6578418	NA	NA	NA	NA	NA	NA
rs11030472	NA	NA	NA	NA	NA	NA
rs6486795	276045.2	<i>ORAI1</i>	1.30E-17	0.22	Whole Blood	CC>CT>TT
	276045.2	<i>ORAI1</i>	9.90E-16	0.29	Esophagus - Mucosa	CC>CT>TT
	276045.2	<i>ORAI1</i>	3.30E-13	0.23	Thyroid	CC>CT>TT
	276045.2	<i>ORAI1</i>	7.60E-08	0.24	Stomach	CC>CT>TT
	276045.2	<i>ORAI1</i>	4.50E-07	0.25	Pancreas	CC>CT>TT
	276045.2	<i>ORAI1</i>	4.80E-07	0.11	Muscle - Skeletal	CC>CT>TT
	276045.2	<i>ORAI1</i>	1.5E-06	0.21	Colon - Transverse	CC>CT>TT
	276045.2	<i>ORAI1</i>	0.000029	0.19	Esophagus - Gastroesophageal Junction	CC>CT>TT
	276045.2	<i>ORAI1</i>	0.000058	0.17	Heart - Atrial Appendage	CC>CT>TT
	276045.2	<i>ORAI1</i>	0.00016	0.13	Esophagus - Muscularis	CC>CT>TT

Source: Expression Quantitative trait loci (eQTL) obtained from <https://gtexportal.org/home>, NA: Not Available

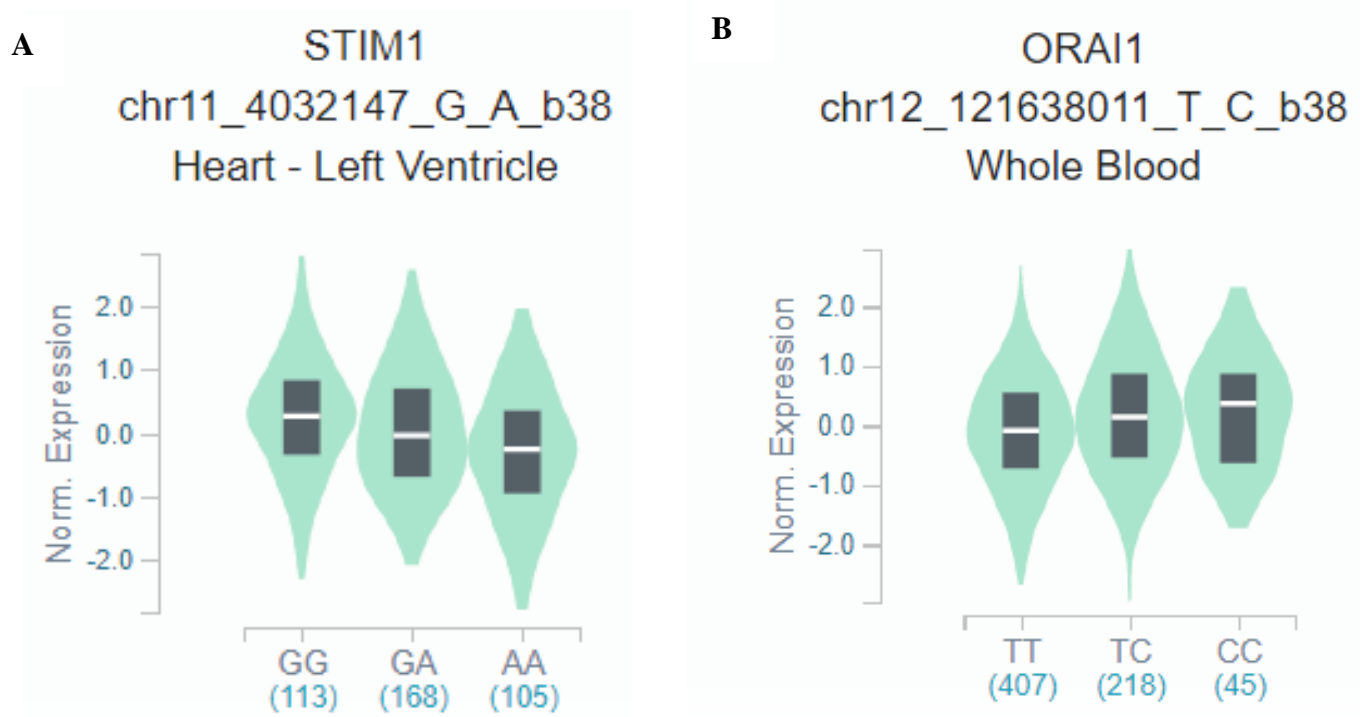


Figure S3. Correlation between genotype and expression in tissue determined by *Cis*-expression quantitative trait loci (*cis*-eQTLs). Figure **A** *STIM1* rs7116520 Homozygous (GG) >Heterozygous (AG)>Homozygous (AA) in heart-left ventricle tissue. Figure **B** *ORAI1* rs6486795 Homozygous CC>Heterozygous (TC)>Homozygous (TT) in whole blood.

Reviewer 2.

Comments and Suggestions for Authors

The manuscript 'Evaluation of genetic susceptibility between store-operated calcium influx pathway (STIM1 and ORAI1) and human hepatocellular carcinoma in patients with chronic hepatitis B infection' aims to identify polymorphisms associated with human hepatocellular carcinoma in 3631 Taiwanese patients. Overall, the manuscript presents data that are important to the field, yet a few points need to be clarified. The specific points that resulted in this conclusion are listed below.

Answer: We sincerely thank the reviewer for the time taken to review our work

Q1: Reviewer #2. The title is not reflective of the research presented in the manuscript. Genetic susceptibility is used in the title however the research is investigating association of SNP with human hepatocellular carcinoma not genetic susceptibility. Genetic susceptibility is also known as genetic predisposition, which cannot be properly evaluated in the current study. Title needs to be revised to reflect the presented study.

A1: We thank the reviewer for the suggestions. To reflect the title of this study, we currently evaluated the Store-Operated Calcium (STIM1 and ORAI1) as the additional supported data of functional cell-based assay. As shown in the current study (**Figure 2**) that we validated the roles of Store-Operated Calcium influx pathway (STIM1 and ORAI1) in HCC cell progression using two liver cancer cell lines (Huh 7 and HepG2). **Figure 2A** showed that the colocalization of STIM1 and Orail was inhibited by 2-APB pre-treatment 30 min in Huh 7 cells. We further analyzed the inhibition effect of Store-Operated Calcium activity suppression on cell migration in Huh 7 and HepG2 cell lines. The results showed that liver cancer cells migration (Huh 7 and HepG2) significantly decreased by 2-APB pre-treatment compared to the control group (**Figure 2B and 2C**). These results indicated that inhibition of Store-Operated Calcium by 2-APB was able to block the liver cancer cells migration. This paragraph has been added in the result part of this study [page 13, lines 310-328] and we also added the functional cell-based assay accordingly in the method section [Pages 5-6, lines 143-173]. In addition, we added the sentences in the discussion part [Pages 15, lines 388-390] and modified the conclusion [Pages 15-16, lines 397-402].

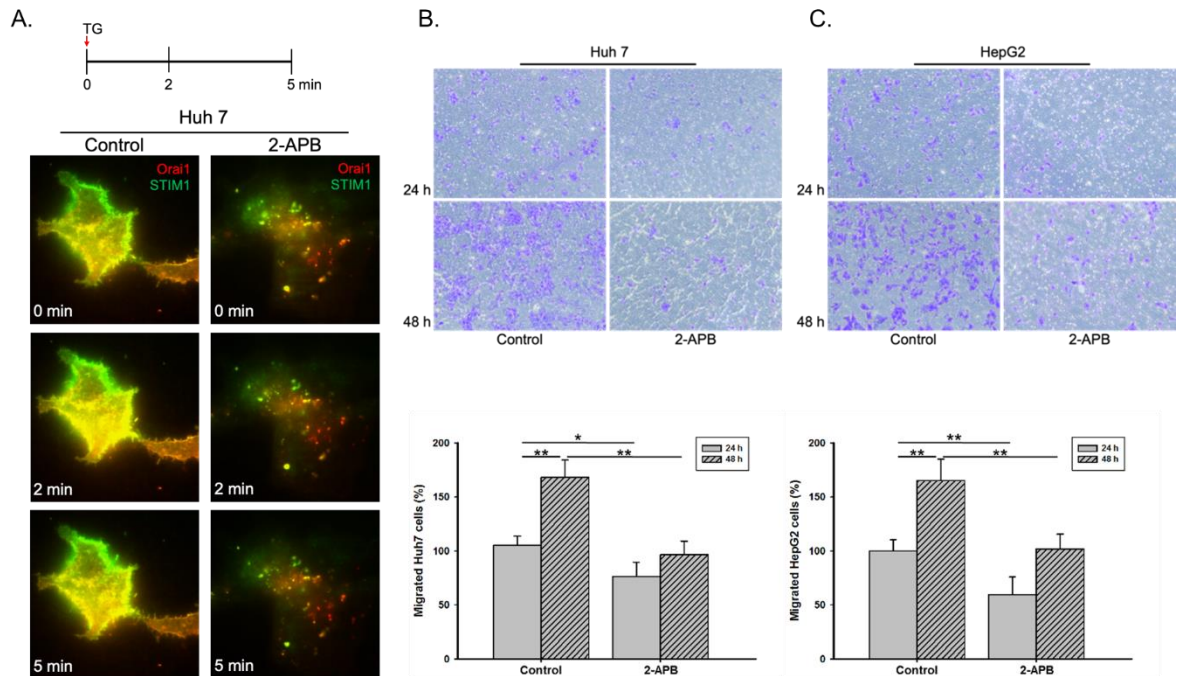


Figure 2. Inhibition of STIM1-Orai1 colocalization by 2-APB pre-treatment leads to reduced liver cancer cell migration ability. The Huh 7 cells were pre-treated with SOC inhibitor, 2-APB for 30 min. (A). The cells were co-transfected with STIM1-YFP and Orai1-mCherry and reseeded on Lab-Tek chambered cover glass for 24h. The inhibition effect of 2-APB on TG-induced SOC activity were observed with time-lapse TIRFM images. The inhibition effect of 2-APB on (B) Huh 7 or (C) HepG2 cell migration ability was examined by transwell migration assay. Statistically significant data are indicated by * for $p < 0.05$ and ** for $p < 0.01$.

Q2: Reviewer #2. Line 28: Remove ‘of’ from ‘in total of 3631’.

A2: We sincerely thank the reviewer for their suggestion. It is a very important point. We already Removed ‘of’ from ‘in total of 3631’ according to the reviewer’s suggestions [Page 1, line 31].

Q3: Reviewer #2. Line 32-33. The authors state that ‘our study revealed that calcium signaling is essential for hepatitis B virus replication’. This is not a valid conclusion of the study presented. While there is published research supporting this conclusion, the current study does not. The current study only evaluates SNP in *STIM1* and *ORAI1* and calcium signaling is not measured. Remove sentence from paragraph.

A3: We sincerely thank the reviewer for taking the time to review our work. According to the reviewer comment, we provided the data of functional cell-based study as showed in the **Figure 2**. Functional studies by both total internal reflection fluorescence microscopy and transwell migration assay confirmed the critical roles of SOC-mediated signaling in the HCC migration. According to the result of current data of functional cell-based assay, we corrected the sentences as presented in the following paragraph [Page 1, lines 35-41].

In particular, our study revealed that calcium (Ca^{2+}) signaling is essential for the migration of HCC, however, genetic polymorphisms of SOC pathway (*STIM1* and *ORAI1*) are not significantly associated with HCC progression. Besides, three SNPs of *STIM1* (rs6578418, rs11030472, and rs7116520) and one SNP of *ORAI1* (rs6486795) with a *borderline significant* trend, might be worth further study. Based on such a comprehensively screening in 3631 patients with chronic hepatitis, we believe that our results are of substantial interest to geneticists especially population geneticists, and clinical physicians.

Furthermore, we modified the conclusion of this study as presented in the following paragraph [Pages 15-16, lines 397-402].

Our study indicated that inhibition of SOC is able to block the cell migration in two liver cancer (Huh 7 and HepG2) cell lines. Although Ca^{2+} signaling is essential for the development of HCC, the genetic polymorphisms of SOC pathway (*STIM1* and *ORAI1*) are not significantly associated with HCC progression after multiple correction. Besides, three SNPs of *STIM1* (rs6578418, rs11030472, and rs7116520) and one SNP of *ORAI1* (rs6486795) with a *borderline significant* trend should be particularly focused in the future study.

Q4: Reviewer #2. Line 130: Specify which Qiagen kit was used.

A4: We thank the reviewer for the suggestions, the sentence has been added in the manuscript [Page 5, line 134].

Q5: Reviewer #2. Line 279-281: The authors state that ‘We acknowledge a lack of calcium concentration data as a limitation of this study, which prevented us from understanding its correlation with HCC progression in CHB patients, even though a previous study revealed that the lowest Ca^{2+} level could be correlated with a decrease in HBV replication’. This is beyond the scope of the current study as the presented research does not evaluate calcium signaling. This sentence and similar statements in the discussion need to be removed.

A5: We are very grateful to the reviewer’s suggestion. We already adjusted the sentences located in the last of discussion part as suggested by the reviewer. The modifications of sentences are presented in the following paragraph [Page 15, lines 390-394].

We acknowledge a weak correlation between *STIM1* and *ORAI1* polymorphisms and the risk of HCC progression in CHB patients may have been due to the modest sample size (3631 CHB patients), which led to a small power in the statistical analysis. Another possibility is that the majority of polymorphisms were located in the non-coding region. The genetic effects of each polymorphism to the expression of store-operated calcium channel are mild.

Q6: Reviewer #2. Font size varies throughout the entire manuscript. Keep font size consistent.

A6: We are very grateful to the reviewer's suggestion. We did it as suggested by the reviewer.

Q7: Reviewer #2. Table 2: Font of column titles is not consistent (minor allelic frequencies). Keep font consistent.

A7: Thank you for your suggestion. We did it as suggested by the reviewer.

Q8: Reviewer #2. Line 137: Change 'an' to 'a'.

A8: Many thanks to the reviewer's suggestions. We have made the correction according to the reviewer suggestion by change "an" to "a" in the sentences [Page 6, line 180]. The revised sentences are as below:

Linkage disequilibrium (LD) was evaluated with Haploview software version. 4.2 (Broad Institute, Cambridge, MA, USA) **by a SNP** analysis of *STIM1* and *ORAI1* polymorphisms together as haplotype blocks.

Q9: Reviewer #2. Line 139: Define OR. OR is not defined until line 162.

A9: Many thanks to the reviewer's suggestions. We have made the correction according to the comment from reviewer by adding the full name of OR, "odds ratio (OR)" [Page 6, line 182].

Q10: Reviewer #2. Line 233: Remove 'of' from 'in total of 3631'.

A10: We thank the reviewer for the suggestions. We have made the correction according to the reviewer's suggestion by removed 'of' from 'in total of 3631' [Page 14, line 343]. The revised sentences are as below:

"In total 3631 patients with chronic hepatitis were recruited."

Process email recording:

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Universitas Ahmad Dahlan Yogyakarta Mail - [Biology] Manuscript ID: biology-947101 - Article Processing Charge Confirmation and Ple...

UNIVERSITAS
AHMAD DAHLAN

LALU MUHAMMAD IRHAM <lalu.irham@pharm.uad.ac.id>

[Biology] Manuscript ID: biology-947101 - Article Processing Charge Confirmation and Please Provide the Invoice Information

2 messages

Amy Tian <amy.tian@mdpi.com>

Wed, Sep 16, 2020 at 7:13 PM

Reply-To: amy.tian@mdpi.com

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Ben Kuen Chen <bkchen58@mail.ncku.edu.tw>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Wei Chiao Chang <weichiao.chang@gmail.com>, Biology Editorial Office <biology@mdpi.com>

Dear Mr. Muhammad Irham,

Thank you very much for submitting your manuscript to Biology:

Journal name: Biology

Manuscript ID: biology-947101

Type of manuscript: Article

Title: Evaluation of Genetic susceptibility between store-operated calcium influx pathway (STIM1 and ORAI1) and human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Authors: Lalu Muhammad Irham, Wan Hsuan Chou, Wirawan Adikusuma, Henry Sung-Ching Wong, Dyah Aryani Perwitasari, Ben Kuen Chen *, Hwai I Yang *, Wei Chiao Chang *

Received: 12 September 2020

E-mails: lalu_irham@pharm.uad.ac.id, ocean.chou@tmu.edu.tw, adikusuma28@gmail.com, mingyue@gmail.com, diahperwitasari2003@yahoo.com, bkchen58@mail.ncku.edu.tw, hiyang@gate.sinica.edu.tw, weichiao.chang@gmail.com

Submitted to section: Cancer Biology,

https://www.mdpi.com/journal/biology/sections/cancer_biology

We confirm that, if accepted for publication, the following Article Processing Charges (APC), 1500 CHF, will apply to your article:

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Total APC: 1500 CHF

Please note that you may be entitled to a discount if you have previously received a discount code. Also note that reviewer vouchers must be applied before acceptance for publication. Vouchers can no longer be applied once an APC invoice has been issued. Reviewer vouchers, IOAP discounts, and vouchers offered by the Editorial Office cannot be applied to one invoice at the same time. You need to select one type of voucher to use. If you need to add any discount or replace the current discount with another type of discount, please contact the Biology Editorial Office as soon as possible.

Please confirm that you support open access publishing, which allows unlimited access to your published paper and that you will pay the Article Processing Charge if your manuscript is accepted.

Meanwhile, please also provide the invoice information so that we can make a record.

Affiliation Department:

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Name:

Invoice Email:

Order Email:

Address:

[Biology] Manuscript ID: biology-947101 - Assistant Editor Assigned

3 messages

Amy Tian <amy.tian@mdpi.com>

Wed, Sep 16, 2020 at 3:24 AM

Reply-To: amy.tian@mdpi.com

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Amy Tian <amy.tian@mdpi.com>, Lalu Muhammad Irham <lalu_irham@pharm.uad.ac.id>, Wan Hsuan Chou <ocean.chou@tmu.edu.tw>, Wirawan Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <miningyue@gmail.com>, Dyah Aryani Perwitasari <diahperwitasari2003@yahoo.com>, Ben Kuen Chen <bkchen58@mail.ncku.edu.tw>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Wei Chiao Chang <weichiao.chang@gmail.com>, Biology Editorial Office <biology@mdpi.com>

Dear Mr. Muhammad Irham,

Your manuscript has been assigned to Amy Tian for further processing who will act as a point of contact for any questions related to your paper.

Journal: Biology

Manuscript ID: biology-947101

Title: Evaluation of Genetic susceptibility between store-operated calcium influx pathway (STIM1 and ORAI1) and human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Authors: Lalu Muhammad Irham , Wan Hsuan Chou , Wirawan Adikusuma , Henry Sung-Ching Wong , Dyah Aryani Perwitasari , Ben Kuen Chen *, Hwai I Yang *, Wei Chiao Chang *

Received: 12 September 2020

E-mails: lalu_irham@pharm.uad.ac.id, ocean.chou@tmu.edu.tw, adikusuma28@gmail.com, miningyue@gmail.com, diahperwitasari2003@yahoo.com, bkchen58@mail.ncku.edu.tw, hiyang@gate.sinica.edu.tw, weichiao.chang@gmail.com

You can find it here:

https://susy.mdpi.com/user/manuscripts/review_info/7b3d5ec09c479a749a6509a712e1523bBest regards,
Ms. Amy Tian
Assistant Editor

--

MDPI Beijing Office Tongzhou, Jincheng Center, Room 2207, Tongzhou District, China

MDPI Biology Editorial Office

St. Alban-Anlage 66, 4052 Basel, Switzerland

E-Mail: biology@mdpi.com<http://www.mdpi.com/journal/biology>**LALU MUHAMMAD IRHAM** <lalu.irham@pharm.uad.ac.id>

Thu, Sep 17, 2020 at 6:04 PM

To: amy.tian@mdpi.com

Cc: Chang wei-chiao <weichiao.chang@gmail.com>, bkchen58@mail.ncku.edu.tw, hiyang@gate.sinica.edu.tw

Dear, Ms. Amy Tian

Thank you for giving us an opportunity to submit our article in "Biology". We attached the invoice information that you might need.

Affiliation Department : Department of Clinical Pharmacy, School of Pharmacy, College of Pharmacy, Taipei

[Biology] Manuscript ID: biology-947101 - Major Revisions (by 20 October)

2 messages

Amy Tian <amy.tian@mdpi.com>

Mon, Oct 12, 2020 at 10:30 PM

Reply-To: amy.tian@mdpi.com

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Lalu Muhammad Irham <lalu_irham@pharm.uad.ac.id>, Wan Hsuan Chou <ocean.chou@tmu.edu.tw>, Wirawan Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <minyingyue@gmail.com>, Dyah Aryani Perwitasari <diahperwitasari2003@yahoo.com>, Ben Kuen Chen <bkchen58@mail.ncku.edu.tw>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Wei Chiao Chang <weichiao.chang@gmail.com>, Biology Editorial Office <biology@mdpi.com>

Dear Mr. Muhammad Irham,

Thank you for submitting the following manuscript to Biology:

Manuscript ID: biology-947101

Type of manuscript: Article

Title: Evaluation of Genetic susceptibility between store-operated calcium influx pathway (STIM1 and ORAI1) and human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Authors: Lalu Muhammad Irham, Wan Hsuan Chou, Wirawan Adikusuma, Henry Sung-Ching Wong, Dyah Aryani Perwitasari, Ben Kuen Chen *, Hwai I Yang *, Wei Chiao Chang *

Received: 12 September 2020

E-mails: lalu_irham@pharm.uad.ac.id, ocean.chou@tmu.edu.tw, adikusuma28@gmail.com, minyingyue@gmail.com, diahperwitasari2003@yahoo.com, bkchen58@mail.ncku.edu.tw, hiyang@gate.sinica.edu.tw, weichiao.chang@gmail.com

Submitted to section: Cancer Biology,

https://www.mdpi.com/journal/biology/sections/cancer_biology

It has been reviewed by experts in the field and we request that you make major revisions before it is processed further. Please find your manuscript and the review reports at the following link:

<https://susy.mdpi.com/user/manuscripts/resubmit/7b3d5ec09c479a749a6509a712e1523b>

Your co-authors can also view this link if they have an account in our submission system using the e-mail address in this message.

Please revise the manuscript according to the reviewers' comments and upload the revised file within 7 days (by 20 October). Use the version of your manuscript found at the above link for your revisions, as the editorial office may have made formatting changes to your original submission. Any revisions should be clearly highlighted, for example using the "Track Changes" function in Microsoft Word, so that changes are easily visible to the editors and reviewers. Please provide a cover letter to explain point-by-point the details of the revisions in the manuscript and your responses to the reviewers' comments. Please include in your rebuttal if you found it impossible to address certain comments. The revised version will be inspected by the editors and reviewers. Please detail the revisions that have been made, citing the line number and exact change, so that the editor can check the changes expeditiously. Simple statements like 'done' or 'revised as requested' will not be accepted unless the change is simply a typographical error.

Please carefully read the guidelines outlined in the 'Instructions for Authors' on the journal website

<https://www.mdpi.com/journal/biology/instructions> and ensure that your manuscript resubmission adheres to these guidelines. In particular, please

Follow up: [Biology] Manuscript ID: biology-947101 - Revision Reminder

2 messages

Amy Tian <amy.tian@mdpi.com>

Fri, Oct 23, 2020 at 1:09 AM

Reply-To: amy.tian@mdpi.com

To: Wei-Chiao Chang <weichiao.chang@gmail.com>

Cc: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>, Wan Hsuan Chou <ocean.chou@tmu.edu.tw>, Wirawan Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <mininyue@gmail.com>, Dyah Aryani Perwitasari <diahperwitasari2003@yahoo.com>, Ben Kuen Chen <bkchen58@mail.ncku.edu.tw>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Biology Editorial Office <biology@mdpi.com>

Dear Dr. Chang,

I hope this email finds you well. This is the follow up with my message on 21 October 2020.

We are writing as we would like to know your revision progress and the possible time that it could be submitted? As the manuscript will be pending major revision for more time, we would like to ask you to upload the revised version to the online system as a new submission and it will be assigned a new manuscript ID.

<http://www.mdpi.com/user/manuscripts/upload/?journal=biology>

We will retain all records of biology-947101. The new submission will be sent to the previous reviewers directly once we receive it.

Thank you very much for your consideration.

Looking forward to hearing from you soon.

Kind regards,
Ms. Amy Tian
Assistant Editor

Biology Impact Factor in 2020: 3.796 (2019 JCR®). For joining the Editorial Board, please contact with biology@mdpi.com

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On 2020/10/21 9:42, Amy Tian wrote:

Dear Dr. Chang,

Thank you very much for your kind reply.

We can give enough time for you to do your studies.

Meanwhile, I have a proposal. As the manuscript will be pending major revision for more time, we would like to ask you to upload the revised version to the online system as a new submission and it will be assigned a new manuscript ID.

<http://www.mdpi.com/user/manuscripts/upload/?journal=biology>

We will retain all records of biology-947101. The new submission will be sent to the previous reviewers directly once we receive it.

Thank you very much for your consideration.

Looking forward to hearing from you soon.

Kind regards,
Ms. Amy Tian
Assistant Editor

Biology Impact Factor in 2020: 3.796 (2019 JCR®). For joining the Editorial Board, please contact with biology@mdpi.com

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On 2020/10/20 19:37, Wei-Chiao Chang wrote:

Dear Amy,

We are doing cell-based functional studies to improve the quality of this manuscript. Could you please extend the deadline of revision ? Hopefully, we will send it back next week.

____ Thank you~

wei chiao

張偉嶠特聘教授/副院長

臺北學大學藥學院

Wei-Chiao Chang (D.Phil.; Oxon)

Distinguished Professor/Vice Dean, School of Pharmacy

Taipei Medical University, Taiwan

Tel: 886-2-27361661 ext.6187 <tel:02%202736%201661>

<http://tw SNP.tmu.edu.tw/>

Amy Tian <amy.tian@mdpi.com> <<mailto:amy.tian@mdpi.com>>> 於 2020年10月 19 日 週一 下午4:29寫道:

Dear Mr. Muhammad Irham,

We sent a revision request for the following manuscript on 13 October 2020.

Manuscript ID: biology-947101
Type of manuscript: Article
Title: Evaluation of Genetic susceptibility between store-operated

calcium
influx pathway (STIM1 and ORAI1) and human Hepatocellular Carcinoma in
Patients with Chronic Hepatitis B Infection
Authors: Lalu Muhammad Irham, Wan Hsuan Chou, Wirawan Adikusuma, Henry
Sung-Ching Wong, Dyah Aryani Perwitasari, Ben Kuen Chen *, Hwai I
Yang *, Wei
Chiao Chang *

Received: 12 September 2020

E-mails: lalu_irham@pharm.uad.ac.id
<mailto:lalu_irham@pharm.uad.ac.id>, ocean.chou@tmu.edu.tw
<<mailto:ocean.chou@tmu.edu.tw>>,
adikusuma28@gmail.com <<mailto:adikusuma28@gmail.com>>,
mingyue@gmail.com <<mailto:mingyue@gmail.com>>,
diahperwitasari2003@yahoo.com <<mailto:diahperwitasari2003@yahoo.com>>,
bkchen58@mail.ncku.edu.tw <<mailto:bkchen58@mail.ncku.edu.tw>>,
hiyang@gate.sinica.edu.tw <<mailto:hiyang@gate.sinica.edu.tw>>,
weichiao.chang@gmail.com <<mailto:weichiao.chang@gmail.com>>

Submitted to section: Cancer Biology,
https://www.mdpi.com/journal/biology/sections/cancer_biology

May we kindly ask you to update us on the progress of your
revisions? If you
have finished your revisions, please upload the revised version
together with
your responses to the reviewers as soon as possible.

You can find your manuscript and review reports at this link:

<https://susy.mdpi.com/user/manuscripts/resubmit/7b3d5ec09c479a749a6509a712e1523b>

Thank you in advance for your kind cooperation and we look forward
to hearing
from you soon.

Kind regards,
Ms. Amy Tian
Assistant Editor

--

MDPI Beijing Office Tongzhou, Jincheng Center, Room 2207, Tongzhou
District,
China

MDPI Biology Editorial Office
St. Alban-Anlage 66, 4052 Basel, Switzerland
E-Mail: biology@mdpi.com <<mailto:biology@mdpi.com>>
<http://www.mdpi.com/journal/biology>

Wei-Chiao Chang <weichiao.chang@gmail.com>

Fri, Oct 23, 2020 at 1:15 AM

To: amy.tian@mdpi.com

Cc: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>, Wan Hsuan Chou <ocean.chou@tmu.edu.tw>, Wirawan
Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <mingyue@gmail.com>, Dyah Aryani Perwitasari
<diahperwitasari2003@yahoo.com>, Ben Kuen Chen <bkchen58@mail.ncku.edu.tw>, Hwai I Yang
<hiyang@gate.sinica.edu.tw>, Biology Editorial Office <biology@mdpi.com>

Hi Amy,
Many thanks for your mail.
I will re-submit our manuscript on 28th Oct.

Best regards,

wei chiao

[Biology] Manuscript ID: biology-998991 - Accepted for Publication

1 message

Amy Tian <amy.tian@mdpi.com>

Thu, Nov 5, 2020 at 7:50 PM

Reply-To: Amy Tian <amy.tian@mdpi.com>, Biology Editorial Office <biology@mdpi.com>

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Wan-Hsuan Chou <s700081@gmail.com>, Yu-Shiuan Wang <yswang1004@gmail.com>, Wirawan Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <miningyue@gmail.com>, Dyah Aryani Perwitasari <diahperwitasari2003@yahoo.com>, Wan Chen Huang <wanchen.huang@gmail.com>, Ben Kuen Chen <bkchen58@icloud.com>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Wei Chiao Chang <weichiao.chang@gmail.com>, Biology Editorial Office <biology@mdpi.com>, Amy Tian <amy.tian@mdpi.com>

Dear Mr. Muhammad Irham,

We are pleased to inform you that the following paper has been officially accepted for publication:

Manuscript ID: biology-998991

Type of manuscript: Article

Title: Evaluation for the Genetic Association Between Store-Operated Calcium Influx Pathway (STIM1 and ORAI1) and Human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Authors: Lalu Muhammad Irham, Wan-Hsuan Chou, Yu-Shiuan Wang, Wirawan Adikusuma, Henry Sung-Ching Wong, Dyah Aryani Perwitasari, Wan Chen Huang, Ben Kuen Chen *, Hwai I Yang *, Wei Chiao Chang *

Received: 28 October 2020

E-mails: lalu.irham@pharm.uad.ac.id, s700081@gmail.com, yswang1004@gmail.com, adikusuma28@gmail.com, miningyue@gmail.com, diahperwitasari2003@yahoo.com, wanchen.huang@gmail.com, bkchen58@icloud.com, hiyang@gate.sinica.edu.tw, weichiao.chang@gmail.com

Submitted to section: Genetics and Genomics,

https://www.mdpi.com/journal/biology/sections/Gene_Genomicshttps://susy.mdpi.com/user/manuscripts/review_info/7b243b2ed7d515d0740beb8ab173d2be

We will now make the final preparations for publication, then return the manuscript to you for your approval.

If, however, extensive English edits are required to your manuscript, we will need to return the paper requesting improvements throughout.

We encourage you to set up your profile at SciProfiles.com, MDPI's researcher network platform. Articles you publish with MDPI will be linked to your SciProfiles page, where colleagues and peers will be able to see all of your publications, citations, as well as your other academic contributions.

We also invite you to contribute to Encyclopedia (<https://encyclopedia.pub>), a scholarly platform providing accurate information about the latest research results. You can adapt parts of your paper to provide valuable reference information for others in the field.

Kind regards,
Ms. Amy Tian
Assistant Editor

--

MDPI Beijing Office Tongzhou, Jincheng Center, Room 2207, Tongzhou District, China

MDPI Biology Editorial Office
St. Alban-Anlage 66, 4052 Basel, Switzerland

[Biology] Manuscript ID: biology-998991 - Final Proofreading Before Publication

4 messages

Amy Tian <amy.tian@mdpi.com>

Fri, Nov 6, 2020 at 2:22 AM

Reply-To: amy.tian@mdpi.com

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Wan-Hsuan Chou <s700081@gmail.com>, Yu-Shiuan Wang <yswang1004@gmail.com>, Wirawan Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <miningyue@gmail.com>, Dyah Aryani Perwitasari <diahperwitasari2003@yahoo.com>, Wan Chen Huang <>wanchen.huang@gmail.com>, Ben Kuen Chen <bkchen58@icloud.com>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Wei Chiao Chang <weichiao.chang@gmail.com>, Biology Editorial Office <biology@mdpi.com>

Dear Mr. Muhammad Irham,

We invite you to proofread your manuscript to ensure that this is the final version that can be published and confirm that you will require no further changes from hereon:

Manuscript ID: biology-998991

Type of manuscript: Article

Title: Evaluation for the Genetic Association Between Store-Operated Calcium Influx Pathway (STIM1 and ORAI1) and Human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Authors: Lalu Muhammad Irham, Wan-Hsuan Chou, Yu-Shiuan Wang, Wirawan Adikusuma, Henry Sung-Ching Wong, Dyah Aryani Perwitasari, Wan Chen Huang, Ben Kuen Chen *, Hwai I Yang *, Wei Chiao Chang *

Received: 28 October 2020

E-mails: lalu.irham@pharm.uad.ac.id, s700081@gmail.com, yswang1004@gmail.com, adikusuma28@gmail.com, miningyue@gmail.com, diahperwitasari2003@yahoo.com, wanchen.huang@gmail.com, bkchen58@icloud.com, hiyang@gate.sinica.edu.tw, weichiao.chang@gmail.com

Submitted to section: Genetics and Genomics,

https://www.mdpi.com/journal/biology/sections/Gene_Genomics

Please read the following instructions carefully before proofreading:

1) Download the manuscript from the link provided at the end of this message and upload the final proofed version at the same link within 24 hours (1 working day). If you experience any difficulties, please contact the Biology Editorial Office.

2) Please use Microsoft Word's built-in track changes function to highlight any changes you make, or send a comprehensive list of changes in a separate document. Note that this is the *last chance* to make textual changes to the manuscript. Some style and formatting changes may have been made by the production team, please do not revert these changes.

3) All authors must agree to the final version. Check carefully that authors' names and affiliations are correct, and that funding sources are correctly acknowledged. Incorrect author names or affiliations are picked up by indexing databases, such as the Web of Science or PubMed, and can be difficult to correct.

After proofreading, final production will be carried out. Note that changes to the position of figures and tables may occur during the final steps. Changes can be made to a paper published online only at the discretion of the Editorial Office. In this case, a separate Correction or Addendum will be published and we reserve the right to charge 50 CHF per Correction (including changes to author names or affiliations).

[Biology] Manuscript ID: biology-998991 - Manuscript Resubmitted

1 message

Submission System <submission@mdpi.com>

Sun, Nov 8, 2020 at 9:19 PM

Reply-To: Amy Tian <amy.tian@mdpi.com>, Biology Editorial Office <biology@mdpi.com>

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Wan-Hsuan Chou <s700081@gmail.com>, Yu-Shiuan Wang <yswang1004@gmail.com>, Wirawan Adikusuma <adikusuma28@gmail.com>, Henry Sung-Ching Wong <miningyue@gmail.com>, Dyah Aryani Perwitasari <diahperwitasari2003@yahoo.com>, Wan Chen Huang <>wanchen.huang@gmail.com>, Ben Kuen Chen <bkchen58@icloud.com>, Hwai I Yang <hiyang@gate.sinica.edu.tw>, Wei Chiao Chang <weichiao.chang@gmail.com>

Dear Mr. Muhammad Irham,

Thank you very much for resubmitting the modified version of the following manuscript:

Manuscript ID: biology-998991

Type of manuscript: Article

Title: Evaluation for the Genetic Association Between Store-Operated Calcium Influx Pathway (STIM1 and ORAI1) and Human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection

Authors: Lalu Muhammad Irham, Wan-Hsuan Chou, Yu-Shiuan Wang, Wirawan Adikusuma, Henry Sung-Ching Wong, Dyah Aryani Perwitasari, Wan Chen Huang, Ben Kuen Chen *, Hwai I Yang *, Wei Chiao Chang *

Received: 28 October 2020

E-mails: lalu.irham@pharm.uad.ac.id, s700081@gmail.com, yswang1004@gmail.com, adikusuma28@gmail.com, miningyue@gmail.com, diahperwitasari2003@yahoo.com, wanchen.huang@gmail.com, bkchen58@icloud.com, hiyang@gate.sinica.edu.tw, weichiao.chang@gmail.com

Submitted to section: Genetics and Genomics,

https://www.mdpi.com/journal/biology/sections/Gene_Genomicshttps://susy.mdpi.com/user/manuscripts/review_info/7b243b2ed7d515d0740beb8ab173d2be

A member of the editorial office will be in touch with you soon regarding progress of the manuscript.

Kind regards,

MDPI

--

Biology Editorial Office

Postfach, CH-4020 Basel, Switzerland

Office: St. Alban-Anlage 66, CH-4052 Basel

Tel. +41 61 683 77 34 (office)

Fax +41 61 302 89 18 (office)

E-mail: biology@mdpi.com<https://www.mdpi.com/journal/biology/>

*** This is an automatically generated email ***

[Biology] Manuscript ID: biology-998991; doi: 10.3390/biology9110388. Paper has been published.

2 messages

Biology Editorial Office <biology@mdpi.com>

Mon, Nov 9, 2020 at 4:04 AM

Reply-To: Amy Tian <amy.tian@mdpi.com>, Biology Editorial Office <biology@mdpi.com>

To: Lalu Muhammad Irham <lalu.irham@pharm.uad.ac.id>

Cc: Biology Editorial Office <biology@mdpi.com>, Amy Tian <amy.tian@mdpi.com>

Dear Mr. Irham,

We are pleased to inform you that "Evaluation for the Genetic Association between Store-Operated Calcium Influx Pathway (STIM1 and ORAI1) and Human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection" by Lalu Muhammad Irham, Wan-Hsuan Chou, Yu-Shiuan Wang, Wirawan Adikusuma, Henry Sung-Ching Wong, Dyah Aryani Perwitasari, Wan-Chen Huang, Ben-Kuen Chen *, Hwai-I Yang *, Wei-Chiao Chang * has been published in Biology and is available online:

Abstract: <https://www.mdpi.com/2079-7737/9/11/388>HTML Version: <https://www.mdpi.com/2079-7737/9/11/388/htm>PDF Version: <https://www.mdpi.com/2079-7737/9/11/388/pdf>

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Kind regards,
Biology Editorial Office
Postfach, CH-4020 Basel, Switzerland
Office: St. Alban-Anlage 66, CH-4052 Basel
Tel. +41 61 683 77 34 (office)
Fax +41 61 302 89 18 (office)
E-mail: biology@mdpi.com
<https://www.mdpi.com/journal/biology/>

biology@mdpi.com <biology@mdpi.com>

Mon, Nov 9, 2020 at 4:06 AM

Reply-To: amy.tian@mdpi.com, biology@mdpi.com

To: lalu.irham@pharm.uad.ac.id, s700081@gmail.com, yswang1004@gmail.com, adikusuma28@gmail.com, miningyue@gmail.com, diahperwitasari2003@yahoo.com, wanchen.huang@gmail.com, bkchen58@mail.ncku.edu.tw, hiyang@gate.sinica.edu.tw, wcc@tmu.edu.tw

Cc: billing@mdpi.com, website@mdpi.com, biology@mdpi.com, amy.tian@mdpi.com

Dear Authors,

We are pleased to inform you that your article "Evaluation for the Genetic Association between Store-Operated Calcium Influx Pathway (STIM1 and ORAI1) and Human Hepatocellular Carcinoma in Patients with Chronic Hepatitis B Infection" has been published in Biology and is available online:

Abstract: <https://www.mdpi.com/2079-7737/9/11/388>HTML Version: <https://www.mdpi.com/2079-7737/9/11/388/htm>

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Wei-Chiao Chang

Department of Clinical Pharmacy, School of
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