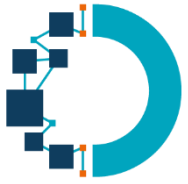


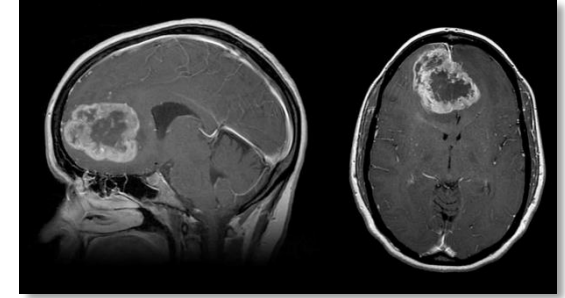
Radiosensibilisation de lignées tumorales gliales par des précurseurs de l'hème et thérapie génique

Mario Terlizzi

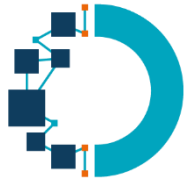




Contexte – le Glioblastome

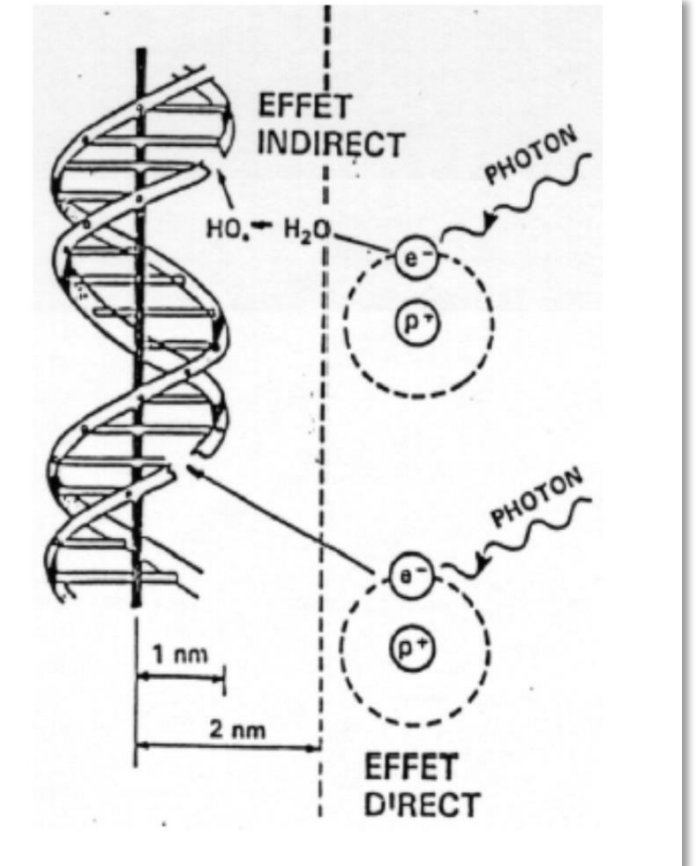
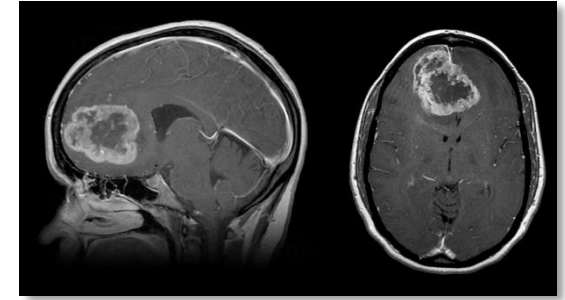


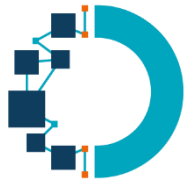
- 1^{ère} tumeur cérébrale maligne chez l'adulte en France ¹
- Standard thérapeutique = Chirurgie + RTCT adjuvante (**Protocole Stupp**) ²
- Mécanismes de **radiorésistance**: hypoxie, cellules souches, altérations génétiques ³



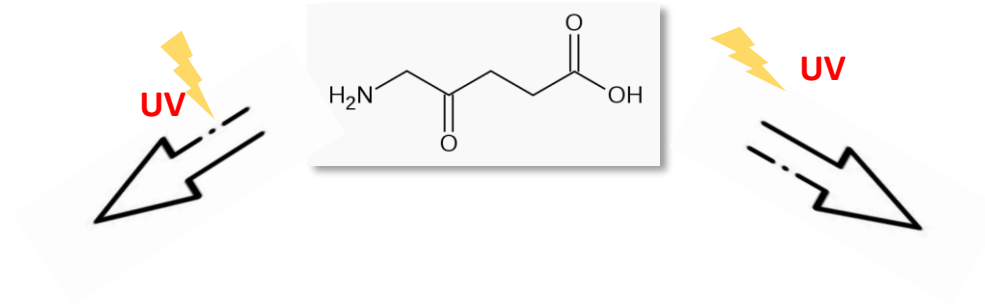
Contexte – le Glioblastome

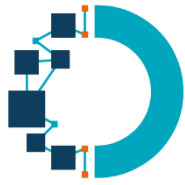
- 1^{ère} tumeur cérébrale maligne chez l'adulte en France ¹
- Standard thérapeutique = Chirurgie + RTCT adjuvante (**Protocole Stupp**) ²
- Mécanismes de **radiorésistance**: hypoxie, cellules souches, altérations génétiques ³



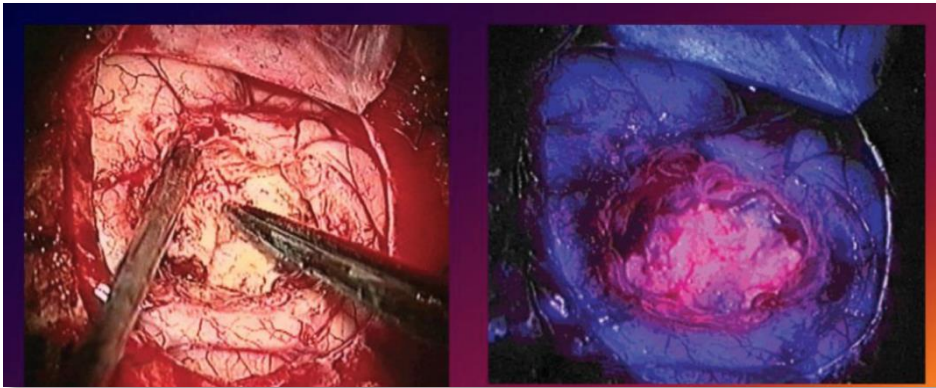
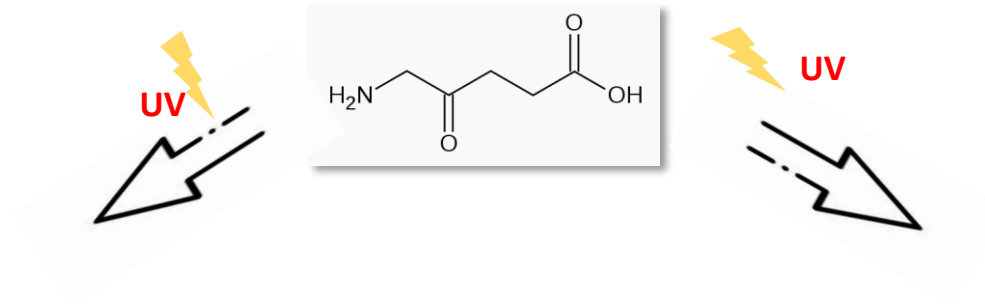


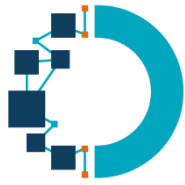
Rationnel: 5-ALA



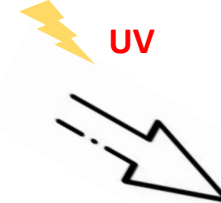
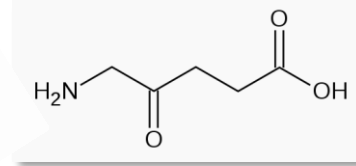


Rationnel: 5-ALA

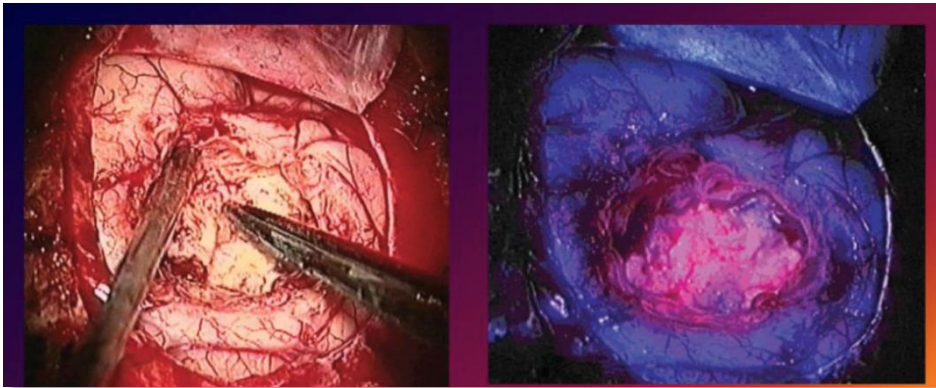


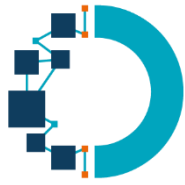


Rationnel: 5-ALA

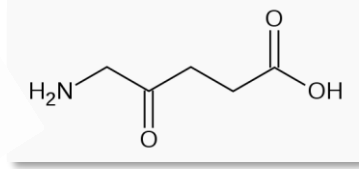


FLUORESCENCE

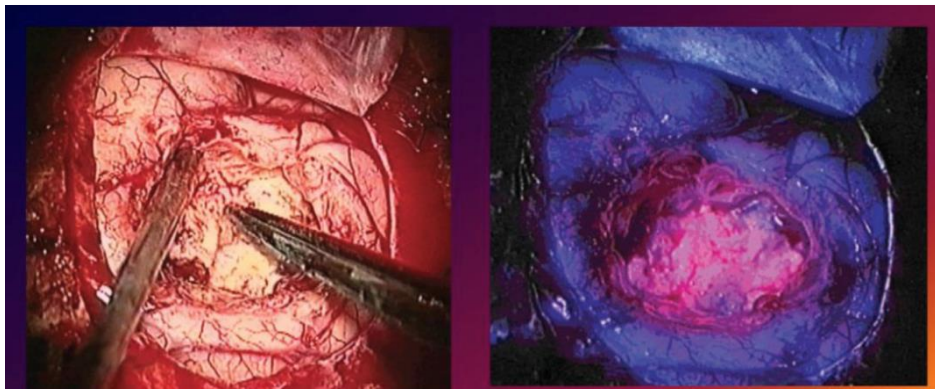




Rationnel: 5-ALA

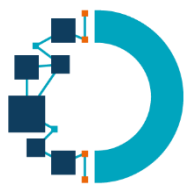


FLUORESCENCE

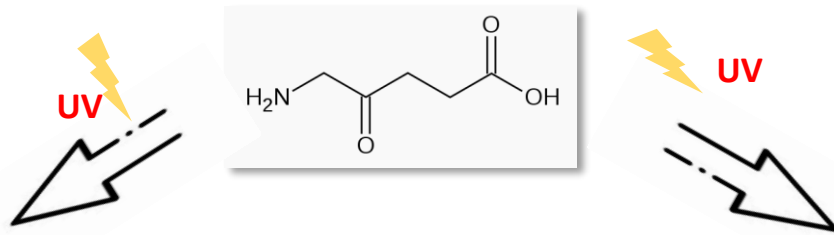


COOPERATION SPATIALE





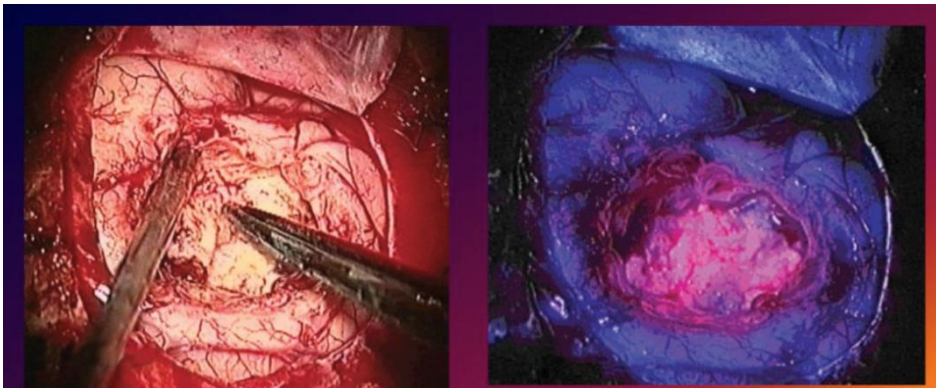
Rationnel: 5-ALA



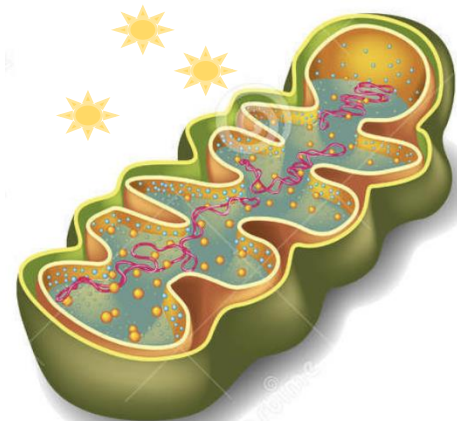
FLUORESCENCE

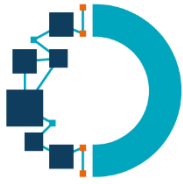


STRESS OXYDATIF

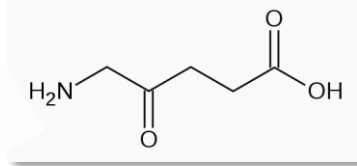


COOPERATION SPATIALE





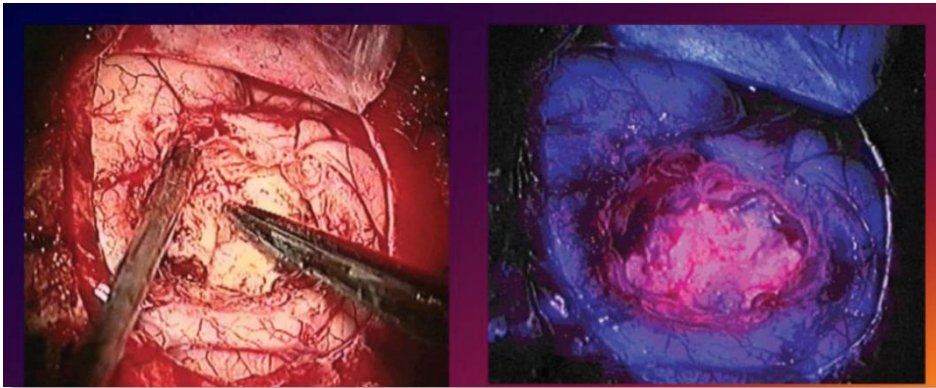
Rationnel: 5-ALA



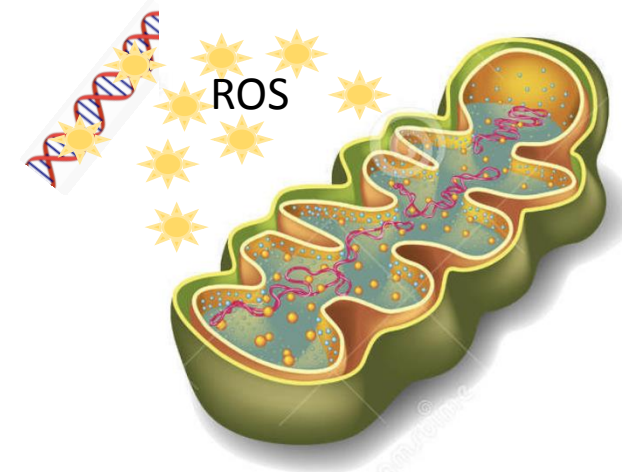
FLUORESCENCE

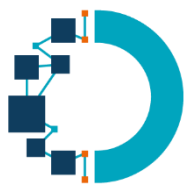


STRESS OXYDATIF

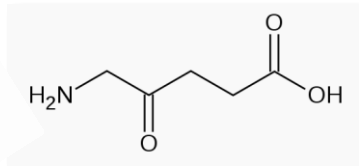


COOPERATION SPATIALE





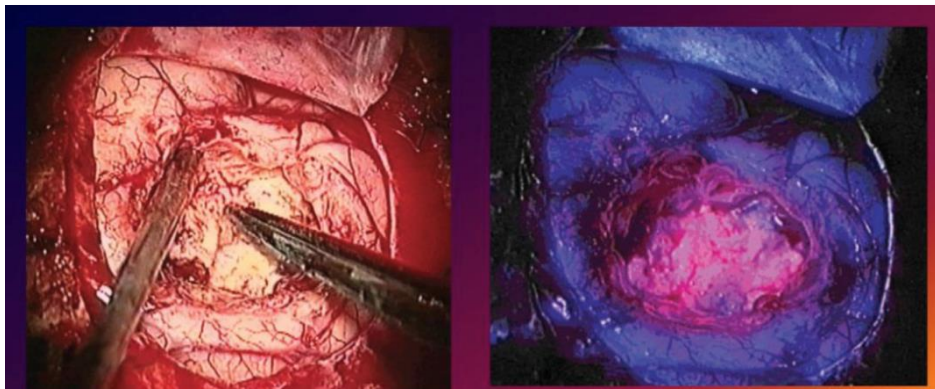
Rationnel: 5-ALA



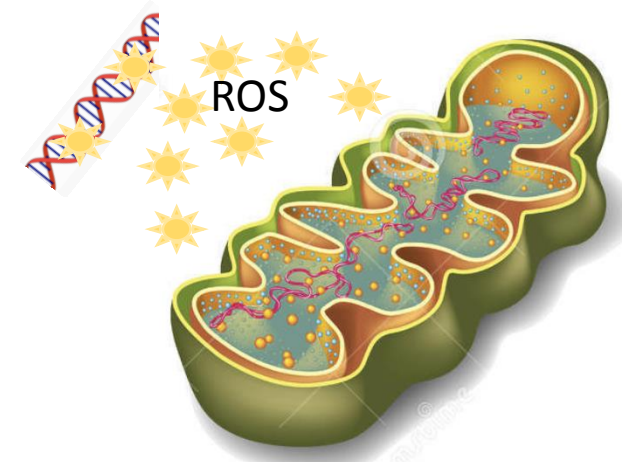
FLUORESCENCE



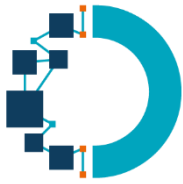
STRESS OXYDATIF



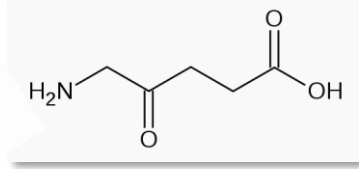
COOPERATION SPATIALE



COOPERATION CHIMIQUE



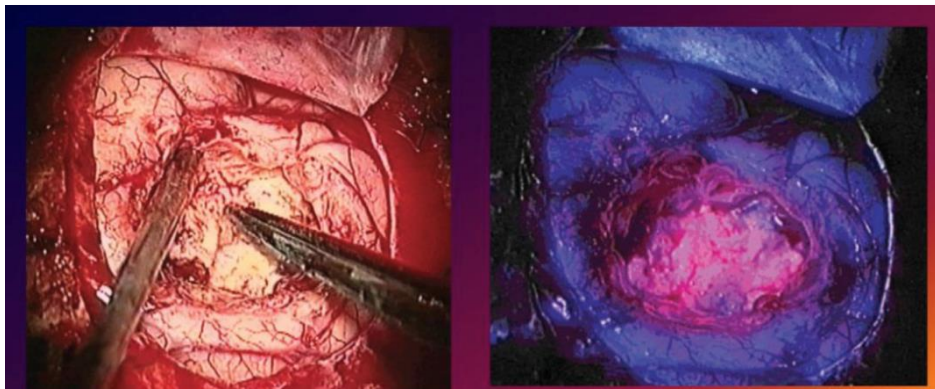
Rationnel: 5-ALA



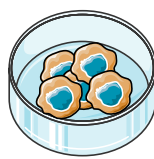
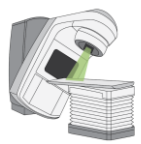
FLUORESCENCE



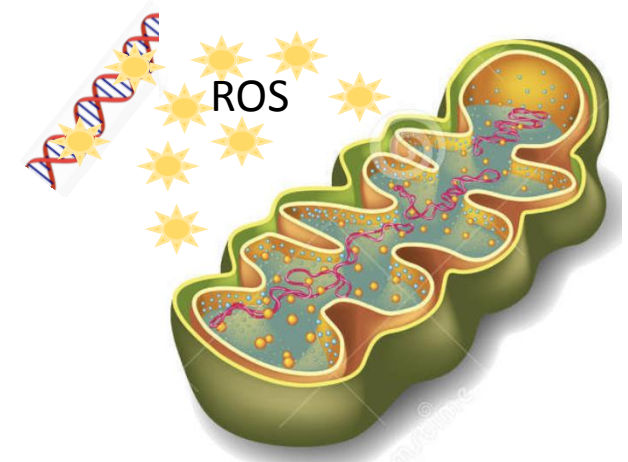
STRESS OXYDATIF



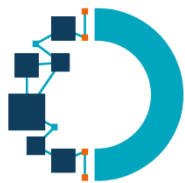
COOPERATION SPATIALE



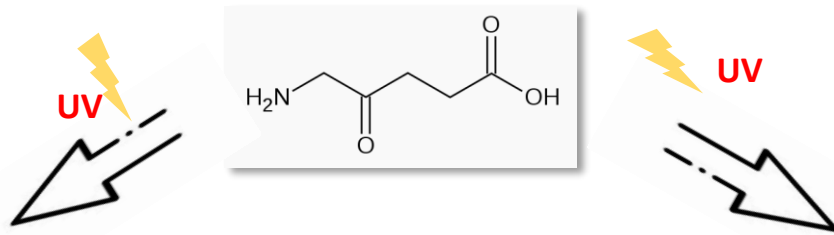
U87



COOPERATION CHIMIQUE



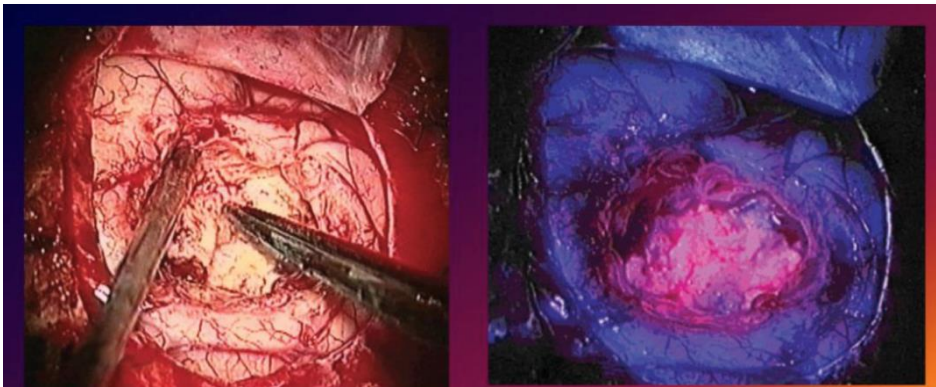
Rationnel: 5-ALA



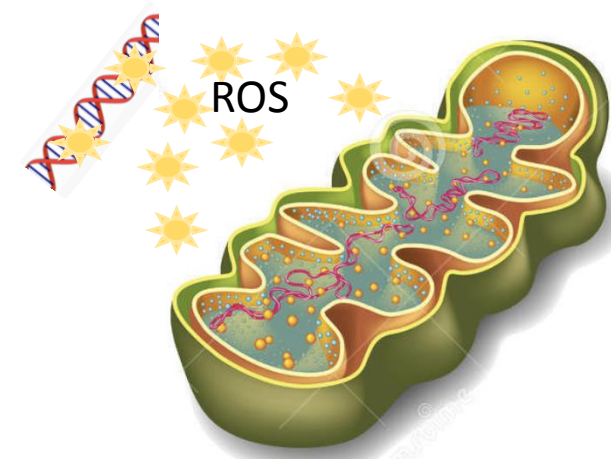
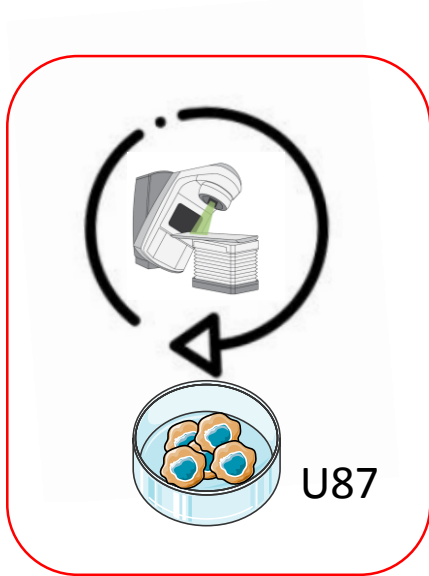
FLUORESCENCE



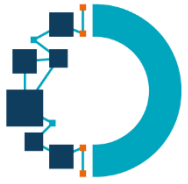
STRESS OXYDATIF



COOPERATION SPATIALE

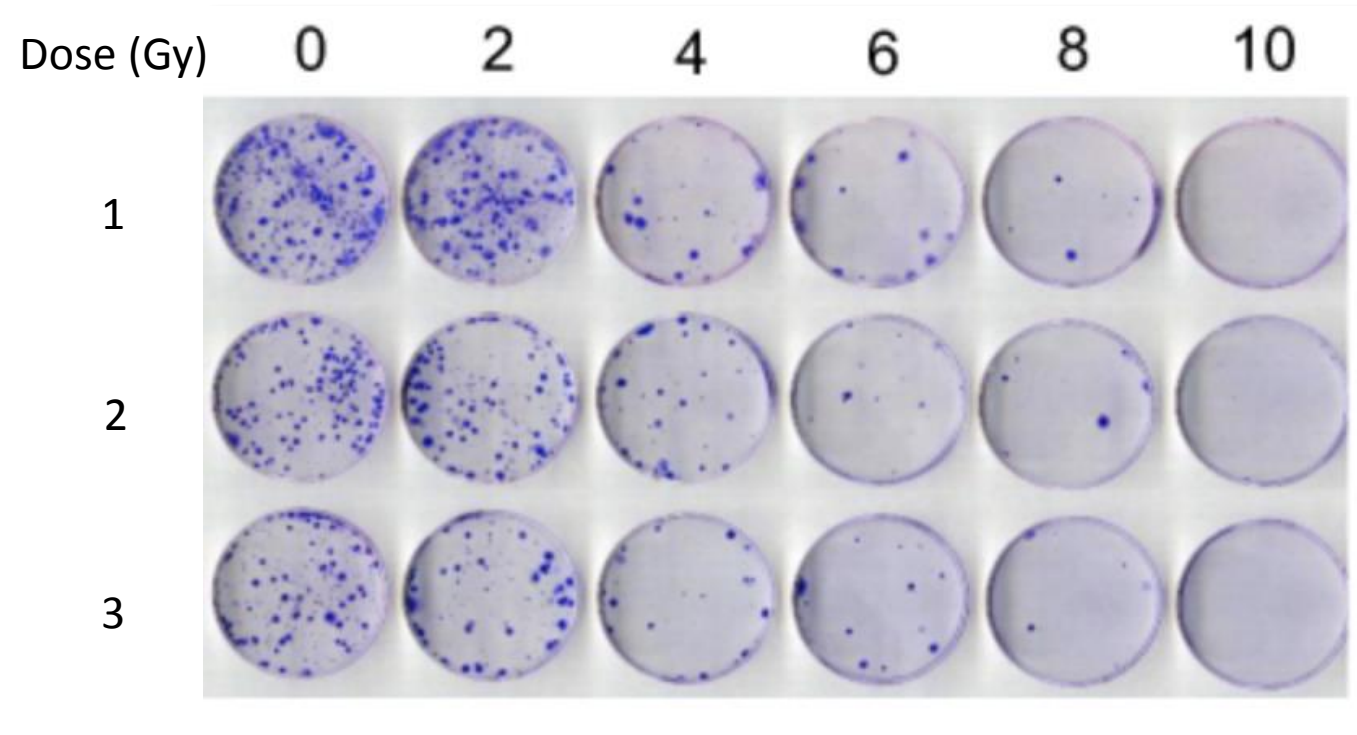


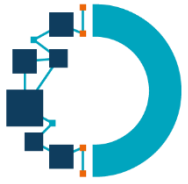
COOPERATION CHIMIQUE



Matériel et méthodes

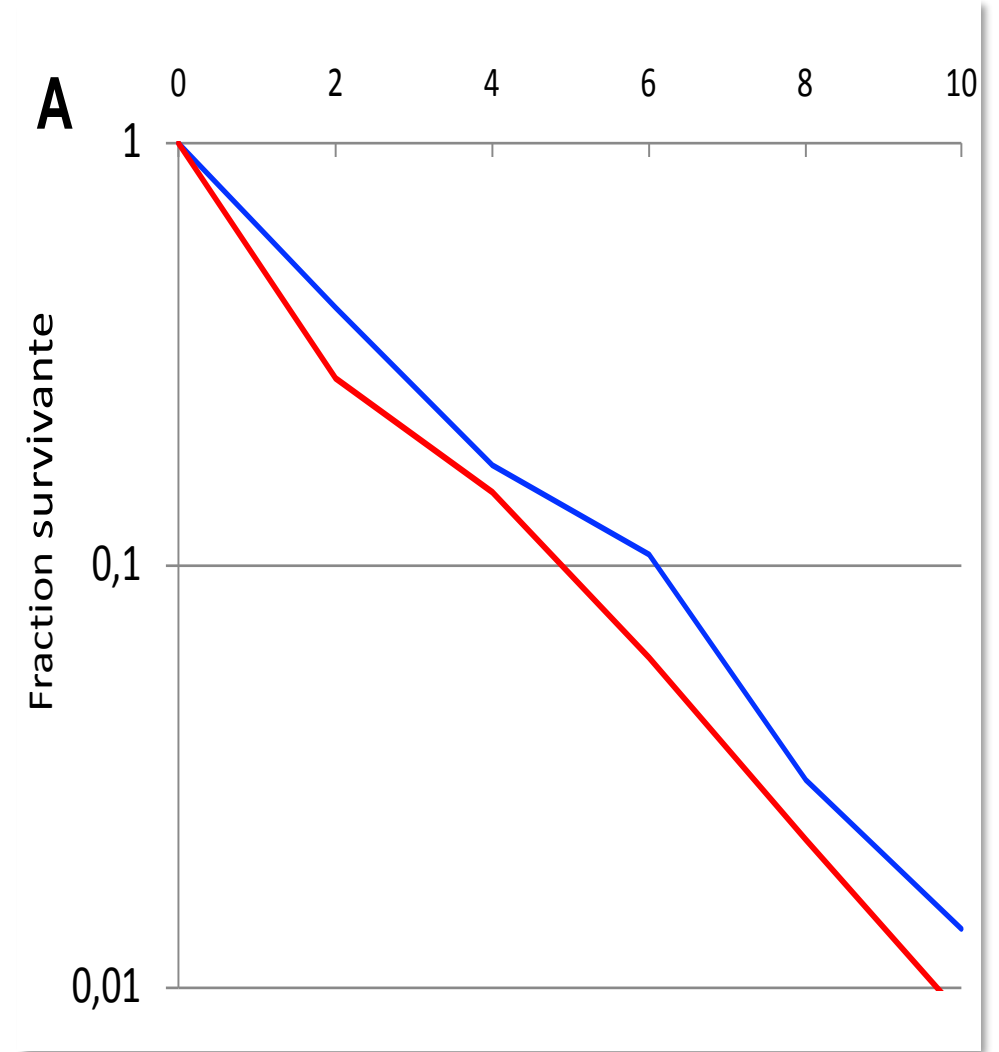
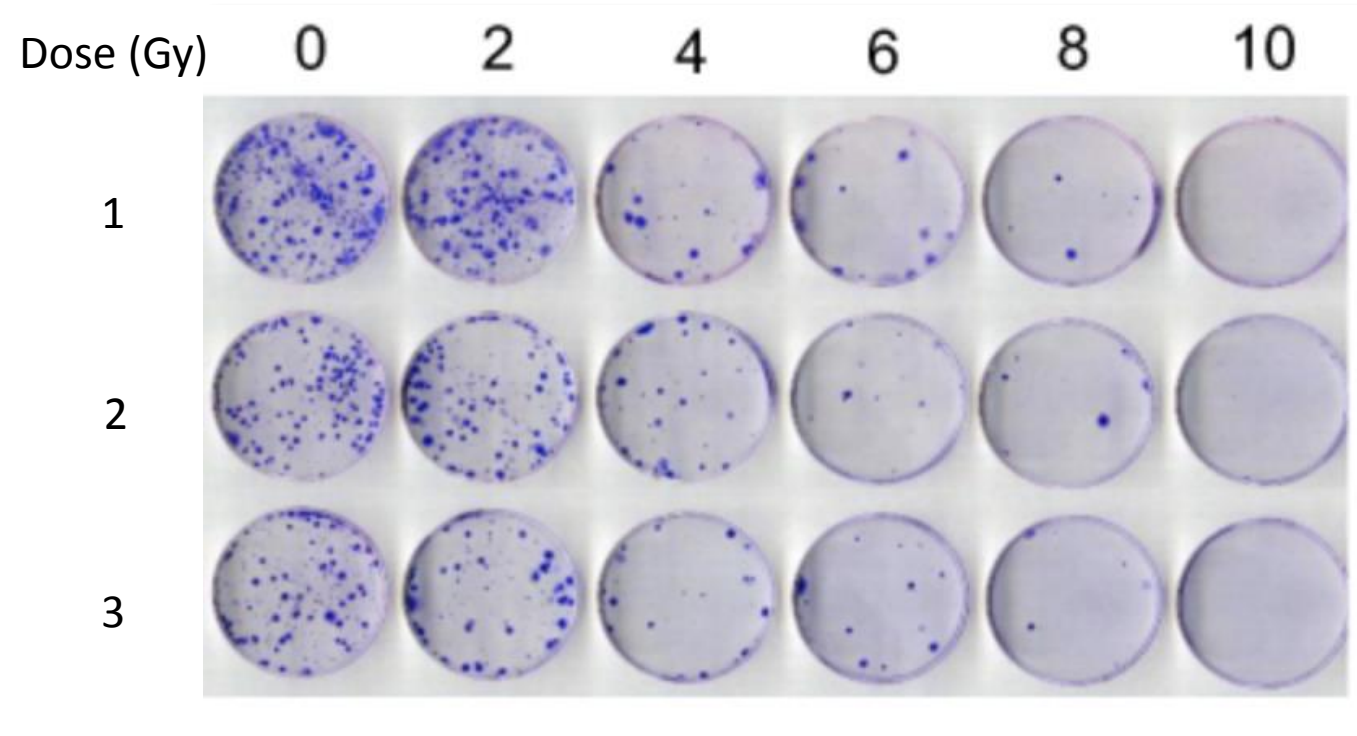
Tests de clonogénicité

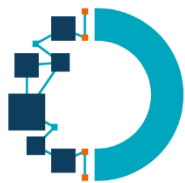




Matériel et méthodes

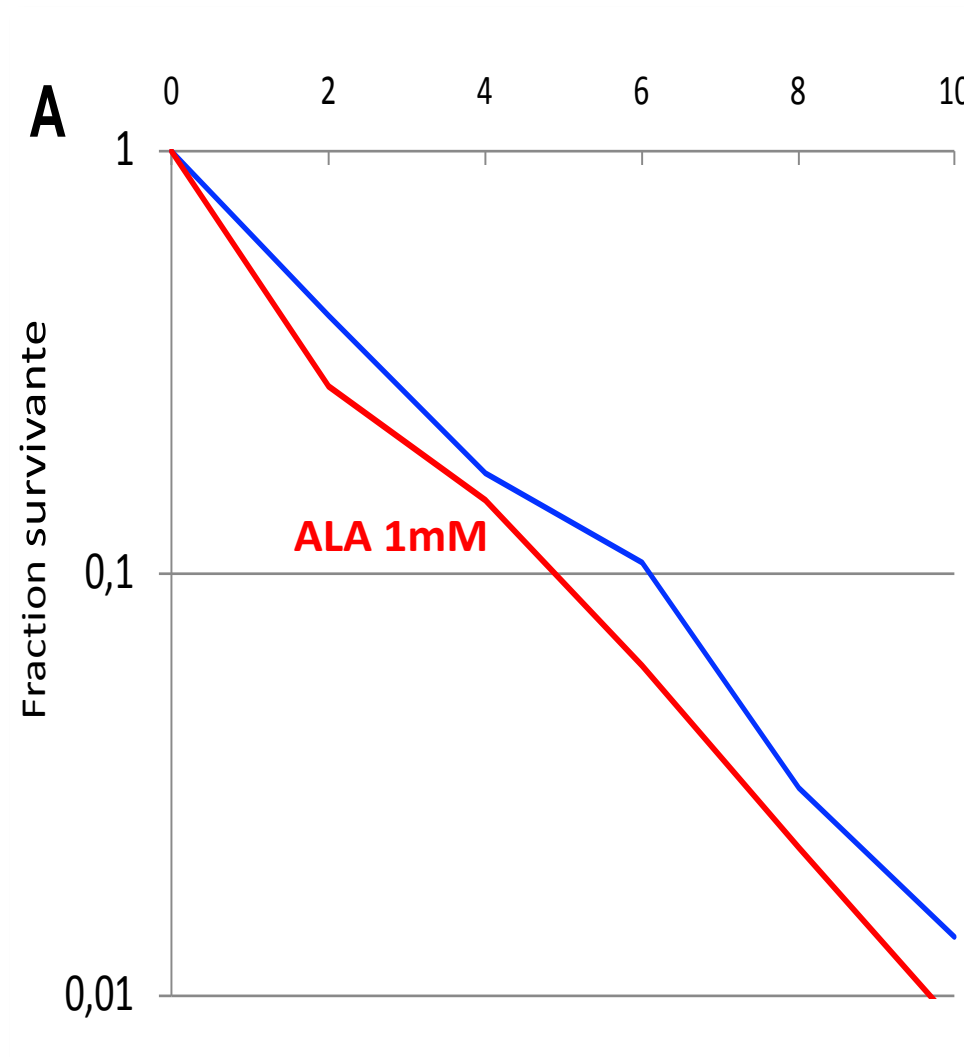
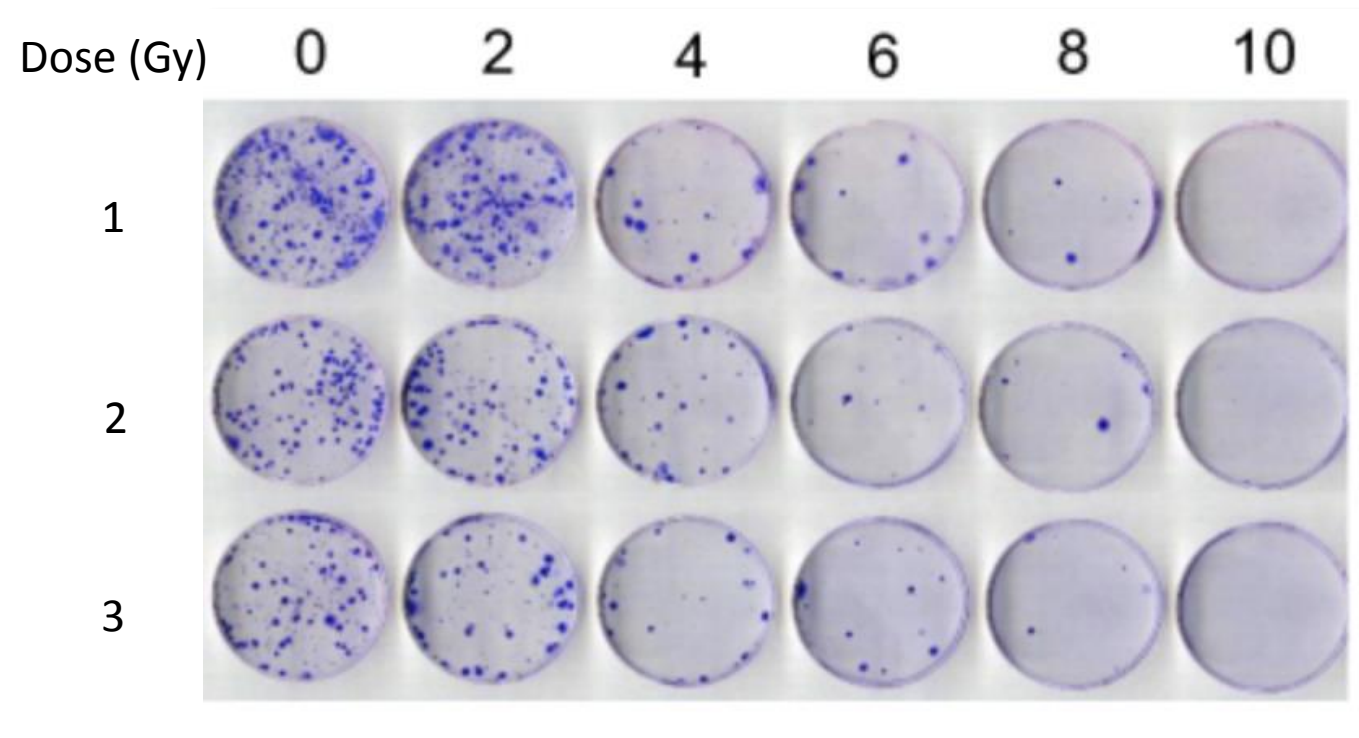
Tests de clonogénicité

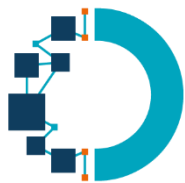




Matériel et méthodes

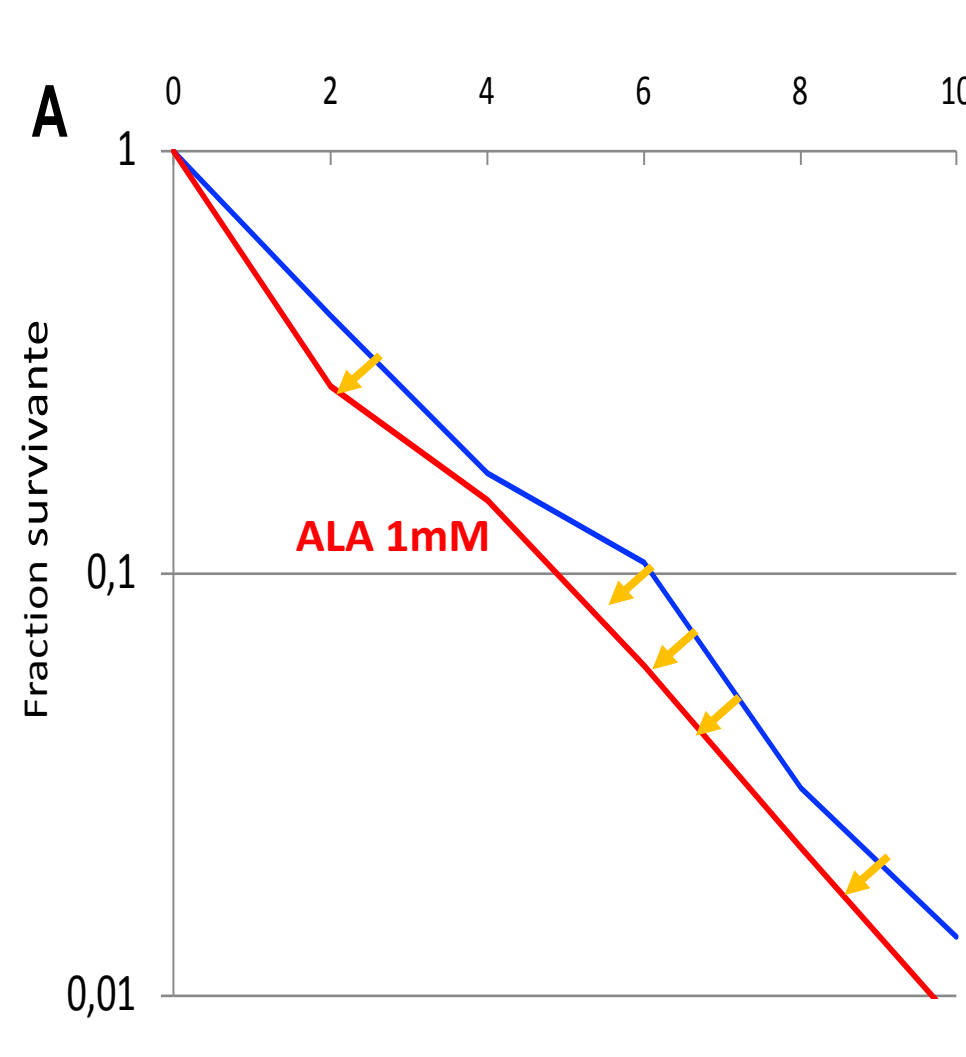
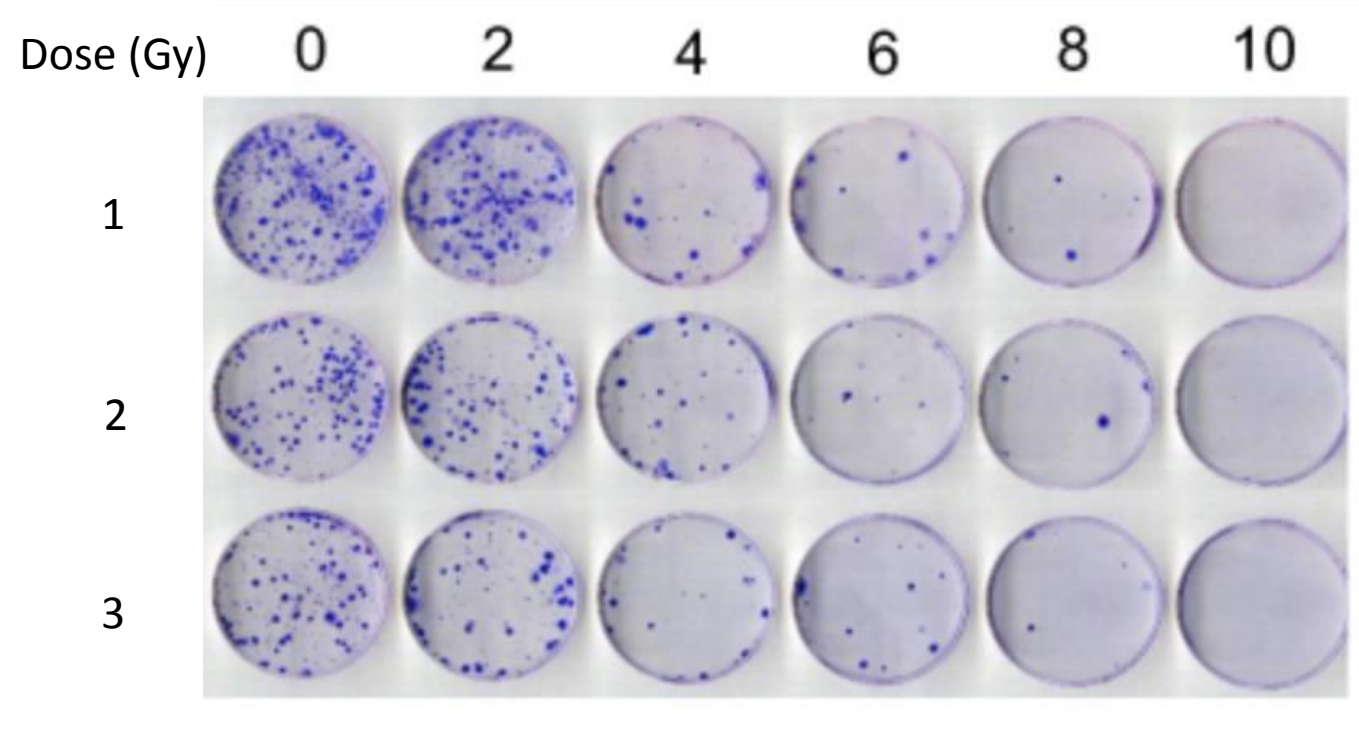
Tests de clonogénicité

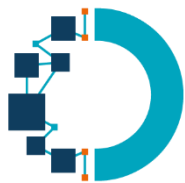




Matériel et méthodes

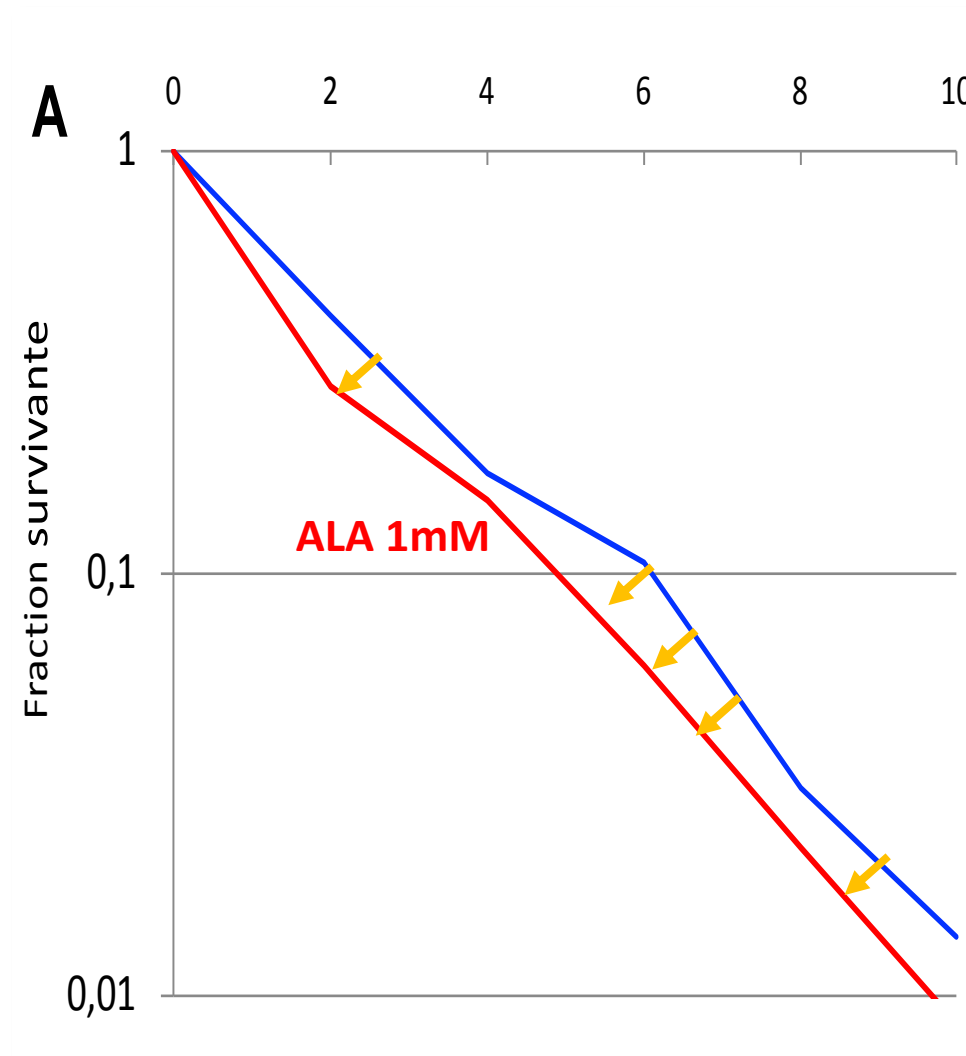
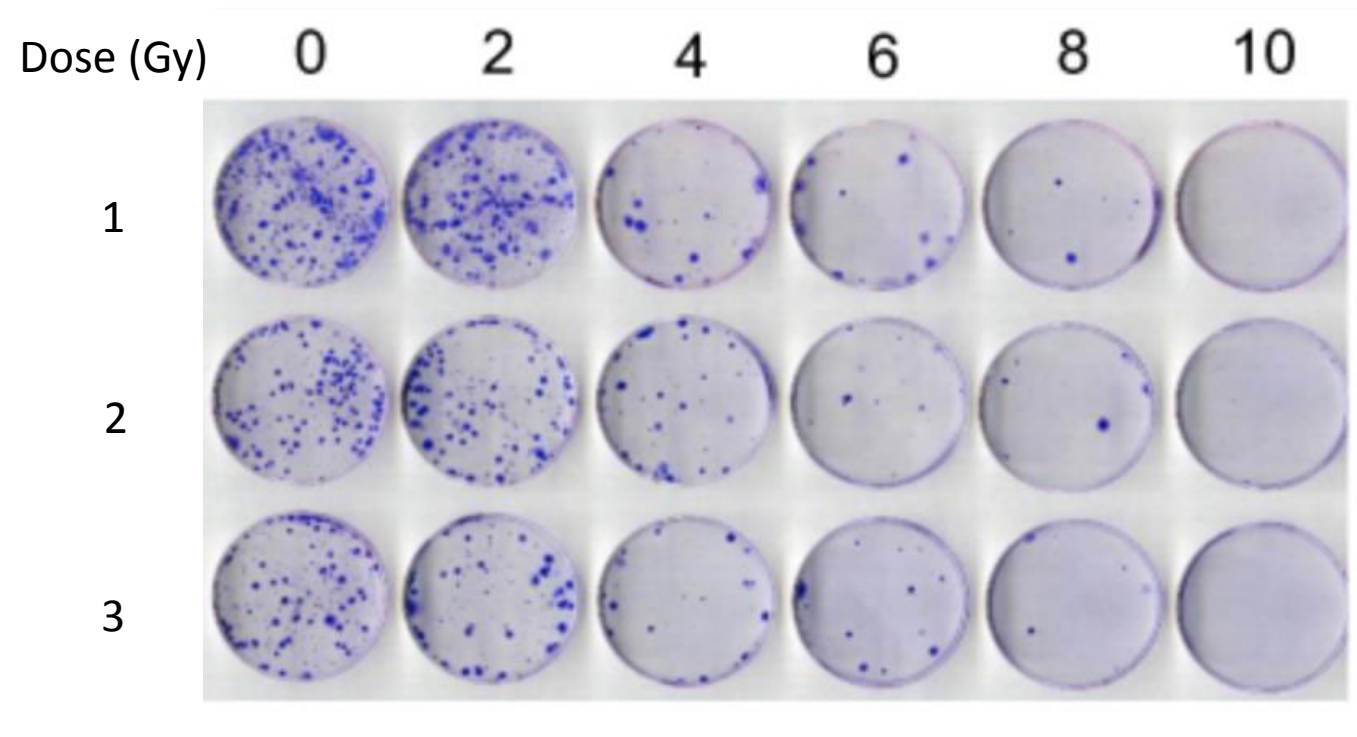
Tests de clonogénicité



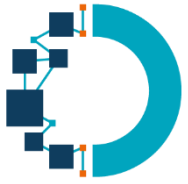


Matériel et méthodes

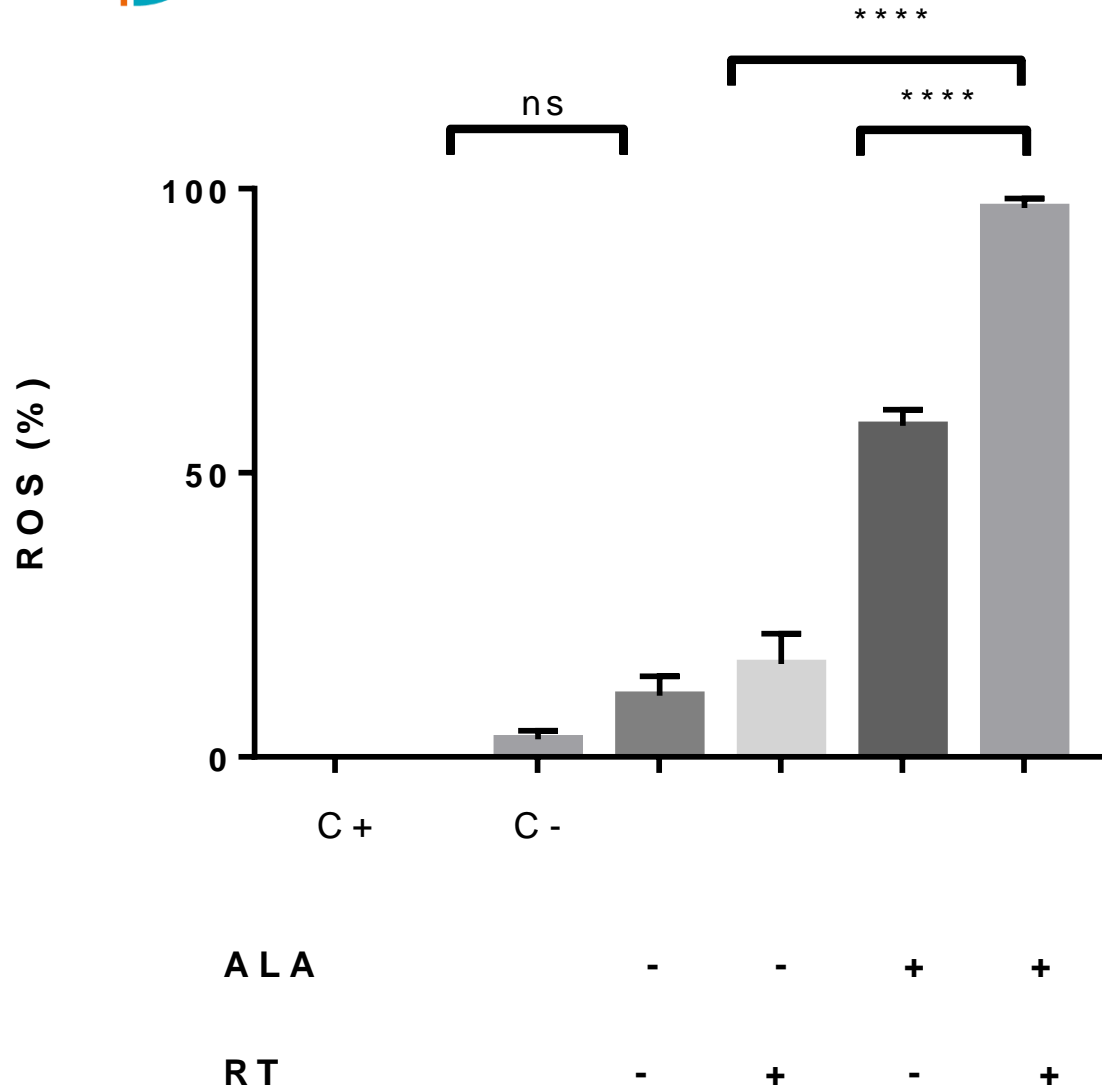
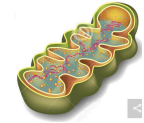
Tests de clonogénicité

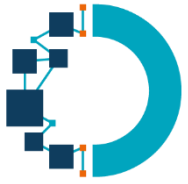


Effet radiosensibilisant de l'ALA

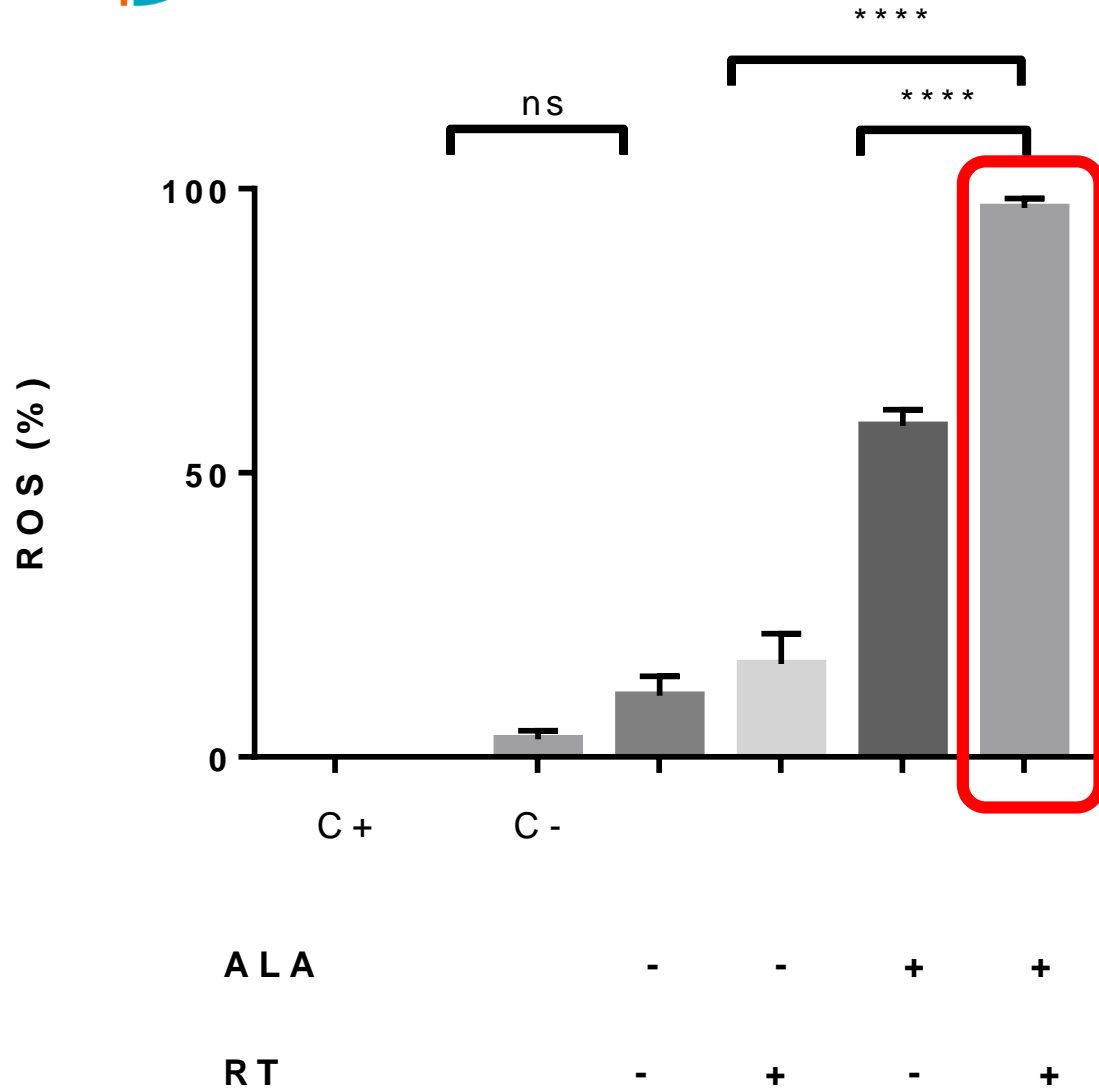
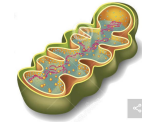


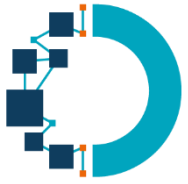
Résultats: mécanisme de radiosensibilisation



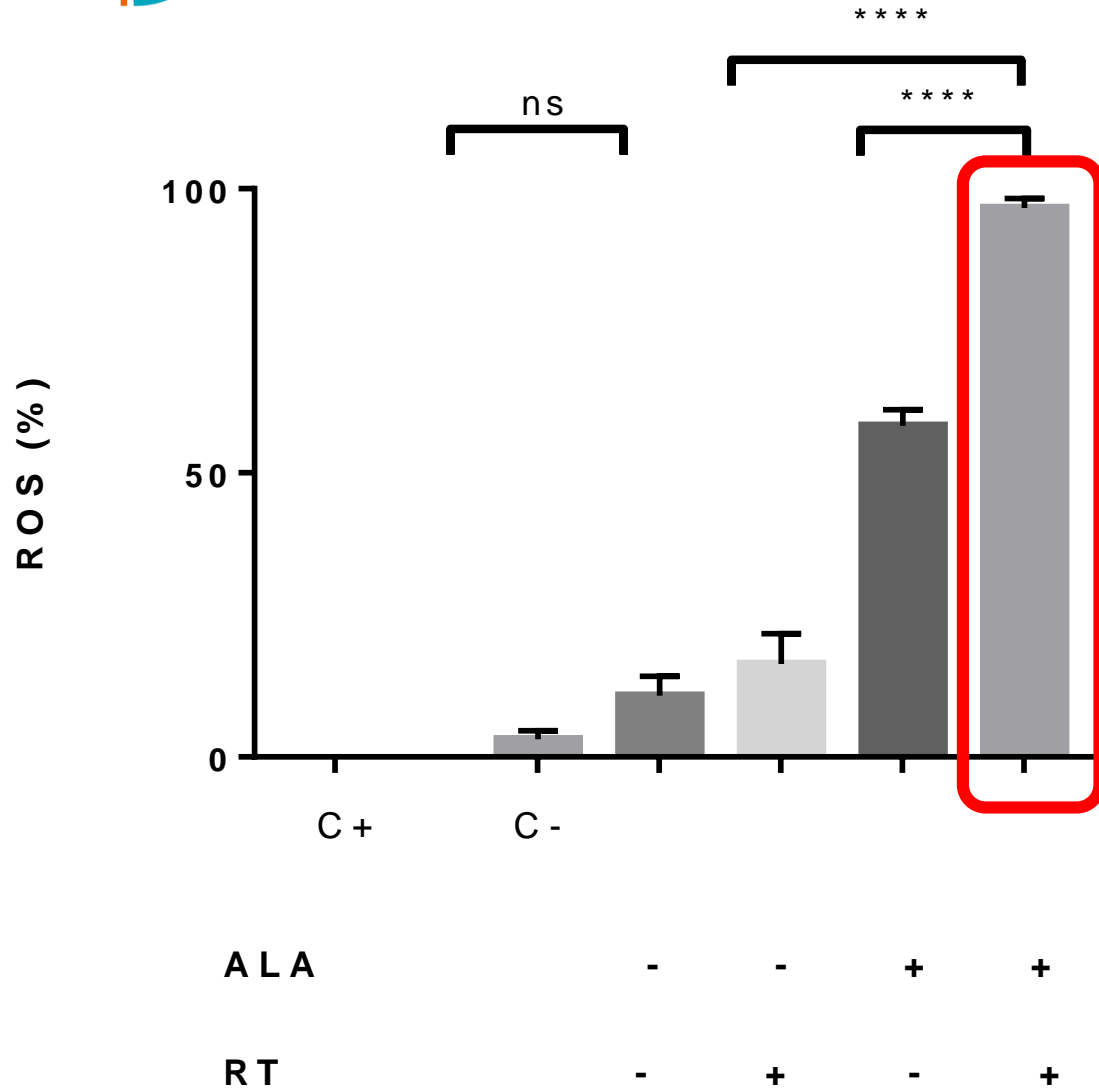
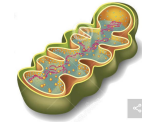


Résultats: mécanisme de radiosensibilisation

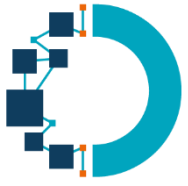




Résultats: mécanisme de radiosensibilisation



- ALA et RT induisent la formation de ROS
- Combinaison ALA + RT induit une forte synthèse de ROS



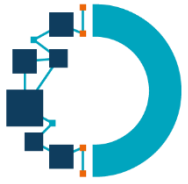
Conclusion

Radiosensibilisation de lignées gliales

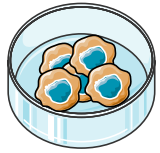
- ALA a un **effet radiosensibilisant**
- **Majoration du stress oxydatif**

=> Validation *in vitro*

=> Transposition en clinique?



Perspectives: transfert *in vivo*



IN VITRO

2019

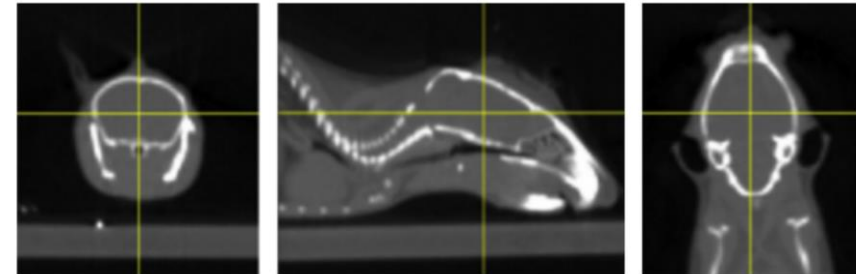


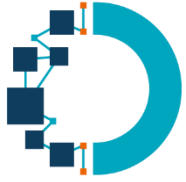
IN VIVO

2019-20



20..





Merci pour votre attention

