

Interaction of new fluorescent 2-quinolinone and coumarin derivatives with phospholipid monolayers and lipid vesicles

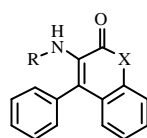
Ana S. Abreu^{1,2,*}, Elisabete M. S. Castanheira¹, B. F. Hermenegildo¹, Maria-João R.P. Queiroz² and Paula M.T. Ferreira²

¹Centro de Física (CFUM), Univ. do Minho, Campus de Gualtar, 4710-057 Braga, Portugal.

²Centro de Química (CQ-UM), Univ. do Minho, Campus de Gualtar, 4710-057 Braga, Portugal.

*anabreu@quimica.uminho.pt

Molecular interactions between organic molecules and phospholipids of various chain lengths have been investigated, either with monolayers at the air-interface or with bilayer vesicles (liposomes) as models of cell membranes [1]. In the present work, the interaction with biomembrane models of a fluorescent 3-amino-4-phenylquinolin-2-one **1** and a 3-(*tert*-butoxycarbonyl)amino-4-phenylcoumarin **2** (Fig. 1), previously synthesized by us [2], were studied. Interactions of both compounds with phospholipid monolayers of egg-yolk phosphatidylcholine (Egg-PC), dipalmitoyl phosphatidylcholine (DPPC) and dipalmitoyl phosphatidylglycerol (DPPG) has been studied by the Langmuir-Blodgett technique (Fig. 2).



1. R = H, X = NH
2. R = Boc, X = O

Figure 1. Structure of compound **1** and **2**.

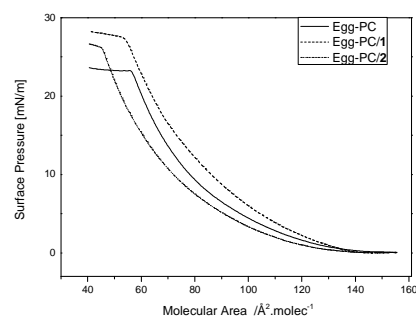


Figure 2: Surface pressure/molecular area isotherms of Egg-PC, Egg-PC/**1** and Egg-PC/**2** at the air-water interface at 22 °C.

Fluorescence emission and anisotropy measurements of **1** and **2** in lipid vesicles were performed below (gel phase) and above (liquid-crystalline phase) the lipid melting transition temperature (Table 1) in order to obtain information about compound interactions with the lipid membranes.

Table 1. Steady-state fluorescence anisotropy (r) values and maximum emission wavelengths (λ_{em}) for compounds **1** and **2** in lipid membranes.

Lipid Membranes	T (°C)	1		2	
		λ_{em}/nm	r	λ_{em}/nm	r
Neat Egg-PC	25	398	0.088	399	0.216
Neat DPPC	25	398	0.059	400	0.164
	55	398	0.045	400	0.149
Neat DPPG	25	394, 509 <i>sh</i>	0.023	400	0.146
	55	394, 503 <i>sh</i>	0.012	398	0.119
DPPC/DPPG (1:1)	25	394, 500 <i>sh</i>	0.025	397	0.177
	55	397, 502 <i>sh</i>	0.012	396	0.157

Acknowledgements: FCT-Portugal, QREN and FEDER/COMPETE through CFUM, CQ-UM, Project PTDC/QUI/81238/2006 and Post-doc. grant of Ana S. Abreu (SFRH/BPD/24548/2005).

[1] Peetla C., Stine A., Labhassetwar V., *Molecular Pharmaceutics* **2009**, 6, 1264-1276.

[2] Queiroz M.-J.R.P., Abreu A.S., Calhelha R.C., Carvalho M.S.D., Ferreira P.M.T., *Tetrahedron* **2008**, 64, 5139-5146.