



## PEER-REVIEW REPORT

**Name of journal:** World Journal of Clinical Oncology

**Manuscript NO:** 33969

**Title:** Further the liquid biopsy: gathering pieces of the puzzle of Genometastasis Theory

**Reviewer's code:** 00503561

**Reviewer's country:** Japan

**Science editor:** Yuan Qi

**Date sent for review:** 2017-06-23

**Date reviewed:** 2017-07-06

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

### COMMENTS TO AUTHORS

Please add the concept of "genometastasis" is a new idea (according to the reference 79) in the abstract. This is very interesting, provocative, and newly twisted idea, possibly raise an argument in the community. I love this kind of statement in a review article and hope the text should be gradual and introductory (junctional part in the section 2.4, and 3 in the flow of the manuscript).



**PEER-REVIEW REPORT**

**Name of journal:** World Journal of Clinical Oncology

**Manuscript NO:** 33969

**Title:** Further the liquid biopsy: gathering pieces of the puzzle of Genometastasis Theory

**Reviewer’s code:** 00503399

**Reviewer’s country:** Greece

**Science editor:** Yuan Qi

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CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input checked="" type="checkbox"/> Rejection
<input checked="" type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input checked="" type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input checked="" type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

**COMMENTS TO AUTHORS**

In this manuscript the authors discuss the uses of liquid biopsies in cancer diagnosis and metastases detection and they focus in their theory (genometastases) which supports that tumor DNA can enter normal cells genetic material leading to the introduction of genetic changes leading to cancer. In more than half of the paper the authors are focused in explaining the uses of liquid biopsies similarly to other reviews and editorials in the literature. “Karachaliou, Niki et al. “Real-Time Liquid Biopsies Become a Reality in Cancer Treatment.” *Annals of Translational Medicine* 3.3 (2015): 36. PMC. Web. 6 July 2017. Pancreatic cancer: Are "liquid biopsies" ready for prime-time? Alexandra R Lewis, Juan W Valle, Mairead G McNamara *World J Gastroenterol*. 2016 Aug 28; 22(32): 7175-7185. Circulating tumor DNA as a liquid biopsy target for detection of pancreatic cancer Erina Takai, Shinichi Yachida *World J Gastroenterol*. 2016 Oct 14; 22(38): 8480-8488. Circulating tumor cells versus circulating tumor DNA in lung cancer—which one will win? Silvia Calabuig-Fariñas, Eloísa Jantus-Lewintre,



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Alejandro Herreros-Pomares, Carlos Camps Transl Lung Cancer Res. 2016 Oct; 5(5): 466–482.” In the other half of the paper the authors try to present their theory of free tumor DNA that can enter normal cells and can lead to metastases and they suggest that the DNA detected from liquid biopsies may be a useful tool to predict metastases (the “genometastases” phenomenon). In my personal opinion the primary target of this group is to prove the genometastases phenomenon since there are many holes in our knowledge in this aspect and then write a review of how to detect it.



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**Name of journal:** World Journal of Clinical Oncology

**Manuscript NO:** 33969

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**Reviewer's country:** China

**Science editor:** Yuan Qi

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CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input checked="" type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
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<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

**COMMENTS TO AUTHORS**

This a very good review for application of liquid biopsy, it not only constitutes a promising tool for cancer diagnostic and patient follow-up but also it may help in the comprehension of metastasis. With this technique, it was found that CTCs are limited in blood, and circulating nucleic acids are much more abundant. Together with the demonstrated capability of circulating nucleic acids to transform susceptible cells, strongly support the theory of genometastasis which sustains that cancer propagation relies on gene transfer from malignant cells to normal cells.