

SEARCH FOR POTENTIAL ANTITUMORAL, ANTIOXIDANT, ANTIINFLAMMATORY, ANTIDIABETIC, ACETYLCHOLINESTERASE AND MIELOPEROXIDASE INHIBITORY NATURAL COMPOUNDS FROM CERRADO AND ATLANTIC FOREST

Dulce Helena Siqueira SILVA

Araraquara Chemistry Institute / Paulista State University (Unesp)



Plants and endophytic fungi-derived extracts and pure compounds

The changing strategies for preservation and sustain the diversity in Brazil in the past few years evidences the intrinsic value of this enormous biological resource of potentially new bioactive compounds, and represents one of the greatest challenges nowadays facing the accelerated process of devastation of several Brazilian biomes. The systematic bioactivity evaluation of crude extracts and pure compounds from São Paulo state biota associated with additional information on chromatographic profile, spectrometric data and/or biological activity shall result in value-added material, which will be available for further studies. The might converge to hits or lead compounds to pharmaceutical, agroceutical, nutraceutical, cosmetics industry, which is expected to bring great contribution for conservation and sustainable economic development from the biodiversity of São Paulo state.

SUMMARY OF RESULTS TO DATE AND PERSPECTIVES

As part of our initial objectives, partial insertion of data from collections of plant material and preparation of plant extracts has been performed, using the BIOprospecTA databank (https://bioprospecta.iq.unesp.br/scyllaProspecta/).

Our bank of plant extracts, currently at Botanic Institute – IBt-SMA, Sao Paulo, has ca. 1,800 extracts from Sao Paulo State main biomes: Cerrado and Atlantic Rainforest. Expansion of this bank has not been significant over the past three years due to CGEN (MMA – Ministry of Environment) restrictions regarding collection and access to biological material. In addition, a collection of endophytic fungi extracts has been organized at NuBBE (Chemistry Institute-UNESP-Araraquara) which has been proven a rich and attractive source of bioactive natural products.

Samples from the bank of extracts have been assayed for antifungal, antioxidant, antimalarial and antitumoral activity, in addition to inhibitors of inflammation-related enzymes myeloperoxidase and cyclooxygenase, and acetylcholinesterase, which is involved in CNS related diseases. This preliminary screening indicated ca. 6% of tested samples presented bioactivity.

Modern phytochemical methodologies, including hyphenated chromatographic/spectroscopic techniques; and for bioactivity evaluation, using fast, sensitive and reproducible preliminary bioassays have been combined for conducting systematic studies, which resulted in the isolation of several bioactive compounds from selected plant species. Such screening allowed the selection of promising crude extracts, which have been chemically investigated in depth and afforded ca. 250 pure compounds. The obtained natural metabolites had their biological/pharmacological properties evaluated, which resulted in antitumor clerodane diterpenes from Casearia sylvestris, cytotoxic piperidine alkaloids from Cassia leptophyla, antioxidant glucosylxanthones from Arrabidea samydoides, antifungal alkyl and benzyl sulphides from Petiveria alliaceae, antifungal, antibiotic and cytotoxic nor-lignans from Styrax ferrugineus and S. camporum (Styracaceae), antibiotic phenetyl fatty acid esthers from *Stemodia foliosa*, cytotosic guanidine alkaloids from Pterogyne nitens (Fabaceae) and Alchornea glandulosa (Euphorbiaceae), antioxidant flavonoids from Nectandra grandiflora and Chiococca braquiata, among others.

Such results have attracted the attention of pharma/cosmetic companies, to carry out co-funded research (FAPESP, FINEP) in our labs aiming the development of products containing value-added material from the Brazilian plant biodiversity.

MAIN PUBLICATIONS

Fernandes DC, Regasini LO, Vellosa JCR, Bolzani VS, et al. 2008. Myeloperoxidase inhibitory and radical scavenging activities of flavones from *Pterogyne nitens*. Chemical and Pharmaceutical Bulletin. **56**.

Regasini LO, Vellosa JCR, Silva DHS, Furlan M, et al. 2008. Flavonols from *Pterogyne nitens* and their evaluation as myeloperoxidase inhibitors. *Phytochemistry*.

Silva DHS, Zhang Y, Santos LA, Bolzani VS, Nair M. 2007. Lipoperoxidation and cyclooxygenases 1 and 2 inhibitory compounds from *Iryanthera juruensis*. *J. of Agricultural and Food Chemistry*. **55**:2569-2574.

Carbonezi CA, Hamerski L, Gunatilaka L, Cavalheiro AJ, Castrogamboa I, Silva DHS, et al. 2007. Bioactive flavone dimers from *Ouratea multiflora* (Ochnaceae). *Brazilian Journal of Pharmacognosy*. **17**:319-324.

Scorzoni L, Benaducci T, Almeida AMF, Silva DHS, Bolzani VS, Gianinni MJ. 2007. The use of standard methodology for determination of antifungal activity of natural products against medical yeasts *Candida sp* and *Cryptococcus sp. Brazilian Journal of Microbiology*. **38**:391-397.

Viegas Jr. C, Silva DHS, Pivatto M, Rezende A, Castrogamboa I, Bolzani VS, Nair M. 2007. Lipoperoxidation and Cyclooxygenase Enzyme Inhibitory Piperidine Alkaloids from *Senna spectabilis* Green Fruits. *J. of Natural Products*. **70(12)**:2026–2028.

Scorzoni L, Benaducci T, Almeida AMF, Silva DHS, Bolzani VS, Mendes-Gianinni MJS. 2007. Comparative study of disk diffusion and microdilution methods for evaluation of antifungal activity of natural compounds against medical yeasts *Candida spp* and *Cryptococcus sp. Revista de Ciências Farmacêuticas Básica e Aplicada*, **28**:25-34.

Teles HL, Hemerly JP, Pauletti PM, Pandolfi JRC, et al. 2005. Cytotoxic lignans from stems of *Styrax camporum* (Styracaceae). *Nat. Prod. Lett.*, Inglaterra. **19**:319-323.

Carvalho PRF, Silva DHS, Bolzani VS, Furlan M. 2005. Antioxidant Quinonemethide Triterpenes from *Salacia campestris*. *Chemistry* & *Biodiversity*, Zurique. **2(3)**:367-372.

Ribeiro AB, Yoshida M, Bolzani VS, Santos LS, Eberlin MN, Silva DHS. 2005. New Neolignan and Antioxidant Phenols from *Nectandra grandiflora*. *Journal of the Brazilian Chemical Society*. **16(3)**:526-530.

Dulce Helena Siqueira SILVA

Instituto de Química de Araraquara Universidade Estadual Paulista (Unesp) Rua Professor Francisco Degni s/n Caixa Postal 355 – Quitandinha CEP 14800-900 – Araraquara, SP – Brasil .55+16-3301-6659 dhsilva@iq.unesp.br