

CENTRA

Computational Oncology Training Alliance

ESR 9 - Evolution of drug resistance on genetic and phenotypic levels

Research project	Tumours often show astonishing genetic and phenotypic plasticity, which drives their capability to evolve into drug-resistant states. The acquisition of drug resistance is the main limitation of targeted cancer therapy. Tumours can grow drug tolerant either a) due to selective pressure on cells carrying specific mutations or b) due to phenotypic reprogramming of cells by external environmental factors imposed by the drugs. The latter may be further shaped by the cellular neighbourhoods, such as the density of the web of stromal cells surrounding the tumour and protecting from the drug. This project aims to investigate how advanced models of tumour evolution should be adapted to better explain risk factors leading to treatment resistance and failure.	
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Host institution	University of Warsaw, Poland Faculty of Mathematics, Informatics and Mechanics	
PhD program	Faculty of Mathematics, Informatics and Mechanics; Institute of Informatics	
Expected results	Mathematical models of tumour evolution, cell composition and emergence of drugresistant mutations Methods to assess the risk of presence of resistant genotypes prior to the start of therapy	
Planned secondments	1) Merck to	learn about drug resistance and its analysis (3 months)
	2) Ardigen to learn software development and large data analysis (1 month)	
	3) KCL-Cri data (3 mo	ck -Ciccarelli to learn resistance modeling and integrate with colorectal cancer nths)
Required profile	Solid background in statistics, data analysis, mathematical modeling, excellence and experience in programming, fluent English, interest in molecular biology and cancer evolution	