



## Non Photochemical Light Induced Nucleation

Anne Spasojevic - de Biré, Aziza Ikni, Bertrand Clair, Nicolas Guiblin, Igor A. Kornev, Xiaoxuan Shi, Jianyao Long, Hongxun Yang, Wenjing Li, Annais Mendre, et al.

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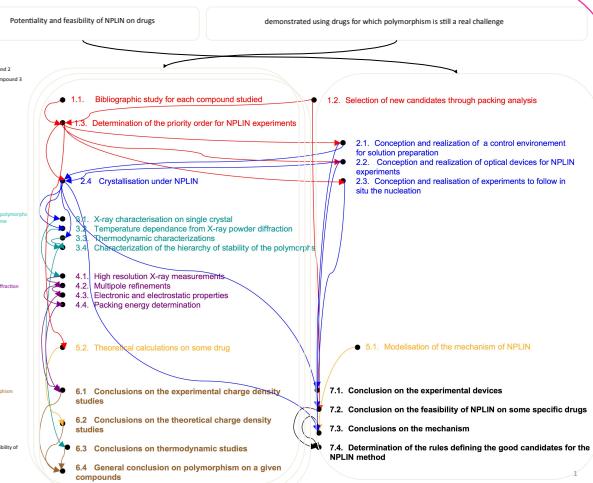
# NPLIN-4-drug

## Non Phototchemical Light Induced Nucleation for drugs

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## Journées Nationales en Nanosciences et Nanotechnologies 2011



### Goals and tasks of the project

#### Polymerism studies important in the industrial pharmaceutical context

Method developed on urea [1], glycine [2], L-histidine [3], Hen Egg White Lysozyme [4], Pentyl cyanobiphenyl [5], KCl [6], protein [7]

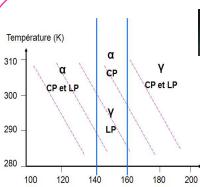
#### Context



- Co-existence of clusters of polarisability tensors of different symmetry: « disk-like » or « rod-like »
- Role of the induced dipolar moment depending on the clusters type

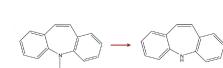
$$\frac{N(\alpha_{\text{parallel}} - \alpha_{\text{perpendiculaire}})^2}{45\epsilon_0 n_0 K_b T}$$

#### Possible mechanism : Kerr effect



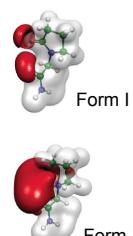
- Switch window size depending on the laser intensity

#### Glycine (task 2.4)

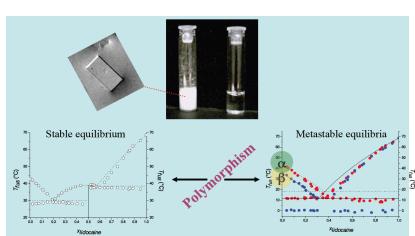


- Essay of crystallization of an anti-epileptic highly polymorphic
- Three different crystals of carbamazepine polymorph III
- 1 and 2 obtained by NPLIN, 3 is a spontaneous crystallization

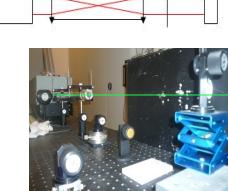
#### Carbamazepine (task 2.4)



#### Piracetam (task 4.)



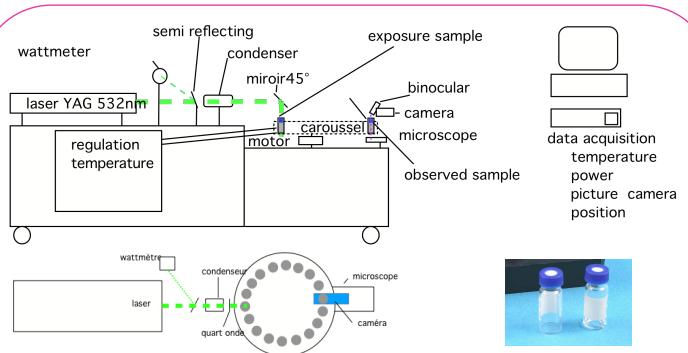
#### Lidocaïne / menthol (task 3.3 3.4)



YAG-Nd laser  $\lambda = 532$  nm, focalization under  $N_2$ , exposition time = 1 mn, energy density = 0.768 GW/cm<sup>2</sup>

Single crystals obtained after 1 or 10 hours

#### NPLIN old experimental devices



#### NPLIN new experimental device (task 2.1, 2.2, 2.3)

[1] J. Prevost, H. Yang, F. Dumas, P. Scouflaire, A. Spasojević – de Biré, « Non Phototchemical Light Induced Nucleation. A tool to crystallize polymorphs on demand ? Application to carbamazepine », XVII Conference of Serbian Crystallographic Society, Ivajnica (Ser), juin 2010

[2] B. Clair, A. Iknî, H. Yang, F. Dumas, P. Scouflaire, A. Spasojević – de Biré « Non Phototchemical Light Induced Nucleation. A tool to crystallize polymorphs on demand ? » CPOSS meeting, London, Grande-Bretagne, 30 mars 2011

[3] A. Spasojević, H. Yang, F. Dumas, P. Scouflaire, A. Spasojević – de Biré « Non Phototchemical Light Induced Nucleation. A tool to crystallize polymorphs on demand ? » Congrès algérien de cristallographie, Kechela, Algérie, 14-16 mai 2011.

[4] Y. Corvis, P. Niguer, P. Espeau, « Phototchemical stability of solid dispersions of a drug and its racemic mixture in stearic acid », J. Pharm. Sci., 2011, 100, 5235-5243.

[5] M.H. Chambrier, N. Boulimida, F. Bonhomme, S. Lebegue, J.M.Gillet, C. Jelsch, N.E. Ghermani « Electron and Electrostatic Properties of Three Crystalline Forms of Piracetam » Crystal Growth & Design, 2011, 11 (6), 2528-2539

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