

Diffuse large B cell lymphoma (DLBCL) in vitro model based on a newly synthesized chitosan/gelatin hydrogel

Gaia Corallo¹, Alessandro Polini², Susanna Anita Pappagallo³, Giuseppe Gigli^{1,2}, Antonella Stanzione², Francesca Scalerà², Maria Carmela Vegliante³, Sabino Ciavarella³, Attilio Guarino³, Francesca Gervaso²

¹ Dipartimento di Matematica e Fisica E. De Giorgi, University of Salento, Campus Ecotekne, via Monteroni, 73100 Lecce, Italy

² CNR NANOTEC – Institute of Nanotechnology c/o Campus Ecotekne, via Monteroni, 73100 Lecce, Italy

³ Hematology and Cell Therapy Unit, IRCCS-Istituto Tumori ‘Giovanni Paolo II’, Bari Italy

Diffuse large B cell lymphoma (DLBCL) is the most common non-Hodgkin lymphoma, accounting for 30% of diagnoses worldwide. The advances in molecular biology techniques have been fundamental to achieve maximum understanding of DLBCL pathophysiology, but conventional 2D cell culture and mice models are not representative of the in vivo physiology. 2D cultures do not mimic growth profiles and cellular organization observed in vivo and lack the heterogeneity of tumor microenvironment (TME). Consequently, extracellular matrix (ECM) analogues can be designed to provide cells with a 3D structure and obtain a more faithful representation of the TME. In this work a chitosan/gelatin-based hydrogel has been developed to provide lymphoma cells with a 3D structure and obtain a more faithful representation of the TME. U2932 lymphoma cells have been encapsulated in the hydrogel, alone or in combination with WPMY-1 stromal cells, showing high viability by live/dead assay. Furthermore, co-culture studies showed the formation of cell spheroids after three days of culture, with respect to U2932 cells encapsulated alone. Based on these results, this hydrogel represents an excellent candidate for establishing a 3D in vitro model of DLBCL.