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# Medicinal Importance of the Genus Agave S. K. HAZRA, A. SAHA, A. K. MAHAPATRA, D. GUPTA, DIPANKAR SAHA, A. K. DAS AND H. S. SEN

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#### Abstract

The leaves root, sap and core of A. sisalana, A. americana, A. cantala, A. angustifolia and A. vera-cruz are used for the treatment of various human diseases, particularly veneral diseases. These five species are rich sources of different sapogenins of medicinal importance, Of the sapogenins hecogenin is the starting material for the synthesis of cortisones, cortisol, prednisolone, prednisone, dexamethagone, beta methasone, triameinalone and others. Similarly, tigiget in is used for prednisolone, hydrocortisone and oral contraceptive cyproterone. Attempts on microbial transformations of other sapogenins are underway. Hecogenin may be converted to pregone derivatives, specially 16 DPA which has great demand in pharma cutical industries. In vitro synthesis of hecogenin from A. sisalana and A. ammtensis is a distinct possibility.

Agaves are a source of food beverage, sugar, syrup, fiber, cellulose, wrap paper for cullinary use, silage for livestock, ornamental plants, wax, methane gas and many other products (1-5). There is a recent trend all over the world for the greater use of plant based drugs of the traditional Indian systems for the treatment of human diseases for various reasons. The detailed account of this ongoing de velopments are available in various literature (6-11). Keeping this upsurge on phytomedicines in view, and the fact that only 5-15% of the approximately 250,000 species of higher plants have been systematically investigated, the potential of others need to be explored for new sources of phytomedicines. This rely on the clues obtained from historic, traditional, folkloric or ethnobotanical uses of plan parts as medicine (6-8). In this context, the genus Agave, being grown in varied edaphoclimatic conditions in India, offers excellent opportunities for direct therapeutic uses. In addition, this genus also has rich load of

various sapogenins which may be used for the manufacture of corticosteroids having great demand in pharmaceutical industries world over.

As we turn our focus on Agave for medicinal materials; it is worthwhile to mention that several modern techniques (NMR, 2DN-MR, multidimensional NMR, HPLC, IRS, UV mass spectrometry) augmented by sophisticated instrumentation, are currently in vogue for isolation and characterization of bioactive principles in other crops. Drug designing, modifications, combinatorial chemistry and other tools are currently being employed for pharmacological analysis (11-14). These techniques, if employed in Agave, will generate quick and accurate information to develop an inventory on Agave for future drug research. The medicinal potentialities of Agave are yet to be explored in a big way because there remained an information gap between detailed ethnomedical uses and chemical constituents of different plant parts. This article has been

designed to fulfill this gap. Finally, the local names of each species have been provided for initial identification but standard botanical description must be followed for commercial exploitation.

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# The Genus Agave, in General

The leaves, as also the root when cut, yield a saccharine juice (nauseous odor and acid taste) which has resolvent and alternative properties (a medicine that makes a change in the functions) especially useful in syphilis. The sap is laxative, diuretic (promotes the flow of urine), emmenagogue (promotes menstrual flow); used in scurvy, scrofula (inflammation of lymph node due to tuberculosis with glandular swelling) and cancer. The large, moist, fleshy leaves are used as poultice; the fresh juice is applied to bruises and contusions (bruise without rupture of blood vessels). The pulp of the leaves placed between folds of muslin is applied to the eye in conjunctivitis (inflammation of the mucous membrane covering the anteri or surface of the eye-ball); it is also used, mixed with sugar, in gonorrhea. A decection (four ounces/one pint of water) of syphilis roots is employed in secondary (15, 16).

Agave sisalana Persine ex. Engelm. Syn.

A. rigida Mill Var sisalana Englem (Sisal)

The juice of the leaves is echolic (moulting effect) and lowers blood pressure. Roasted sisal leaves are used to remove headache and pain in knee joints. The juice extracted from roasted leaves, mixed with 10—20 black pepper seeds, is used to treat swelling of throat of cattle. The sisal dust may cause pulmonary fibrosis to the workers engaged in

making ropes and twines. During decortication of leaves for fiber extraction, a pulpy material is separated out which contains, apart from usual chemical, hecogenin (0.1%) in cobination with tigogenin, wax and 11% fer mentable sugars, the latter is, however, not profitable to prepare alcohol (13—19).

#### Chemical Constituents

A process for isolation of hecogenin from juice has been standardized (17, 20); 5α-Pregn-16-en-3β-Ol-20-one, mp 202 C, was isolated (21,22).

Neotigogenin, sisalagenin, gloriogenin, 9, 11-dehydro hecogenin, diosgenin and yamogenin isolated from leaves; two new trihydroxysapogenins-hainangenin and hongguangenin-isolated along with known compounds—tigogenin, neotigogenone, neotigonenin, sisalagenin, hecogenin,  $5\alpha$ -pregnan- $3\beta$ ,  $20\beta$ -diol, 12-epirockogenin, chlorogenin and  $\beta$ -sitosterol from leaves (23, 24).

#### Occurrence and Distribution

Sisal extensively cultivated in India in Nil Dongri, Jharsuguda, Manasanikani, Sonpur and Kardadhaman areas of Orissa, and to certain extent in Palamau district of Bihar, Bilaspur of Madhya Pradesh and pockets of Tamil Nadu, Andhra Pradesh, Karnataka, Maharastra and West Bengal.

#### Local Names

English: Indian agave, sisal, sisal hemp, sisal agave, Bahama hemp; Bengali: sisal; Assam: sonn.

#### Agave americana Linn

If the central bud be lopped off at the flowering season, the cut stem discharges freely a sour liquid, which ferments rapidly and forms the Pulque Beer of the Spaniards, or, by distillation, a kind of brandy known as Mexical. Beside this, in Mexico, a strong spirit is prepared from the sap, known as

Mezcal, also a kind of brandy of 80 degrees of strength, a sweet, thick substance resembling honey, a concentrated gum used in medicine, brown-sugar, leaf sugar, sugarcandy, and vinegar of excellent quality, so that this is considered one of the most important economic plants of Mexico (15).

Leaves are used as laxative, emmenagogue, diuretic and also useful in scurvy. They are employed as a resolvent in syphilis, scrofula and cancer. Slices of leaves applied as poultice (16) and used in bruises and rheumatism (pain and stiffness in the joints or muscles of the body). The pulp of leaf is a good remedy for gonorrhoea. The leaf-juice is used for warts, cancerous ulcers and putrid tumors (25).

The roots are diuretic, diaphoretic (promotes sweating), antisyphilic and are said to find their way to Europe mixed with Sarsaparilla. The core of the plant is used for the treatment of ascites (abnormal accumulation of serous fluid in peritoneal cavity), dropsy (neurological disorder), venereal sores, dysentery (15, 20). Various parts of plant are used for ascites (abnormal accumulation of serous fluid in peritonial cavity), anasarca (accumulation of fluid in subcutaneous connective tissues), toothache; and as antiseptic, antipyretic and alternative. In homocopathy, mother tincture is used in the sore in the mouth, in scurvey, in afflictions, in teeth, pains and swelling in legs, in constipation, gonorrhoea (25).

Chemical Constituents

Steroid sapogenins—hecogenin (0.15%), 5., 25 D spirostan-3β-cl-12-one (0.17%) mp 252 C, chlorogenin, 5., 25D spirostan-3β, 6. diol (0.13%), and two other compounds, mp 259 C and 273 C isolated from leaf. Seeds yielded neotigogenin, kammogenin along with hecogenin. Altogether, ten steroidal saponins

-agavosides A, B, C, C', D, E, E, G, H and I—isolated from the leaves; common aglycone was hecogenin, while sugars were glucose, galoctose, xylose and rhamnose. Structures of most agavosides A, B, C, C', D, E, H and G elucidated as Agavoside A (R=Gal); Agavoside B (R=Gal (4→1) Glu; Agavoside C  $(R \pm Gal (4 \rightarrow 1) Glu (4 \rightarrow 1) Glu)$ ; Agavoside  $C'(R=Gal(4\rightarrow I))Glu(4\rightarrow I)Glu(2\rightarrow I)Glu);$ Agavoside D (R=Gal (4-1) Glu (4-1) Glu  $[(2\rightarrow 1) \text{ Glu}] (3\rightarrow 1) \text{ Rha}$ ; Agavoside E (R=  $Gal(4\rightarrow 1)$   $Glu(4\rightarrow 1)$   $Glu[(2\rightarrow 1)]$   $Xyl](3\rightarrow 1)$ Rha (4→1) Rha; Agavoside G (R=Gal (4→ 1) Glu  $(4\rightarrow 1)$  Glu  $f(2\rightarrow 1)$  Xy $f(3\rightarrow 1)$  Rha); Agavoside H ( $R = Gat (4 \rightarrow 1) Glu 4 \rightarrow 1$ ) Glu  $[(2\rightarrow 1) \text{ Xyl}] (3\rightarrow 1) \text{ Rha } (4\rightarrow 1) \text{ Rha respecti-}$ vely and anticancer activity was found to be positive (22, 26, 27). Cholorogenin, kaemferol-3-glucoside and 3-rutinoside isolated from flower (25).

# Occurrence and Distribution

The century plant is extensively used for hedges in many parts of India such as east and west coasts, upper gangetic plains and Punjab. It is at home under widely different climatic conditions, various altitudes and poor agroresource base.

# Local Names

English: The century plant, american aloe, carata; Sanskrit: Kalakantala; Hindi: Bauskeora, Barakanwar, Kantala; Bengali: Bilatipat, Koyan; Gujreti: Jangli Kunrara, Jangli Kunvara; Telegu: Kithanara, Rakashimatalu; Tamil: Rakashimatalu; Mafayalam: Bhuttale, Budkkattalenaru; Uriya: Birhotokumari; Punjabi: Kantala.

Agave angustifolia Haw, syn. A. wightii
Drummond and Prain

Lodhas use dried leaf powder in warm water (3:1) as laxative. Leaves contain hecogenin, tigogenin and gitogenin (16). Lodhas

apply root paste as oure for septic wounds. During treatment period, they do not allow the patient to take sugar and chilli. Santhals apply juice obtained from squeezing fermented leaves about 5 ml with paste of 21 black peppers as cure for swelling of throat of cattle (28). The juice of leaves is applied to bruises and roots are diuretic and diaphoritic (20). The plant extract is blood pressure depressor and found to be anti-microbial in vitro (25).

## Occurrence and Distribution

It is naturalized in the sub-Himalyan tract and outer Himalayas and many other parts.

#### Local Names

English: The dwarf aloe; Sanskrit: Kantala, Telegu: Bulurakkasi, Kittanara, Samata, Tamil: China erumaikkattalai, Seemakathaalai, Hindi: Khetki.

## Agave cantala Roxb.

The leaves are used in boils, burn, cut and wound and as diuretic, antiseptic, blood purifier, laxative and analgesic, remedy for urinary infection, a general tonic. The leaf-paste is used to relieve pain and decoction is used for the treatment of scanty menstruation (25). Roots are anthelmintic (destroys or expels intestinal worms), used in diarrhea, fever, goiter (chronic enlargement of the thyroid gland), paralysis and skin diseases. The plant is used by the Santhals in anasarca ascites, dropsy, dysentery and syphilis (19).

The leaves contain hecogenin (0.15%) which can be recovered from the wastes after fiber extraction. Leaves also contain saponasess suitable for conversion of *Discorea* saponin to diosgenin (29). Presence of diosgenin (0.03%) along with hecogenin reported (22, 30).

#### Chemical Constituents

New saponin—cantalanin A isolated from leaves and characterized (27), isolation and structure elucidation of cantalanin B from fru-

its were done (31),  $\beta$ -sitosterol, tigogenin, hecogenin, 9-dehydrohecogenin, gitogenin, manogenin, 9-dehydromanogenin and chlorogenin isolated from leaves (32); new steroidal saponin (I) isolated from roots and its structure determined (33); tigogenin, diosgenin, hecogenin, gitogenin and chlorogenin isolated from root stalks (34).

Occurrence and Distribution

East and west coast, upper Gangetic parts and Panjab.

#### Local Names

Bengali: Bilati anarash, Punjab: Kantala, Kitki.

Agave vera-cruz Mill

Stem is a rich and cheap source of polyfructosans of the type of insulin and its isomers. These branched fructans have been indicated to be an useful chemotaxonomic aid (35). The plant is used as pergative (20).

#### Chemical Constituents

The juice of leaves contains hecogenin (0.1%) only (17, 20). Extraction and isolation process for hecogenin from juice standardized (17). 9 (II) dehydrohecogenin isolated from leaves; trisaccharides— I-kestose and neokestose-tetrasaccharides-nystose and another related to neokestose isolated from stem (23, 27, 36).

#### Occurrence and Distribution

Andhra Pradesh, Maharashtra, Karnataka and Madhya Pradesh, Assam, West Bengal, Mumbai.

# Local Names

Hindi: Kuwarbuti, Rakaspatta, Tamil: Kattazha, Telegu; Kattalai.

# Sapogenins and Future Prospects

Corticosteroids, sex hormones and contraceptive pills belong to steroid drugs, manufactured from hecogenin (C<sub>27</sub>H<sub>42</sub>O<sub>4</sub>) and diosgenin, belonging to the group of "sapogenins" Sapogenins has the basic structure of steroid

- 4. Hazra S. K., A. K. Das and S. Das.
  2001. In vitro synthesis of hecogening from sisal (Agave sisulana Perrime ex Engelm). Plant Tissue Culture. 11:77

  —84.
- 5. Hazra S. K., S. Das and A. K. Das. 2001. In vitro shoot induction from rhizome buds of sisal (Agave sisalana Perrex Engelm) for obtaining quick planting material. J. Plant Biol. 28: 165-171.
- Chatterjee A and S. C. Pakrashi. 1991.
   The treatise on Indian medicinal plants, volume I. Publ. Inform. Direct., CSIR, New Delhi, India.
- 7. Leung Y. A. and S. Foster. 1995. Introduction. Enclycopedia of common natural ingredients. A Wiley Inter Sci. Publ., John Wiley and Sons, Inc., London, UK.
- 8. Natesh S. 2001. The changing scenario of herbal drugs: Role of botanist. Phytomorphology. Pages 75—96 in N. S. Rangaswamy, editor. Golden Jubilee Issue 2001: Trends in plant sciences.
- 9. Cragg M. G. and D. J. Newman. 2002. Biodiversity: medicinals for the millennia. Sci. Cult. 68: 3-10.
- 10. Ghosal S. 2002. Herbal health-products in global perspective. Sci. Cult. 68: 33

   40.
- 11. Ser S. 2002. Legacy of international health care: drug resistance, drug development and therapeutic uses. Sci. Cult. 68: 45-55.
- 12. Ganguly S. 2002. Combinatorial chemistry—emerging technology in synthesis of new drug molecules. Sci. Cult. 68: 56-58.
- CSIR. 1948 The wealth of India. CSIR, New Delhi, India.
- 14. Lock G. W. 1962. Sisal, Longmans,

- Green and Co. Ltd., London, UK.
- Watt G. 1972. A dictionary of the economic products of India. Cosmo Publ. 1:133-144.
- Ambasta S. P. 1992. The useful plants of India. CSIR, New Delhi, India.
- 17. Murthy M. S. and S. S. Lokras.. 1993.

  A comprehensive report on technologies developed for sisal based industries. Indian Inst. Sci., Bangalore, India, pp. 1—86.
- Mahapatra A. K., D. Gupta and A. Saha. 1999-2000. Annual report. NATP on Plant Biodiversity. CRIJAF, 20.
- Pakrashi S. C. and S. Mukhopadhyay.
   2001. Medicinal and aromatic plants of West Bengal (Midnapore). DST-NES, Govt. of West Bengal and West Bengal Acad. Sci. Tech., Kolkata, India.
- Chopra R. N., S. L. Nayar and I. C. Chopra. 1980. Glossary of Indian medicinal plants. CSIR. New Delhi, India.
- Rubin M. 1967. Recovery of tigogenia acetate and 5α-pregn-16-en-3β-ol 2-one from natural plant sources. U. S. 3, 303, 187.
- Rastogi R. P. and B. N. Meherotra. 1991.
   Compendium of Indian medicinal plants.
   Central Drug Res. Inst., Lucknow and CSIR, New Delhi, India. 1:15-16.
- 23. Rastogi R. P. and B. N. Meherotra. 1993. Report on medicinal plants. Central Drug Res. Inst., Lucknow and CSIR, New Delhi, India. 2:20—22.
- 24. Chu-Pu T., C. Yen-Yong, and L. Huant. 1976. Studies on the steroidal sapogenins of the Agave plant IL. Isolation and identification of the steroidal sapogenins from leaves of Agave sisalana Perrine. Hua Hsueh Pao. 34:179—196.
- 25. Guhabakshi D. N., P. Sensarma and D.

- C. Pal. 1999. A lexicon of medicinal plants in India. Naya Prakash, Calcutta, India.
- Jain S. K. 1991. Dictionary of Indian folk medicine and ethnobotany. Deep Publ, New Delhi, India.
- 27. Rastogi R.P. and B. N. Meherotra. 1993. Compendium of Indian medicinal plants. Central Drug Res. Inst., Lucknow and CSIR, New Delhi, India.
- Pal D. C. and S. K. Jain. 1998. Triba medicine. Naya Prakash, Kolkata, India.
- Chopra R. N., I. C. Chopra and B. S. Verma. 1959. Supplement to glossary of Indian medicinal plants. CSIR, New Delhi, India.
- Chakraborty R. N., M. N. Mitra and D. Chakraborty. 1959. Steroid sapogenins from Indian Agaves (sisal). Bull. Calcutta. Sch. Trop. Med. Hyg. 7: 5.
- 31. Varshney I. P., D. C. Jain and H. C. Srivastava. 1982. A triglucoside of hecogenin from fruits of Agave cantala. Phytochemistry 21: 239—240.
- 32. Sharma S. C., O. P. Sati and H. C. Sharma. 1982. Study of the natural constituents of *Agave cantala* Roxb. (leaves). Pharmazie 37: 305-306.
- Sharma S. C. and O. P. Sati. 1982. A spirostanol glycoside from Agave can-

- tala. Phytochemistry 21: 1820—1821.
- 34. Sati O. P. and G. Pant. 1983. Steroidal constituents of *Agave cantala* Roxb (root stocks). Pharmazie 38: 353.
- 35. Sims I. M., A. G. Cairns and R. H. Furneaux. 2001. Structure of fructans from excised leaves of New Zealand flax. Phytochemistry 57: 661—668.
- Lambertus D., P. J. Kamerling, F. G. J. Vliegenthart and N. H. Satyanarayana. 1977. Oligosaccharides isolated from Agave vera cruz. Carbohydr. Res. 54: 275-284.
- Patel A. V, G. Blunden and T.A. Crabb.
   1994. Transformations of solasodine and derivatives of hecogenin by Cunninghamella elegans. Phytochemistry 35: 125-133.
- 38. Smith M. A. L., H. Kobayashi, M. Gawienowski and D. P. Briskon. 2002. An in vitro approach to investigate medicinal chemical synthesis by three herbal plants. Plant Tiss. Org. Cult. 70: 105—111.
- 39. Andrijany V. S, I. Gunawan and L. A. Adi Soehono. 1999. Simultaneous effect of calcium, magnesium, copper and cobalt ions on sapogenin steroids content in callus cultures of Agave amaniensis. Plant Cell Tiss. Org. Cult. 55: 103—108.