

<u>Apostolou P.¹</u>, Ntanovasilis D.A. ¹, Papasotiriou I.¹, ¹Research Genetic Cancer Centre S.A. Industrial Area of Florina, 53100 Florina, Greece





Liquid Biopsy, personalized treatment and response to Hyperthermia: **Challenges and Perspectives**

Contents



- Hyperthermia and Cancer
 - Heat Shock Proteins
 - Liquid biopsy and CTCs
 - Study Results

Hyperthermia





(Skitzki et al. 2009)

Heat Shock Proteins





(Chatterjee, Burns Int. J. Mol. Sci. 2017)

HSP27 (HSPB1):

Up-Regulation associated with

- Cancer progression
- Metastasis
- Poor prognosis
- Resistance

Heat Shock Proteins Centre Research Genetic Cancer Cell growth R Survival (CD44) Motility Apoptosis Hsp70 Invasion (mitochondria, lysosomes) (Met, Smad2, tTG, FAK, Wasf3, MMPs, endocytosis) Inflammation & Angiogenesis

HSP70 (HSPA1A):

- Metastasis
- Anti-apoptotic properties

Heat Shock Proteins





(Eskander, Tewari, Journal of Gynecologic Oncology 2014)

HSP90:

• Implicated in all hallmark traits of cancer

Liquid Biopsy and CTCs



(Chaffer, Weinberg Science, 2011)



Liquid Biopsy-Detection

- Bead-Base method
- Microscopy-based method
- Size-based method
- PCR-based method

- Viability?
- Purity?
- Noise?

Liquid Biopsy-Detection

- Using parameters like FS, SS and fluorescence we can detect multiple antigens inside each cell.
- There are two approaches to detect CTCs: positive selection and negative selection.
- FC can provide information about quantity and quality of CTCs

Journal of Cancer Therapy, 2015, 6, 543-553 Published Online July 2015 in SciRes. <u>http://www.scirp.org/journal/jct</u> <u>http://dx.doi.org/10.4236/jct.2015.67059</u>

5. Conclusion

In conclusion, this study demonstrates that it is possible to detect CTCs with higher sensitivity (86.2%) and specificity (83.9%) compared with routine clinical methodologies. The parameters may vary depending on the antibody panel used; however, using flow cytometry to identify CTCs has proven to be efficient. These results suggest that further studies are required to improve the accuracy by which CTCs and CTC subtypes can be identified by flow cytometry and thereby improve our ability to detect and follow the progression of cancer.







Samples Tested





^{*} Breast CTCs per 7.5ml- Other types per ml

















Concluding Remarks



- HSP correlated with resistance to hyperthermia
- Liquid biopsy based on CTCs study can be used for prediction response in different types of therapy
- CTCs study for thermo-tolerance

Selected Reference



- Lianos GD, Alexiou GA, et al. The role of heat shock proteins in cancer. Cancer Lett. 2015 May 1;360(2):114-8. doi: 10.1016/j.canlet.2015.02.026. Epub 2015 Feb 23.
- Skitzki JJ, Repasky EA, and Evans SS. Hyperthermia as an immunotherapy strategy for cancer. Curr Opin Investig Drugs. 2009 Jun;10(6):550-8.
- Horowitz M, Robinson SD. Heat shock proteins and the heat shock response during hyperthermia and its modulation by altered physiological conditions. Prog Brain Res. 2007;162:433-46.



THANK YOU FOR YOUR TIME AND PATIENCE

IN CASE OF QUESTIONS AND INQUIRIES PLEASE CONTACT WITH

question[at]rgcc-genlab.com