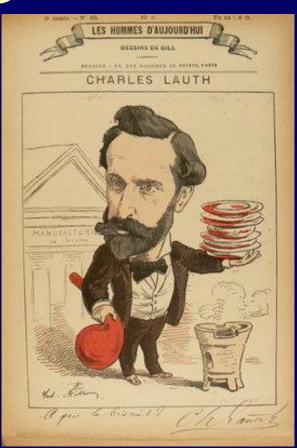
I am cofounder of ABBY, a company developing imipramine blue, a triphenylmethane, for the treatment of glioblastoma

"Where there is no interest, there is no conflict" Raymond Schinazi, PhD, codiscoverer of Emtriva

Charles Lauth, discoverer of gentian violet



Lauth, C. (1867), <u>"On the new aniline dye, 'Violet de</u> <u>Paris'"</u>, Laboratory, **1**: 138–139 Available OTC as 1% or 2% solutions, creams, ointments and pessaries (\$2-\$6)

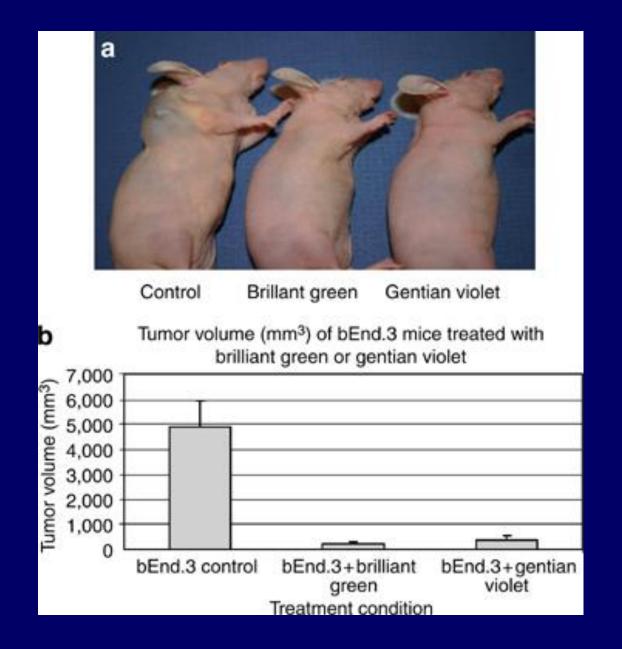
• Other names:

-andergon
-aniline violet
-basic violet 3
-Brilliant violet 58
-hexamethyl pararosaniline Cl
-meroxylan
-pyoktanin
-crystal violet

-etc.....







Perry BN et al, JID 2006

Infantile Hemangioma

- Most common tumor of infancy
- Usually absent at birth→ appears within 1st weeks of life→ undergoes rapid prolif→ spontaneous involution of years
- Result in a range of complications, from cosmetic to life threatening







Before and after Gentian Violet to ulcerated hemangioma (1 month of treatment, Leslie Lawley, M.D.)



Gentian violet downregulates Angptl4

Proof of concept using Triphenylmethanes in Humans (Gentian violet Plus imiquimod)

Hypothesis: Combining Imiquimod (TLR7 agonist, Interferon inducer) with Gentian violet (NADPH Oxidase inhibitor) will result in superior immune response than imiquimod alone

Three bulky melanoma metastases after debulking

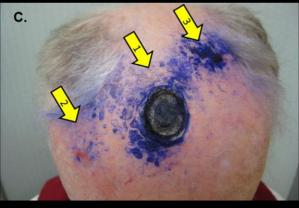
<u>Combination therapy of imiquimod and</u> <u>gentian violet for cutaneous melanoma metastases.</u> **Arbiser** JL, Bips M, Seidler A, Bonner MY, Kovach C. J Am Acad Dermatol. 2012 Aug;67(2):e81-3. doi

> Lack of recurrence at 6 months following gentian violet and imiquimod

Three bulky melanoma metastases at presentation







Regression of diffuse B-cell lymphoma of the leg with intralesional ger violet.

Rao S, Morris R, Rice ZP, Arbiser JL.

Exp Dermatol. 2017 Aug 20. doi: 10.1111/exd.13418

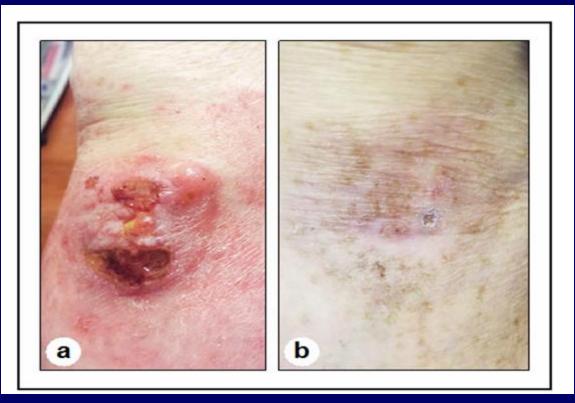
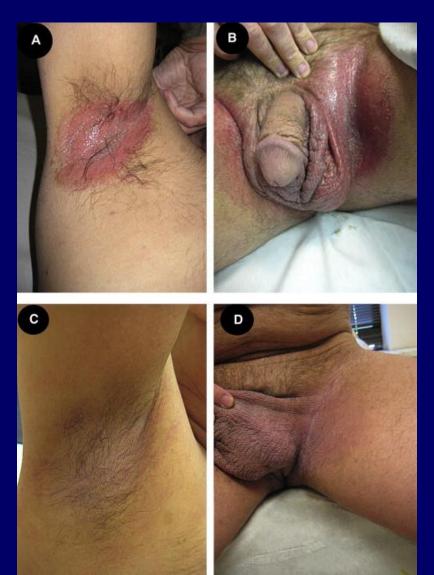


Figure 1: Primary Cutaneous Diffuse B-cell lymphoma, leg type: Clinical appearance of the patient at presentation a), and lesion at four months follow up b)

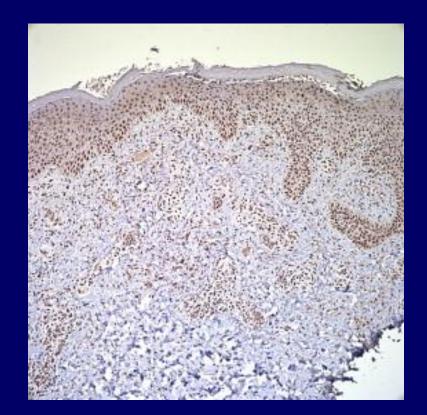


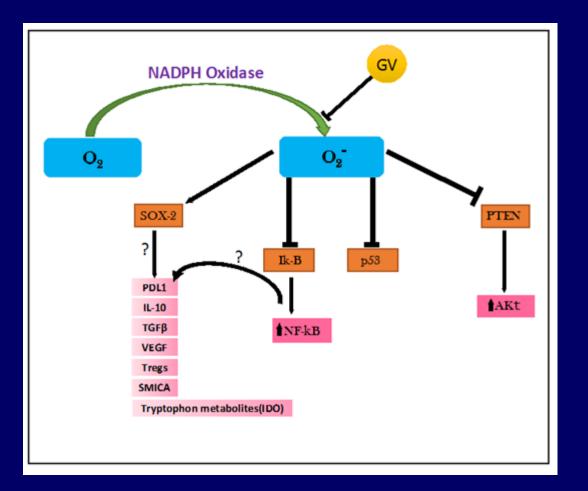
JAAD Case Report 2019 May; 5(5): 413–415. **Treatment of recalcitrant mycosis fungoides with topical gentian violet**

Anti-inflammatory effects of gentian violet in humans



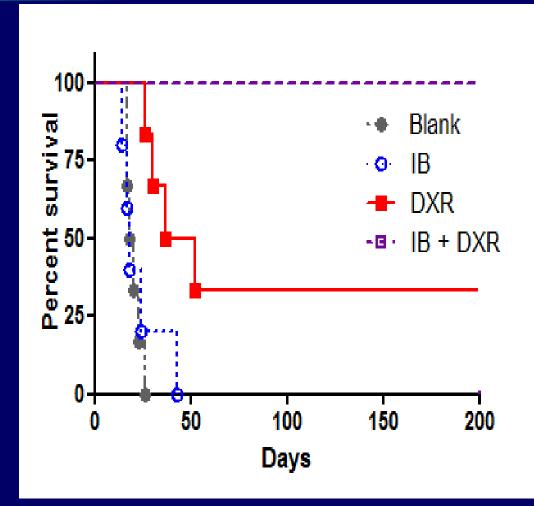
Journal of the American Academy of Dermatology Volume 63, Issue 3, September 2010, Pages 537-539





Generation of reactive oxygen species results in activation of NF κ B, Akt and suppression of wild type p53. In addition, SOX-2 may increase the production of immunosuppression factors.

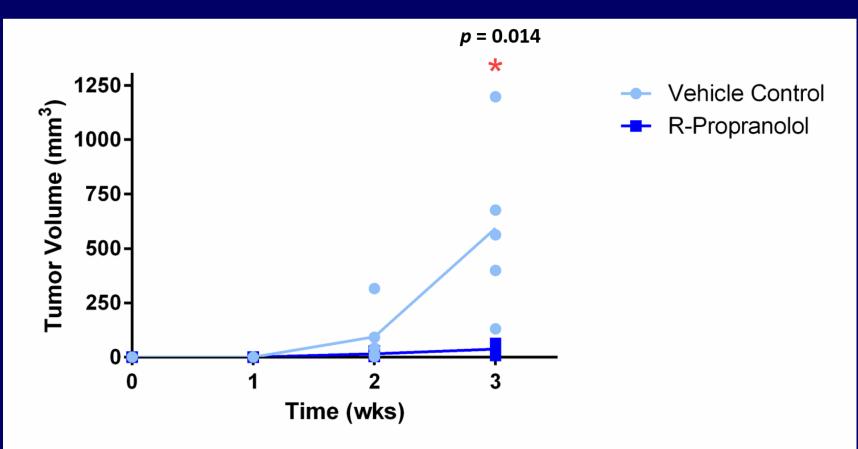
In Vitro Studies (Glioblastoma in Rats) - Delivery of NANO-IB Before DXR Yields Increased Survival



Anti-Invasive Adjuvant Therapy with Imipramine Blue Enhances Chemotherapeutic Efficacy Against Glioma Science Translational Medicine 28 Mar 2012:

ABBYMED

R-Propranolol has antitumor activity in vivo



NPJ Precis Oncol. 2019 Nov 1;3:27. doi: 10.1038/s41698-019-0099-9. eCollection 2019

<u>Summary</u>

- GV has been used for many years
- Antimicrobial/antimycotic actions are well documented
- Discovery of GV as NADPH oxidase inhibitor leads to use as anti-inflammatory and anti-tumor agent
- Novel uses for the future.....novel delivery modes Precursor to systemic NADPH oxidase inhibitors for cancer and inflammation