



JUNE 2020

CONTENTS

"Annie Montague Alexander: A Multifaceted Life"-Part 2 - 1 "I Knew That" - 4 Meeting Topics - 5 Appreciation - 5 Adobe House Report - 5

President's Message - 6

EVENTS

JUNE

18 – Chapter Meeting, 6 pm Program, 7 pm



ZOOM Meeting 7 pm (See page 4)

SEPTEMBER

17 – Chapter Meeting, 6pm Program, 7pm

Annie Montague Alexander (1867-1950): A Multifaceted Life Part 2: The Partnership of Alexander and Kellogg

by Nancy Nies

IN THE EARLY YEARS OF THE TWENTIETH CENTURY, IT WAS considered inappropriate for a single woman to accompany a group of men on a scientific expedition. So, in 1908, when Annie Alexander planned

to finance and take part in a bear-hunting trip to Alaska — for the **UC Berkeley Museum of Vertebrate Zoology** (**MVZ**), which she had founded that same year — she sought a female companion. When she met **Louise Kellogg** (1879-1967), a teacher thirteen years her junior, she found a kindred spirit who would not only join her on the Alaska trip, but also become her partner — in work and in life — over the next 42 years, until Annie's death.

Fauna, Farming, Fossils . . .

That two-month expedition, to Alaska's Prince William Sound area, gave Louise her first experience with field work, and she proved an energetic, enthusiastic participant. The party brought back an eight-foot male grizzly bear and a smaller female, two hundred small mammals and a great number of birds for the museum's collection. After that Alexander and Kellogg would work on their own, collect-

d he

Annie Alexander (left) and Louise Kellogg (right). (Illustration by **Katrin Em**ery for palaeopoems.com)

ing birds and small mammals on many field trips throughout western North America and abroad. Though they were to make regular trips to Hawaii, which Annie would always consider her *"only real home,"* the

...Annie and Louise chose to collect in remote areas where the plants were less well known ... women would live in California for the rest of their lives.

In 1911 Annie bought 525 acres on **Grizzly Island** in **Suisun Bay**. Reclaiming Sacramento River delta land, the women turned the undeveloped acreage into a farm. There, they made their home and

raised cattle, pigs, barley, oats, hay — and, notably, their own AK brand of asparagus. At the time, the area was remote and wild. Annie sent speci-

The California Native Plant Society is a non-profit organization dedicated to the conservation of California native plants and their natural habitats, and to increasing the understanding, appreciation, and horticultural use of native plants.



Photo by Ashley Olive

Prunus andersonii (desert peach) close-up, 21 April 2018

mens trapped there to "her" museum, which led to the identification of six new subspecies of birds and mammals. But Alexander had not forgotten her first love, paleontology.

In 1921 she founded the **University of California Museum of Paleontology (UCMP)** on the Berkeley campus. She and Kellogg then embarked on what Stein calls "*a spate of paleontological expeditions.*" Among them was a trip to **Last Chance Gulch** in **Kern County**, where they found fossils dating from the Lower Pliocene. Their most significant fossil work was in 1931, when they found both carnivore and ungulate fossils in Miocene formations near **Barstow**. Annie wrote that the experience involved "*half a month with pick and shovel, digging a camel out of the ground*. . .", as well as painstaking work on their knees with jackknife and chisel.

Prunus andersonii (desert peach), *Leptosyne bigelovii* (coreopsis), *Yucca brevifolia* (Joshua tree) at Walker Pass, 21 April, 2018, just as described by Alexander and Kellogg in April 1940.

And . . . Flora!

Although Alexander and Kellogg had done some previous plant-collecting, it was not until 1939, after decades of labor-intensive field work in zoology and paleontology, that the women turned their full attention to botany. Collecting plant specimens required much less labor and equipment, and allowed them to see familiar places in a new way. Also, Annie and Louise chose to collect in remote areas where the plants were less well known, and were particularly drawn to **California's Mojave Desert** and **Eastern Sierra**.

The women's botanical field notes from 1928 to 1968, mostly from trips in California and Nevada, are held at the University and Jepson Herbaria. See <u>http://ucjepsarchives.berkeley.edu/archon/?p=collections/</u> <u>findingaid&id=99&q=</u>. Botanist **Tom Schweich** has made Alexander and Kellogg's Eastern Mojave field notes (1939-1949) available online. On April 12-13, 1940, Annie and Louise recorded sighting thirty plants in the **Walker Pass** and **Inyokern** areas of **Kern County.**

Their field notes for April 12 read: "From Bakersfield 50 miles to Isabella, following up Kern R. through canyon. Isabella in long valley. Onyx at upper end. Start collecting at 3750 ft. alt. Conspicuous bush in full bloom – wild peach – from this point on, nearly to summit of pass. . . **Prunus andersonii**. . . **Coreopsis bigelovii**. . . Summit "Walker Pass - 5250 ft. A clump of Joshua trees on east slope just below summit. Bare granite sand hillsides. A yellow bloom far up slope facing south." I feel certain that Annie and Louise would be pleased to see the same plants blooming in the same location — in a photo taken at Walker Pass on an April day nearly eighty years after their visit.

The Herbaria's website credits the women with collecting more than 300 plants, representing nearly 200

> species, in what is today **Mojave National Preserve.** Their most important botanical find, in 1949, was *Swallenia alexandrae* (**Eureka dunegrass**), a rare genus in the grass family with only one known species, found only in the Eureka Valley sand dunes in **Death Valley National Park**.

The women's beautifullyprepared botanical specimens, often of unusual or undescribed species, included maps, photos and meticulous field notes. **Alice Eastwood**, curator of botany at the **Cali**-



fornia Academy of Sciences, wrote to Alexander in 1942, "*The specimens are the finest ever and we are very grateful to you and Miss Kellogg.*"

A Lasting Legacy

By the 1940s, Alexander and Kellogg were renowned at Berkeley — for both the quantity and quality of their natural-history collections. In all, they donated over 20,000 specimens to the MVZ, and around 1,500 (plus many uncatalogued ones) to the UCMP. In addition, Annie and Louise contributed almost 18,000 plant specimens to the University Herbarium. There are seventeen taxa — fossils, birds, mammals and plants — as well as a lake in Alaska, named in honor of Alexander, and several other taxa named in honor of Kellogg.

The MVZ website states that '[w]hile Alexander was not herself a scientist, it was her vision, her money, the specimens that she collected, and her financial and political acumen that created the natural history museums on the Berkeley campus and that permitted men like Joseph Grinnell and John C. Merriam to achieve distinction in their disciplines."

The work of Alexander and Kellogg was and is valuable for understanding speciation and evolution, as well as for preserving endangered species. Stein writes that Alexander made so many collections in order to "create a record of the beauty and natural diversity from which she derived so much pleasure, but which she recognized was rapidly being lost in the wake of development and human population growth."

Along with their numerous contributions to science, Alexander and Kellogg played a role in the advancement of women — in both scientific study and participation in field research. Until the 1940s, women were rare in the graduate programs in zoology and paleontology, and were not allowed to go on MVZ-sponsored field trips. In the 1930s Alexander and Kellogg invited the first two women graduate students to accompany them on expeditions, and in the early 1940s sponsored field trips for female graduate students. In 1948 Alexander established two scholarship funds — one in paleontology and the other in vertebrate zoology — for graduate students of both sexes, whom she considered her "investments." These funds were, of course, in addition to Alexander's previous generous donations and endowments to both museums. Kellogg, in her will, was to leave an endowment to the University Herbarium.

Annie and Louise's last adventure together — a threemonth trip in 1947 to **Baja California**, where Annie celebrated her 80th birthday — provides another example of their encouragement of women. They

Swallenia alexandrae (Eureka dunegrass) Death Valley National Park, — 26 April 2011.

invited forty-year-old **Annetta Carter**, who worked at the herbarium but had never been on a field trip because of her sex, to join them on the expedition, from which the three women returned with more than 4,600 sheets for the herbarium.

The experience was life-changing for Carter, who would return frequently to Baja California and become an authority on the region's flora. On subsequent trips there, Carter would collect new species and name one in memory of each of her mentors: *Bouvardia alexanderae* and *Acacia kelloggiana*. The legacy of Annie Alexander and Louise Kellogg, says Stein, endures "*in the lives of women who seek to follow their own star to the fullest degree possible.*" \clubsuit

Author's note: Many thanks to **Clyde Golden** for introducing me to Annie Alexander and Louise Kellogg, and to **Barbara K. Stein** for her outstanding book about their lives and accomplishments.



CNPS is the leader for providing reliable information on California native plants and plant conservation. Comprehensive information about California's flora and vegetation communities is available throughout the state for conservation and educational purposes. CNPS's leadership influences personal ethics and actions, as well as public policy for native plant protection.



"I KNEW THAT"

by Yvonne Turkal

T HIS COLUMN IS ABOUT EDIBLE PLANTS, with a disclaimer. If you decide to investigate further, the information presented here comes from <u>Edible and Useful Plants of California</u>, Charlotte Bringle Clarke, Univ. of California Press, 1977 and from the website <u>calscape.org</u>. The Kern California Native Plant Society isn't responsible for any of your results.

CALIFORNIA WILDROSE (Rosa californica)



"There are nine species in California,.... All have edible, though not always palatable, fruits and flowers, R. woodsii with straight prickles" (Why not call them thorns?) "has an unusually pleasant fragrance."¹ (What does that mean?)

The californica spe-

cies is beautiful and grows through the coast and foothills of California, up to elevations of 6000 feet. It blooms May through August enjoying a drought, but they will thrive being planted by a water source.²

Flowers are generally flat with a pink color from almost white to a deep magenta, producing rose hips containing yellow seeds.

Clarke reports that it is often used as a barrier in a garden where it can form a large thicket. If you want to grow one in your yard, it can also grow from a shrub to 8-10 feet tall.

Historical Uses?

"The fruit of the Wild Rose, known as the hip, is said to contain more vitamin C, calcium, phosphorus, and iron than oranges. During World War II in England, the hips were gathered for their abundance of these essential vitamins and minerals."¹

Mimulus Memo — June 2020

When the dry inner seeds are removed, rose hips can make jelly, tea and/ or sauce. Backpackers like the plant because the fruits remain on the bushes from year to year and can be soaked in water, used, and eaten, resembling a small dry apple in appearance and taste.

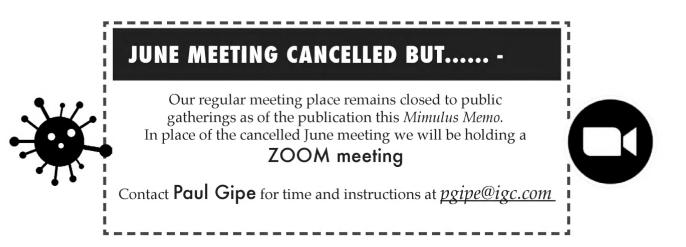


"The Cahuillas picked the buds and ate them raw or soaked the blossoms in water to make a drink. Some tribes made a tea from the roots for colds and from the leaves and fruits for pains and colic." (Doctors should hand a new mother one of these plants — just in case.) "The wood was used for arrow shafts, and the bark fiber made twine and other goods."¹ These rose thickets also provided great shelter and nesting sites for the Audubon Cottontail and the wood rat, among other western animals.

"I knew that" 🕁

References

- 1 Clarke, Charlotte Bringle 1977. *Edible and Useful Plants of California*. University of California Press, Berkley, CA.
- 2 California Native Plant Society CALSCAPE calscape.org <u>https://www.calscape.org/Rosa-californica-(California-Wildrose)?srchcr=sc5ed2ec98d455b</u>



Chapter Meetings

upcoming **TOPICS**

- Thursday, June 18, 2020 7 pm The planned program is cancelled. In its place will be a ZOOM meeting (*See page 4 for contact info*)
- Thursday, September 17, 2020 Presenter: Gary Adest River Ridge Institute (Springville) Topic: Ranching without Cows— Sustainable and Regenerative Land Management
- Suggestions for presenters and topics? Contact Paul Gipe <u>pgipe@igc.org</u> or Richard Spjut <u>richspjut@gmail.com</u>
- All chapter meetings are held the **3rd Thursday** of each month usually at 1300 17th Street, Room 1A or 1B, Bakersfield, CA. Check website for any change of venue.

Meeting times:

6 pm — Discussion groups on plant identification and native plant gardening

7 pm — Program presentation



- ... John Kalk, Rich Spjut and Ken Owen all of whom had their program presentations cancelled due to COVID-19 precautions.
- ... **Yvonne Turkal** for authoring a new *Mimulus Memo* column on edible plants.
- ... Don Turkal for keeping us up-to-date on activities around the Adobe House 🏠



ADOBE HOUSE

by Don Turkal

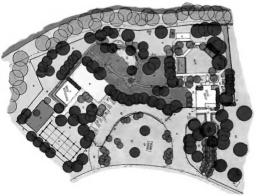
THE ADOBE HOUSE LANDSCAPE PLANS HAVE been completed. **Bill Cooper**, co-founder of the **Kern River Parkway Foundation** is working toward securing a land lease for additional landscaping.

Diane Mitchell and I suggested a demo garden to be used as an educational tool to highlight native plants different from those at CALM; and possibly to re-vegetate along the riparian area. Diane pointed out the riparian area may be problematic due to erosion, steep slopes, and lack of an irrigation system.

Diane attended a meeting in March with Kern County staff, **Alvin Nuval** representing **Chattel Architecture**, and representatives of the Kern River Parkway Foundation. It was recommended that the **Hart Park Working Group** be formalized as a subcommittee of the Kern River Parkway Foundation.

As a qualified historic architectural firm, Chattel submitted paperwork for the house to be placed on the **Register of Historic Buildings** and is preparing plans for its restoration.

Volunteers will assure that this needed educational tool is a success for Bakersfield and Kern County. Diane Mitchell and Don Turkal are representing Kern CNPS. Stay tuned for updates. ☆

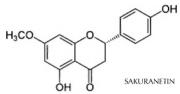


6



Eriodictyon californicum (California yerba santa), shrubs (left) near McCloud River, Shasta Lake May 12, 2009. Flowers close-up (right) Old Kern Canyon Rd. – 27 Apr. 2013. Leaves have been used by native American tribes for treating colds, coughs, fevers, rheumatism, asthma, pulmonary, dermatological and stomach problems.¹ Species one of six among 97 samples (WBA-656, SPJ-10440) I recommended on a shipping

list for antiviral and antitumor screening in addition to the chemopreventive screening at the Ohio State University School of Pharmacy. Twelve flavonoids found to inhibit metabolism of the carcinogen benzo[a]pyrene by hamster embryo cells in tissue culture.² The flavanone, sakuranetin, inhibited the response to saccharin by more than 50%.³ Additionally, sakuranetin has shown antiproliferative activity against human H₂CO cancer cell lines typical for B16BL6 melanoma, esophageal squamous cell carcinoma (ESCC) and colon cancer (Colo 320), and reported to have "antiviral activity towards human rhinovirus 3 and influenza B virus."4



President's Message: Antiviral Plants: Chemical in Kern County Plant Could Treat COVID-19? by Rich Spjut

Y CAREER HAS BEEN LARGELY DEVOTED **1** to discovery of new chemotherapeutic agents from plants including bryophytes, and also from lichens, cyanobacteria and marine algae. The collection strategy has been based on taxonomy, recognizing that the genus is the lowest level of chemo-taxonmic diversity,⁵ while reviewing plants used in traditional medicine.6 Most samples I collected were screened for antitumor, antibiotic⁷, antiviral, antiprotozoal⁸, and chemopreventive agents;9 fewer for isolation

Nearly half... of all chemo-therapeutic agents used in medical prescriptions or ... treatments can be attributed directly or indirectly to discoveries from natural products.

and identification of carotenoids¹⁰ and endophytes and endolichenic microbes¹¹ that produce the antitumor active compounds. An example of an endolichenic fungus is Penicillium aurantiacobrunneum, cultured from the lichen Niebla homalea, that led to discovery of four

new mycotoxins related to citreoviridin, one of which showed selective cytotoxicity towards ovarian and breast cell line assays.¹² Higher plants from which anticancer compounds have been isolated often harbor endophytic fungi and bacteria that produce the same or similar compounds.¹³ Moreover, antitumor activity in many moss species appears related to their microbial associations.^{14,15,16} Nearly half ("49%") of all chemotherapeutic agents used in medical prescriptions or prescribed treatments can be attributed directly or indirectly to discoveries from natural products; the active chemicals are "small molecules," in contrast to large molecules (nucleic acids, proteins, polysaccharides).¹⁷

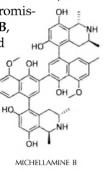
Recently, our chapter activities, like those of other CNPS chapters, have been curtailed by the spread of a coronavirus, SARS-CoV-2 — that causes COVID-19 classified in the genus Betacoronavirus, one of six genera in the family Coronaviridae. More than 30 families of viruses are recognized to infect humans and other vertebrates as outlined on the website "viral zone,"¹⁸ which classifies viruses by nomenclatural taxonomy, and by other criteria such as their host — including invertebrates, plants, fungi, eukaryotic microorganisms, bacteria and archaea. From my experience, I wondered to what extent antiviral compounds have been isolated from plants and lichens and advanced to clinical trials. A brief review follows.

One can easily find online reports of antiviral activity in plants and lichens; for example, 795 plant species reportedly used for treating HIV¹⁹ included 23 species — mostly nonnative — found in Kern County.²⁰ But it must be kept in mind that very few reach human clinical trials; for example, ~10% of 270 lichen species that I collected for the National Cancer Institute (NCI) from the United States and Mexico during 1986 – 1992 were "active" in HIV assays due to sensitivity to polysaccharides²¹ whose immunomodulating effects are already generally known in metabolism and food but nonetheless of pharmacological interest.^{22,23} On the other hand,

many of 344 medicinal plant compounds reported to show activity in HIV, herpes and other viral assays²⁴ have been clinically evaluated in cancer chemotherapy.

Antiviral plant compound - MICHELLAMINE B

During 1987–1996, the NCI screened more than 60,000 plant extracts in HIV assays.²⁵ Of four promising discoveries, one was michellamine B, a naphthyl-isoquinoline dimeric alkaloid isolated from leaves of a rare tropical rain forest woody vine (liana) that was later discovered to be a species new to science, Anscistrocladus korupen-HO sis (Ancistrocladaceae), found only in southwest Cameroon. The compound was active against many strains of HIV-1 and anti-HIV-2; however, in animal

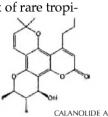


preclinical trials it had a very narrow therapeutic dosage between efficacy and toxicity.²⁶

Antiviral plant compound - CALANOLIDE A

Another NCI anti-HIV discovery, (+) calanolide A, a pyranocoumarin isolated from the latex of rare tropi-

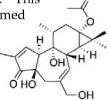
cal forest tree, Calophyllum lanigerum (Calophyllaceae), in Sarawak, Malaysia, advanced to HIV Phase I clinical trials with promising results, but development "delayed due to its low therapeutic index (range: 16-279), non-ideal antiviral activity, and the



complexity of its extraction from plants;" however, a calanolide derivative is being investigated in animal studies for the treatment of HIV in combination therapy.²⁷

Antiviral plant compound - prostratin

A third anti-HIV compound of interest is prostratin — an unusual phorbol ester (tetracyclic diterpene) in not being tumorogenic as opposed to many phorbol esters in Euphorbiaceae and Thymelaeaceae that cause tumors. Prostratin was isolated from the stembark of Homalanthus nutans (Euphorbiaceae), a rain forest tree in central Polynesia, Samoa.²⁸ This species — originally described and named Croton nutans by J.G.A. Forester in н 1786 from a specimen he and his father collected on Tonga — has ŌΗ since been recognized to occur in Fiji, Society Islands, New Caledonia, and OH New Hebrides,²⁹ distinguished from the PROSTRATIN



closely related and more widely distrib-

uted *H. populneus*.³⁰ In Fiji the local inhabitants prepare an aqueous infusion of the leaves and drink it for stomach troubles, and take a fruit decoction for relief of urination problems (ibid.). Similar uses have been noted in Samoa. An expert on the vegetation, flora and



Above: Aerial view of smokebush species, Conospermum stoechadis ssp. stoechadis - Aug 1992. Below: ground view, northern sand plains between Perth and Geraldton, Western Australia. This species appears to be the most conspicuous and common of the 20 smokebush species in section Foliosa, but not known to contain conocurvone. A survey was conducted by Richard Spjut and Ross Smith of World Botanical Associates (WBA) under a NIH-NCI Master Agreement Award to identify where best to collect Conospermum species for anti-HIV screening of concurvone.

ethnobotany of Samoa, W. Arthur Whistler, who was born near Death Valley, recently died from coronavirus, apparently infected while visiting in the state of Washington. Prostratin is in clinical trials in which its efficacy has been compared with that of a polyketide, bryostatin, discovered in a bryozoan, Bugula neritina,³¹ the compound also in clinical trials for treating cancer. 32

Antiviral plant compound - conocurvone

A fourth NCI anti-HIV compound, conocurvone, is a naphthoquinone, isolated from the root of a smokebush species, Conospermum incurvum (Proteaceae),

that I collected on the northern sand plains of Western Australia (WA) during Aug-Sep 1981.33 The WA field work came to a sudden halt when I was informed — that after 20

H₃¢ CONOCURVONE CHa

years - the NCI terminated their coop-

erative agreement with the USDA/ARS at the start of the new fiscal year. At that moment it was uncertain whether any of the 758 samples would be screened for their intended purpose — as new anticancer drugs especially since no money appeared available to ship the samples to the US. The curator of the WA herbari-



Smokebush AIDS drug pact reached

BY BRENDAN NICHOLSON CLINICAL trials of an AIDS drug derived from a chemical found in WA's native smokebush might be carried out experimenting with new approaches to the smokebush chemical, conocurrone. Even if the research was unsuccessful, the effort could produce invaluable information about the AIDS virus.

Top: *Conospermum incurvum*, near Perth, Western Australia, photo documented by herbarium specimen, SPJ 12464. Root contains conocurvone. **Bottom:** News article from a WA newspaper regarding agreement between the NCI and Australian Medical Research and Development (AMRAD) to pursue research on conocurvone HIV activity.

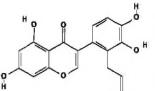
um expressed his disappointment, while also conveying that they could not use or store the samples. It was further disappointing in that I had had to amend my license to allow for the collection of root samples.

However, I successfully lobbied to get the samples shipped to my office facility in Maryland at the Beltsville Agricultural Research Center. After accounting for their identification and dry weight, they were shipped to the extraction laboratory in Wisconsin from where many went to Purdue University and to Research Triangle Institute (RTI) in North Carolina. Monroe Wall at RTI, who isolated the anticancer drug taxol from the Pacific yew, Taxus brevifolia, based on original activity in the KB assay,³⁴ reported KB activity in nine WA species.³⁵ The KB test results for 181 of 340 samples at Purdue University were sent to me in 1986 by Thomas McCloud, a graduate student of Dr. John Cassady; these results were published on the WBA website in 2014.³⁶ Although Conospermum was not among the samples screened against KB, the test results indicated the significance for collecting root; 15 of 24 (63%) of the active species in WA woody plants had activity only in root.

The WA samples, among many others at Purdue University, were retrieved and brought to the NCI Natural Products Repository in Frederick, MD by Dr. McCloud for anti-HIV screening during the time when the NCI was developing their antitumor 60-cell-line assays. Discovery of anti-HIV activity in the smokebush led me to visit WA in 1991 for recollections, encouraged by WA authorities. Additional material needed in 1992 led to a partnership, World Botanical Associates-Australia. Agreements were initiated with WA government authorities, including a NCI letter of Collection, and proposals were sought with Kings Park for cultivation of smokebush in tissue cultures. Several hundred species also were collected for preliminary screening of novel bio-active agents. But only a temporary agreement could be reached. After I left WA, conocurvone continued to be pharmacologically evaluated in Australia, while I have not seen reports on its advancement in HIV screening. Nevertheless, concurvone has emerged as "one of the most notable members" of the naphthoquinone class of compounds that — in its synthesis — has led to a "series of novel dimeric naphthoguinones" that show "selective cytotoxic to human acute myeloid leukemia (AML), breast and prostate cancer cell lines." ³⁷ A "relatively simple naphthoquinone, ß-lapachol" "obtained from the bark of the South American lapacho tree, (Handroanthus *impetiginosus*), has been in phase II clinical trials in the US for advanced solid tumors.³⁸

New leads gleaned from databases...

As field exploration of plants continue to find leads to new therapeutic drugs, chemists using computer algorithms search "ligand databases" ³⁹ on the genomic, structural, reactive and binding properties of known molecules; in essence, they virtually screen compounds for new drug therapies, ranking the results as one might find from a search in Google looking for specific information using key words. Recently, chemists at universities in China and Saudi Arabia - in collaboration – reported screening 32,297 "potential antiviral phytochemicals" for treating COVID-19 by targeting the enzyme that controls coronavirus replication as seen in the severe acute respiratory syndrome coronavirus (SARS-CoV), which has a very similar genome sequence to SARS-CoV-2 ("96.08%"). Their highest ranking compound was a 7-Hydroxy isoflavone – 5,7,3',4'-tetrahydroxy-2'-(3,3-dimethylallyl) — isolated from the root of Psorothamnus arborescens var. minu-



tifolius (Mojave indigo bush), the species and variety found here in Kern County.⁴⁰

The isoflavone was discovered by chemists at the Ohio State University–School of Pharmacy as a result of screening plants for activity against leishmaniasis, a disease caused by an infected sand fly bite, inoculating the victim with the protozoan (Leish*mania*).⁴¹ The isolation of the compound was guided by fractionation of the active extracts prepared from 670 g of dried root (WBA-4841-13) of P. arborescens var. minutifolius collected near the Kern County line (Nine-Mile Canyon) "in desert-chaparral transitional vegetation by Dr. Richard Spjut (WBA 4841-13, SPJ-15357), World Botanical Associates (Bakersfield, CA)." Dr. Salem in her Ph.D. dissertation (2005, Ohio State University) had reported the isoflavone and other flavonoids, and also those reported earlier from the aerial parts of P. polydenius. She concluded — in her abstract: "From the results [of screening 174 biodiversity samples collected and 149 extracts from selected plants in China used in Chinese medicine], the plant genus Psorothamnus was identified as a promising source of potential new antiparasitic compounds."

Of additional interest is that the percentage of antiparasitic actives was higher in my selection of samples (19/174, 11%) than in the selection from plants used in Chinese medicine (10/149, 6.7%), and that the sample of the invasive white horehound (*Marrubium vulgare*), I collected in Nevada (leaf-flower-fruit, 65 g) showed the "same activity" as the sample from China (native). As reported by Dr. Salem, my collection strategy focused "on samples not generally collected for the National Cancer Institute (NCI) based on the NCI requirement of 500 g" with emphasis on "root and stem bark from shrubs, small annual herbs, and flower and fruit parts of all plant species," samples each weighing "30–100 g."

The WBA samples Dr. Salem reported on were collected from San Diego County to southern Oregon and in western Nevada during June 14 to July 10, 1998. An example of a small annual is *Mimulus bigelovii*; a 45 g sample (WBA-3664) of the entire herb led to discovery of selective antileishmanial active compounds in the species. This was followed by two recollections, one in 2003 (WBA 4840-11, 130 g) and another in 2005 from limestone talus slopes west of Sandy Valley NV in the Mesquite Mts. in California along Kingston Rd near the state line; active C-geranyl flavonoids were isolated from 1.1 kg (dried) of the whole plant (WBA-5257-11, SPJ-15900).⁴²

Environmental variations

Finally, it should be recognized that plants contain many secondary metabolites in different parts of the plant in various concentrations, and the species itself may not always produce the active compounds in <u>all</u> environments. An example is **white corn lily**, *Veratrum californicum* (Melanthiaceae). I conducted surveys in the western US in 2004–2011. A semi-synthetic derivative of the active alkaloid, **cyclopamine**, depended on



Mojave Indigo Bush (*Psorothamnus arborescens* var. *minutifolius*). Highest-ranked plant worldwide for containing a chemical in the root — **isoflavone**, 5,7,3',4'-tetrahydroxy-2'-(3,3-dimethylallyl) — that might prove useful for treating COVID-19. Shrub (above) and close-up flower (below) taken in western Mojave Desert, Short Canyon, Kern County – March 2016 and April 2019. Related species, *P. polydenius*, used by the Paiute and Shoshoni for colds, measles, smallpox, pneumonia, whooping cough, and tuberculosis;¹ its active flavonoids were reported 2004 in the Journal of Natural Products (68: 108–111) by Drs. M. M. Salem and K. A Werbovetz.

the plant source for synthesizing the derivative; the alkaloid concentrated in the root-rhizome-bulb. Plants that produced the alkaloid were primarily found in a narrow geographic range of the species, and only in specific types of vegetation.⁴³ Even among clones, the yield in cyclopamine varied, suggesting the active compound further depended on mycorrhizal association.

References

- 1 Moerman DE. 1998. Native American Medicinal Plants. An Ethnobotanical Dictionary. Timber Press, Portland.
- 2 Liu YL DK Ho, JM Cassady, VM Cook, WM Baird. 1992. Isolation of potential cancer chemopreventive agents from *Eriodictyon californicum*. J. Nat. Prod. 55(3): 357–363. Notes: 0.4–1 kg collected of leaf with twig in Oregon just south of Cave Junction along Hwy 199, June 30, 1988 (WBA-656, SPJ-10436). WBA is abbreviation for World Botanical Associates, followed by accession number given to the plant part(s); SPJ is abbreviation originally assigned by the New Crops Research Branch, USDA Agricultural Research Service (ARS) for collections by Richard Spjut followed by his collection (voucher specimen) number.
- 3 Fletcher JN, AD Kinghorn, JP Slack, TS McCluskey, A Odley, Z Jia Z. 2011. J. Agric. Food Chem. 59 (24):13117–13121. Note: 1.875 kg of dried leaf (WBA-4400-33) and other plant plants—collected by R. Spjut near Whiskey Town Reservoir in northern California, Sep 2002.
- 4 Stompor M. 2020. A review on sources and pharmacological aspects of sakuranetin. Nutrients 12(2) 513.
- 5 Spjut RW. 1985. Limitations of a random screen: Search for new anticancer drugs in higher plants. Econ. Bot.: 39: 266–288.
- 6 Spjut RW. 2005. Relationships between plant folklore and antitumor activity: An historical review. Sida 21(4): 2205–2241.
- 7 Jean-Jacques ME, AL Okunade, AM Clark, "Antimicrobial and chemical investigation of *Vermilacinia leopardina*." 19th MALTO Medicinal Chemistry-Pharmacognosy Meeting, Little Rock, AR, May 17-19, 1992. Note: An unpublished report; the ms name V. *leopardina* published in 1996 by Spjut in Sida Misc.14, preceded by the genius name published in1995, *Flechten Follmann*: 37–51).
- 8 Salem MM. 2005. Identification of Antikinetoplastid Compounds from *Psorothamnus polydenius* and *P. arborescens*. Ph.D. Dissertation, 214 pp. Ohio State University, Committee: KA Werbovetz (advisor), A Douglas Kinghorn, C-s Chen. **Note:** 240 samples collected June-July1998 by R. Spjut, WBA in Laurel MD, shipped to Dr. John Cassady, Dean of the School of Pharmacy, OSU, Sept.2, 1999, cited by M Salem.
- 9 Cassady JM. 1990. Natural products as a source of potential cancer chemotherapeutic and chemopreventive agents. J. Nat. Prod. 53(1): 23–41.
- 10 Czeczuga B, BD Ryan, RW Spjut, J-A Flock, WA Weber, CW Beasley, RE Showman, RD Worthington, VL Boucher. 1997. Carotenoids in lichens from the United States of America and Mexico. Feddes Repertorium 108 (5-6): 401–417.
- Suryanarayanan T S, N Thirunavukkarasu. 2017. Endolichenic fungi: the lesser known fungal associates of lichens. Mycology 8(3): 189–196.
- 12 Tan CY, F Wang, GD Anaya-Eugenio, JC Gallucci, KD Goughenour, CA Rappleye, RW Spjut, EJ Carcache de Blanco, A Douglas Kinghorn, LH Rakotondraibe. 2019. Pyrone and sterol constituents of *Penicillium aurantiacobrunneum*, a fungal associate of the lichen *Niebla homalea*. J. Nat. Prod. 82 (9): 2529-2536.
- 13 Uzma F, CD Mohan, A Hashem, NM Konappa, S Rangappa, PV. Kamath, BP Singh, V Mudili, VK. Gupta, CN Siddaiah, S Chowdappa, AA. Alqarawi, EF Abd-Allah. 2018. Endophytic fungi—Alternative sources of cytotoxic compounds: A review. Front Pharmacol. 9: 309.
- 14 Spjut RW, JM Cassady, T McCloud, DH Norris, M Suffness, GM Cragg, CF Edson 1988. Variation in cytotoxicity and antitumor activity among samples of a moss, *Claopodium crispifolium* (Hook.) Ren. & Card. (Thuidiaceae). Econ. Bot. 42(1): 62–72.
- 15 Spjut RW, DGI Kingston, JM Cassady. 1992. Systematic screening of bryophytes for antitumor agents. Tropical Bryology 6: 193–202.
- 16 Suwanborirux K, C-J Chang, RW Spjut, JM Cassady 1990. Ansamitocin P-3, a maytansinoid, from *Claopodium crispifolium* and *Anomodon attenuatus* or associated actinomycetes. Experimentia 46: 117-120.

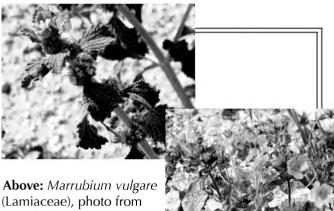
- 17 Newman DJ, GM Cragg. 2016. Natural products as sources of new drugs from 1981 to 2014. J Nat Prod. 79(3):629-61. Note: Table listing anti-viral drugs; none associated with Kern plants mentioned below.
- 18 "Baltimore classification" of viruses. 2017, <u>https://viralzone.expasy.org/656</u>. Accepted by The International Committee on Taxonomy of Viruses (ICTV), https://talk.ictvonline.org/. The disease name, COVID-19, was designated by the WHO. The '19' designated for the year, 2019, that the virus was first seen.
- 19 Salehi B, NV Anil Kumar, B Sener, M Sharifi-Rad, M Kılıç, GB Mahady, S Vlaisavljevic, M Iriti, F Kobarfard, WN Setzer, SA Ayatollahi, A Ata, J Sharifi-Ra. 2018. Medicinal plants used in the treatment of Human Immunodeficiency Virus. Int. J. Mol. Sci. 19: 1459 (60 pp.).
- 20 Notes: Antiviral plants from the preceding reference in the Kern County Flora, L. Maynard Moe, CNPS 2016: Ailanthus altissima, Apium graveolens, Cannabis sativa, Capsella bursa-pastoris, Eclipta prostrata, Equisetum arvense, E. hyemale, Foeniculum vulgare, Galium aparinae, Geum macrophyllum, Glycyrrhiza glabra, Hypericum perforatum, Koelreuteria paninculata,* Lonicera japonica, Melia azedarach, Mentha arvensis, Morus alba, Platycladus orientalis,* Portulaca olercea, Prosopis glandulosa, Raphanus raphanistrum, Rumex crispus, Typha domingensis. Asterisk denotes cultivated plants in Kern County not mentioned in the flora. Olomucine, isolated from Raphanus sativus, is a potent inhibitor of cyclin-dependent kinases (Cdk), which regulates transcription, messenger RNA processing, and the differentiation of nerve cells proteins, and along with the related "purvalanol, are in clinical trials. GM Cragg, DF Newman. 2010. Plant natural products in anticancer drug discovery. Current Org. Chem. 14.
- 21 Notes: Anti-HIV active agents isolated by the NCI from lichen recollections obtained by R Spjut from the United States during 1989 due to polysaccharides (GM Cragg, comm. 1990-91). Ramalin in *Ramalina terebrata* reported by Suh et al. 2017 shows promise for treating colorectal cancer. Molecules 22(8): 361.
- 22 Karunaratne DN, RGU Jayalal, V Karunaratne. 2012. Lichen polysaccharides. IntechOpen: DOI: 10.5772/5102.
- 23 Van Dam, JEG, L AM van den Broek, CG Boeriu. 2017. Polysaccharides in human health care. NPC Natural Product Communication 12(6): 821–830.
- 24 Perez G, RM 2003. Antiviral activity of compounds isolated from plants. Pharmaceutical Biology 41(2): 107-157. Note: Examples of well-known antiviral anticancer plant compounds recognized here are camptothecin and 10-methoxycamptothecin (Camptotheca acuminata, Nyssaceae), emetine (Cephaelis ipecacuanha, Rubiaceae), fagaronine (Zanthoxylum zanthoxyloides (Rutaceae, related to Z. gilletti = Fagara macrophylla), vincaleucoblastine (Catharanthus roseus, Apocynaceae), triptonines (Tripterygium wilfordii, Celastraceae), podophyllotoxin (Podophylum peltatum, Berberidaceae), chaparrinone (Castela spp., Simaroubaceae). Additionally, hypoxanthine in sugar beet, Beta vulgaris (Chenopodiaceae) mentioned without specific antiviral activity and caffeine are reported to suppress the growth of Coxsackie-virus, Echonovirus, Herpes, Poliovirus, vaccinia and influenza virus. Also, dextran sulphate is an anti-HIV active sulphated polysaccharide, reported from Prunella vulgaris, Viola yedoensis, and Alternanthera philoxeroides).
- 25 Cragg GM, DF Newman. 2005. International collaboration in drug discovery and development from natural sources. Pure Applied Chemistry 77(11): 1923–1942.
- 26 Mahomoodally MF, A Gurib-Fakim. 2013. Harnessing traditional knowledge to treat existing and emerging infectious diseases. In M Rai and K. Kon, eds., *Fighting Multidrug Resistance with Herbal Extracts, Essential Oils and Their Components*, Pp. 223–233. Academic Press.
- 27 Wu Xiangmeng, Zhang Qinghao, Guo Jiamei, Jia Yufei, Zhang Ziqian, Zhao Manman, Yang Yakun, Wang Baolian, Hu Jinping, Sheng Li, Li Yan. 2017. Metabolism of F18, a derivative of Calanolide A, in Human Liver Microsomes and Cytosol. Frontiers in

Pharmacology: 8: 479. DOI=10.3389/fphar.2017.00479, https:// www.frontiersin.org/article/10.3389/fphar.2017.00479

- 28 Gustafson KR, JH Cardellina II., JB McMahon, RJ Gulakowski, J Ishitoya, Z Szallasi, NE Lewin, PM Blumberg, OS Weislow, JA Beutler, RW Buckheit, Jr., GM Cragg, PA Cox, JP Bader, and MR Boyd. 1992. A nonpromoting phorbol from the Samoan medicinal plant *Homalanthus nutans* inhibits cell killing by HIV-1. J. Amer. Chem. Soc. 35: 1978–1986.
- 29 Smith AC. Flora Vitiensis Nova II: 558–562, as *Omalanthus* (without the H).
- 30 Esser H-J. 1997. A revision of Omalanthus (Euphorbiaceae) in Malesia. Blumea 42: 421–466 cited in Flora Malesiana, referenced by Welzen, P.C. van (ed.). Flora Malesiana Euphorbiaceae. Naturalis Biodiversity Center, Leiden. <u>www.nationaalherbarium.</u> <u>nl/euphorbs</u>, last updated 8 June 2017. Note: *H. populneus* was first described and named Stillingia populnea by Geiseler in Croton. Monogr. 80 (1807); it cannot be a synonym of *H. nutans* as indicated on many websites.
- 31 Desimio MG, E Giulian, AS Ferraro, G Adorno, M Doria. 2018. In vitro exposure to Prostratin but not Bryostatin-1 improves Natural Killercell functions including killing of CD4+ T Cells harboring reactivated Human Immunodeficiency Virus. Front. Immunol. 9:1514. doi: 10.3389/fimmu.2018.01514
- 32 Newman DJ. 2005. The bryostatins. In: *Anticancer Agents from Natural Products*, eds. GM Cragg, DGI Kingston, and DJ Newman. CRC Press, pp. 137–150.
- 33 Decosterd LA, I C Parsons, KR Gustafson, JH Cardellina II, JB McMahon, GM Cragg, Y Murata, LK Pannell, JR Steiner, J Clardy, M Boyd. 1993. Structure, absolute stereochemistry, and synthesis of conocurvone, a potent, novel HIV-inhibitory naphthoquinone trimer from a *Conospermum* sp. J. Amer. Chem. Soc. 115: 6673–6679. **Note:** The original identification in Sep 1981 was *C. incurvum*. During Spjut's visit to WA in 1992, he received a manuscript copy from Eleanor Bennett who was revising the genus *Conospermum* for the Flora of Australia (1995: 16: 224–271, 453–457). She recognized many new species and subspecies among which one collection in 1991 could be assigned to the unpublished name *C. unilaterale*. During 1991, Spjut collected samples of three other less related smokebush species; none were reported to have conocurvone, whereas the most closely related *C. brachyphyllum*, reportedly has conocurvone in the root.
- 34 **Note:** The KB assay is a culture of human cancer cells of the nasopharynx in artificial media. Activity in KB is defined by a concentration level of the test substance at which 50% of the cancer growth is inhibited (assumed to be proportional to the amount of protein synthesis), expressed as ED50 ≤20 µg/m. The KB assay was routinely employed by the NCI and affiliated institutions in preliminary screening of natural products from 1960–1979.
- 35 Wall ME, H Taylor, MC Wani. 1987. Plant antitumor agents, 24. Rapid 9-KB Assay. J. Nat. Prod. 50: 764–766.
- 36 Spjut RW 2014. A taxonomic review of Western Australian plants screened in KB Cell Culture and other bioassays in the search for new anticancer drugs. World Botanical Memoirs 1: 1–41. <u>http://www.worldbotanical.com/australia.htm#australia</u>. Note: Many Australian plants reportedly used in Aboriginal medicine occur in the subtropical woodland and forests regions, compared to relatively few reports for the Mediterranean flora (p. 29, empirical *obs.*).
- 37 Pidugu LSM, JCE Mbimba, M Ahmad, E Pozharski, EA Sausville, A Emadi, EA Toth. 2016. A direct interaction between NQO1 and a chemotherapeutic dimeric naphthoquinone. BMC Structural Biology (2016) 16:1, DOI 10.1186/s12900-016-0052-x (10 pp.)
- 38 Cragg GM, DF Newman. 2010. Plant natural products in anticancer drug discovery. Current Org. Chem. 14: 14(16):1781-1791. Note: synonym: *Tabebuia avellanedae* (Bignoniaceae).
- 39 Note: Ligand is like a key (small molecule) to a lock (protein) and the unlocking as molecular docking. In virtual screening the problem then is to identify the lock, find the key and determine its correct relative orientation to which way to turn to unlock (molecular docking), or a "best-fit" orientation of a ligand that

binds to a particular protein of interest, such as glove to a hand as in the *OJ Simpson Trial*. Ref.: Morris, GM, Lim-Wilby, M. 2003. Molecular docking. Methods Mol. Biol. 443:365–82 (revised from Wikipedia).

- 40 Qamara MT ul, SM Alqahtanic, MA Alamri, L-L Chen, 2020, in press, online Mar 26. Structural basis of SARS-CoV-2 3CLpro and anti-COVID-19 drug discovery from medicinal plants. J. Pharmaceutical Analysis. <u>https://www.sciencedirect.com/science/article/pii/S2095177920301271</u>. Notes: Closely related to Bat-CoV, SARS-CoV. Eight other plant species scored in the coranaviral screening: *Phaseolus vulgaris* (kidney, pinto, navy, etc. beans) 3,5,7,3',4',5'-hexahydroxy flavanone-3-O-beta-D-glucopyranoside, *Camellia sinensis* (tea plant) myricetin 3-O-beta-D-glucopyranoside, *Myrica cerifera* (southern wax myrtle) myricitrin, *Glycyrrhiza uralensis* (Chinese licorice) licoleafol, *Amaranthus tricolor* (mostly horticulture) amaranthin, *Fraxinus sieboldiana* (Siebold ash), calceolarioside B, *Phyllanthus emblica* (Indian gooseberry), (2S)-eriodictyol 7-O-(6''-O-galloyl)-beta-D-glucopyranoside, *Hyptis atrorubens* (marubio oscuro) methyl rosmarinate.
- 41 Salem MM, KA Werbovetz. 2006. Isoflavonoids and other compounds from *Psorothamnus arborescens* with antiprotozoal activities. J. Nat. Prod. 69: 43–49.
- 42 Salem MM, J Capers, S Rito, KA Werbovetz. 2011. Antiparasitic activity of C-geranyl flavonoids from *Mimulus bigelovii*. Phytother Res. 25(8):1246-9. **Note:** *Mimulus bigelovii* recently reclassified, *Diplacus bigelovii*.
- 43 Spjut RW. 2010. Potential harvest sites of *Veratrum californicum* in relation to its taxonomy, geographical distribution, and presence of cyclopamine and cycloposine. Society for Range Management: Symposium—Medicinal Uses of *Veratrum*. Annual Meeting with Weed Science Society of America, Denver, CO, Feb 7–11 (30 minute oral presentation, no abstract, unpublished). Note: See also Mimulus Memo, June 2011, and WBA website, *Veratrum*, <u>http://www.worldbotanical.com/veratrum.htm</u>



Above: Marrubium vulgare (Lamiaceae), photo from near Mojave, California, May 2005. An invasive species that showed antiparasitic activity. Native to Europe, SW Asia to NW China



(Xinjiang). Tea is prepared from the dried plant for debility and colds, also used in certain candies for coughs and sore throat, or as an expectorant, diaphoretic, and laxative (Flora China 17: 104. 1994).

Below: *Mimulus bigelovii* (**Bigelow's Monkeyflower**, Phrymaceae, *Diplacus bigelovii* var *bigelovii*), eastern Mojave Desert, Mesquite Mts.,CA - Mar 2005 contains active antiparasitic C-geranyl flavonoids. See Calflora for additional images at this location including voucher specimen for the original active sample investigated for new compounds to treat leishmaniasis.

Individual – \$45 Family or Library – \$75

CNPS-Kern Chapter c/ o Dinah Campbell, Editor 3806 Dalehurst Drive Bakersfield, CA 93306

mimulus.memo@gmail.com



Inside this Issue:

MEETING DATES & TOPICS A ANNIE MONTAGUE ALEXANDER, PART II PRESIDENT'S MESSAGE: ANTI-VIRAL PLANTS I KNEW THAT: EDIBLE PLANTS COLUMN ADOBE HOUSE PROJECT UPDATE