Bitter Melon

500 mg per capsule  
100 or 300 capsules per bottle

Concentrated Herbal Formula

For Professional Use Only

INDICATION:
Bitter Melon is intended to be used to supplement the diet when following programs designed to maintain blood sugar levels.

EXPLANATION:

In our Bitter Melon Extract, we use the whole plant; leaves, seeds, vine, and melon. Research has found that the leaves are nutritious sources of calcium, magnesium, potassium, phosphorus, and iron; both the edible fruit and the leaves are great sources of the B vitamins.

Diabetes
The diabetes disorder stems from the way the body processes carbohydrates. Normally, carbohydrates are converted into a form of sugar called glucose, which floats along in the bloodstream until the pancreas goes into action. The pancreas produces insulin, a hormone that signals body cells to soak up glucose. Once inside the cell, glucose is either used to produce little or no insulin or else becomes resistant to the hormone’s action and can’t compensate. Either way, the glucose can’t get into the cells; it accumulates in the blood and is later expelled in the urine. In short, blood sugar rises while cells starve.

Both kinds of diabetes (Type I is regarded as insulin-dependent diabetes mellitus, Type II is considered to be non-insulin dependent diabetes mellitus) can be managed quite well with dietary changes, moderate exercise, effective weight reduction, and natural supplements such as herbs, vitamins, minerals, amino acids, and enzymes. Herbs, in particular, are especially beneficial in dealing with this particular health problem. What one would find in medicinal plants isn’t so much as a “cure”, as a safe reliable way to better manage body blood sugar levels.

Diabetics who have routinely used other herbs have reported what may be described as a “seesaw” action in their hypoglycemic activities: an alternating influence in which their hypoglycemic effects may be felt sometimes, but not other times. What would be nice is to have an herb that is always consistent in its blood sugar-lowering properties.

The herb that meets this criterion in every way is Bitter Melon. There are several different kinds of Bitter Melon throughout the world, but the Chinese Bitter Melon (K’u Gua) has proven to be the most effective and consistent in stabilizing blood sugar levels.

Indian biochemists reported isolating a hypoglycemic peptide from the fruit, seeds, and tissue of Momordica, which they labeled Polypeptide-p (also called p-insulin or v-insulin). Based on a medical hypothesis, it could exert a direct action on the islets of Langerhans located within the pancreas that are the source of insulin and glucagon (a pancreatic hormone); in so doing, it might stimulate the release of insulin when needed most. Bitter Melon also relieves the dehydration that frequently accompanies diabetics. Our extraction and concentration process yields one gram of extract from 25 grams of fresh plant material.

Anti-HIV effects
Bitter Melon contains three anti-HIV proteins: alpha- and beta momorcharin, and MAP-30. They have been tested in vitro and found to possess inhibitory effects on HIV-infected macrophages and T-cells and inhibit cell to cell infection and syncytial formation. They can inactivate ribosome’s functions in HIV-infected cells. Thus they can stop protein synthesis and kill infected cells. Significantly, these proteins do not appear to be toxic against uninfected cells.

Anti-viral effects
These proteins isolated from extracts of Bitter Melon have been found to inhibit replication of herpes simplex (HSV-1) and polio virus.
**INGREDIENTS:**

<table>
<thead>
<tr>
<th>Latin Name</th>
<th>Pinyin Name</th>
<th>English Name</th>
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<tbody>
<tr>
<td>Momordica Charantia</td>
<td>K’u Gua</td>
<td>Bitter Melon</td>
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**RECOMMENDED DOSAGE:**

2-10 capsules per day taken with meals. A typical course of therapy would be to start by taking 1 capsule in the morning and the evening for the first week. Increase to include noontime dosage the second week and subsequently increasing capsules by one at each time interval until desired results are achieved.

**CAUTIONS AND CONTRAINDICATIONS:**

**WARNING:** Not to be used during pregnancy.

**ANALYSIS OF FORMULA:**

**Actions of Ingredients:**

Momordica Charantia  
(Bitter Melon)  
_\textit{K’u Gua}_

A bitter, pungent, warming herb that acts as an expectorant, soothes irritated tissues, reduces inflammation, clears toxins, and has anti-viral properties.
ABSTRACT

Clinical study of an insulin-like compound obtained from vegetable source (vegetable insulin) was carried out on nine patients with diabetes mellitus. The active hypoglycaemic principle, purified protein extract, was obtained from fruits as well as from tissue cultures of the plant *Momordica charantia* L. This extract was homologous to insulin obtained from animal pancreas. It showed a consistent hypoglycaemic effect in patients with diabetes mellitus. The average fall in blood sugar level at the peak effect of vegetable insulin was found to be statistically significant. The onset of action was within 30-60 min with the peak effect six hours after the administration of the dose of plant insulin. No hypersensitivity reaction to this extract was observed in the group of patients studied.

INTRODUCTION

A crude extract obtained from the fruit of a plant known a *Momordica charantia* L. (bitter gourd), has been shown to possess hypoglycaemic activity when tested in rabbits (1, 2, 3). However, this extract was also found to have many side effects, including uterine haemorrhage in gravid female rabbits. An intraperitoneal injection of this extract invariably caused the death of the experimental animal. Khanna & Mohan (4) were able to extract an abortifacient factor present in this crude extract and isolated diosgenin from the fruit as well as from the in vitro tissue cultures of this plant. Subsequently they were able to extract out the active principle in a pure protein from (vegetable insulin, v-insulin) which could be used as a hypoglycaemic agent in human beings after intraperitoneal injection of this extract invariably caused the death of the experimental animal. Khanna & Mohan (4) were able to extract an abortifacient factor present in this crude extract and isolated diosgenin from the fruit as well as from the in vitro tissue cultures of this plant. Subsequently they were able to extract out the active principle in a pure protein from (vegetable insulin, v-insulin) which could be used as a hypoglycaemic agent in human beings after

<table>
<thead>
<tr>
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<th>Name</th>
<th>Sex</th>
<th>Age (y.)</th>
<th>Body height (cm)</th>
<th>Body weight (kg)</th>
<th>Duration of disease (y.)</th>
<th>Type of diabetes</th>
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<td>JC</td>
<td>M</td>
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<td>2</td>
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<td>50</td>
<td>162</td>
<td>60.5</td>
<td>1</td>
<td>Mature onset</td>
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Table II. Dose of vegetable insulin, according to blood sugar levels

<table>
<thead>
<tr>
<th>Severity of diabetes</th>
<th>Fasting blood sugar level (mg%)</th>
<th>Dose of vegetable insulin</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mild</td>
<td>Less than 180</td>
<td>10 units</td>
</tr>
<tr>
<td>Moderate</td>
<td>180 – 250</td>
<td>20 units</td>
</tr>
<tr>
<td>Severe</td>
<td>250 and above</td>
<td>30 units</td>
</tr>
</tbody>
</table>
These patients were placed in two groups, depending on their stage of carbohydrate decompensation, as follow:

(i) Overt, clinical diabetes, i.e. patients with elevated fasting and random blood sugar levels (more than 110 mg per 100 ml). They were further subdivided into juvenile and maturity onset types, depending on the age at onset of diabetic symptoms.

(ii) Asymptomatic diabetes, i.e. patients with normal blood sugar levels (less than 110 mg per 100 ml) but with an abnormal intravenous glucose tolerance test.

There were 6 patients with juvenile diabetes, one with maturity onset and 2 with asymptomatic diabetes mellitus (Table I). All antidiabetic medication in the form of insulin or oral hypoglycaemic agents was stopped 24 hours prior to the study. A fasting blood sugar sample was taken at 7 a.m. before giving v-insulin. The dose of v-insulin was varied according to the severity of the disease and was decided upon arbitrarily according to the fasting sugar level (Table II). Vegetable insulin was administered subcutaneously at regular intervals. The first three samples were drawn at half-hour intervals (7:30 a.m., 8:00 a.m. and 8:30 a.m.) to establish the onset of action and the subsequent samples were collected at 11:00 a.m., 1:00 p.m., 3:00 p.m. and 7:00 p.m. to determine the peak effect and duration of action of v-insulin. All samples were collected intravenously. Blood sugar estimation was done on whole blood by the method described by King & Wooten (7). All the subjects were kept fasting during the interval for collection of samples and only plain lemon water was given if desired by the patients. A provision was kept for the administration of glucose in the event of the development of hypoglycaemic symptoms.

Five healthy volunteers (control group I) and 5 patients with overt diabetes mellitus (control groups II) served as controls. A placebo injection was used. The control subjects were kept fasting and blood samples were collected and analysed in a similar way. The administration of v-insulin to control subjects was avoided due to its inherent hypoglycaemic properties, as evaluated in animal experiments.

**RESULTS**

The placebo injection in the control groups did not produce any appreciable reduction in blood sugar levels at different intervals. A definite hypoglycaemic effect of v-insulin was observed in the patient group in this study. The onset of vegetable insulin effect was observed within ½-1 hours, with the peak effect after 4 hours in 6 juvenile diabetics, after 6 hours in 2 patients with chemical diabetes mellitus, and after 12 hours in one patient with maturity onset of diabetes mellitus.

All values in this study were analysed statistically by applying the paired t-test and the calculated t was 3.3 and the tabulated t was 2.3 at d.f 8*, which was highly significant for the diabetic patient group compared with the healthy and diabetic controls during peak hours.

The hypersensitivity reactions were conspicuously absent after administration of vegetable insulin and there was no local reaction at the site of injection.

**DISCUSSION**

The present investigation revealed that vegetable insulin has a consistent hypoglycaemic effect in patients with diabetes mellitus. The onset of action is similar to that of standard zinc crystalline insulin (30-60 min). However, the peak effect of vegetable insulin was seen after 4-12 hours as compared with, for 2-3 hours regular insulin. The greatest fall in blood sugar levels observed in the patient group was found to be statistically significant. There were no anaphylactic reactions to vegetable insulin however, as regards its long-term use, further studies are required in order to evaluate its antigenic properties.

The availability of vegetable insulin should open new horizons in the treatment of diabetes mellitus, especially where it is taboo to use animal products. Since the active principles are derived from a vegetable source, it can be obtained in abundance. Further clinical trials are needed in order to establish its duration of action, assay, antigenicity and various effects on intermediary metabolism in human beings.

**ACKNOWLEDGEMENT**

The authors are grateful to Dr P. Khanna, Department of Botany, University of Rajasthan, Jaipur, for providing the vegetable insulin used in this study.

**REFERENCES**

INTRODUCTION: Herbal medicine was the main form of treatment for illness before the advent of Western medical practice, and still is today in 75 to 90% of the rural areas of the world (1). Each area of the world has its own form of ethnomedicine based on the plants found in that area (1,2,3,4,5). Japan and China are two countries with a long history of exchanging herbal lore extending over many thousands of year’s (3,6). Their particular form of herbal medicine is called Kampo (3,6). The integration of Western medicine and Kampo in Japan, although slow, was not entirely antagonistic (6). In 1774, Gentaku Otsuki, pointed out in one book that both Eastern and Western medicine have their weaknesses and encouraged practitioners of both systems to adopt a positive attitude towards the West so that medicine could eventually evolve into a better system (6).

The first example of the Integration of Eastern and Western medicine was achieved by Dr. Seishu Hanaoka (1760-1835). He developed the general anesthetic, tsusensan, from Kampo herbs and used it to perform a mastectomy for cancer (6). Since that time, the scientific study of many herbal medicines have revealed biologically active compounds that have been adopted into Western pharmaceuticals. The drugs isolated from plants are particularly useful as they are more often complex compounds yet they occur in a biologically active form. Because of their complexity, it is also very unlikely that the plant compounds would have been synthesized a priori in the search for new compounds (7). When herbal medicine leads are exploited, broad classes of compounds can be developed. A few examples of the variety of common pharmaceuticals of herbal origin are outlined below.

The well-known bronchodilator and antitussive ephedrine, comes from the herb Ephedra sinica, commonly used in kampo prescriptions for conditions such as bronchial asthma (8). The discovery of ephedrine led to the development of all the narcotic analgesics (7). The bark of Cinchona is found to contain quinine which led to the development of the antimalarial drugs (3,7). The traditional herbal extract from rhubarb (Rheum spp.) contains several active compounds. These compounds mediate several medicinal effects of rhubarb such as its purgative action (sennosides), its anti-bacterial, anti-fungal, and antitumor activities (anthraquinones), Its anti-Inflammatory activities and analgesic activities (lindelypin), and Its Improvement of lipid metabolism (stilbenes) and nitrogen metabolism (rhatannin) (9).

Vincristine, derived from the Madagascar periwinkle (Catharanthus roseus), is highly effective and extensively used in the treatment of leukemias (8,10). However, the yield of vincristine from the periwinkle Is extremely low and every available soil suitable for the growth of this plant is under development (1).

Many plant compounds have been identified with antiviral or cytotoxic antitumor characteristics (11-20). One such plant, Momordica charantia, will be discussed in detail in the following section.

Momordica charantia

The plant Momordica charantia belongs to the family cucurbitaceae and is known by many common names: balsam pear, African cucumber, cindemor, bitter apple, bitter melon, bitter gourd, carilla plant, wild cucumber, margose. concombre African (4), the herb Kuguza (China) (21), and Karela (Pakistan) (22). It is common in China, India, and Africa where it has a history of several medicinal uses (4). The species has become naturalized in the southern United States where the unripe fruit is eaten and used in home remedies for treating colds (3).

Momordica charantia is not only used for medicinal purposes, but also as a general insecticide, which is beneficial in tropical countries where insects are a common vector of many diseases (4).
Yet others (28,29), found no hypoglycemic activity in an aqueous extract of the dried fruits of the bitter melon on alloxan induced diabetic mice. Alloxan induces diabetes by destroying all β-cells of the pancreas. This indicates that the bitter melon hypoglycemic activity may work to some extent via these cells. This is partially supported through experiments conducted by Welinhinda et al. (30). These workers demonstrated that an aqueous extract of the unripe fruits of the bitter melon is a potent stimulator of insulin release from β-cell-rich pancreatic islets isolated from obese-hyperglycemic mice. This induced release of insulin is not suppressed by epinephrine nor by the removal of calcium. These authors suggest that the insulin releasing effect is a result of membrane perturbations as the bitter melon extract is found to mimic the action of saponin in inhibiting the Ca²⁺/H⁺ exchange mediated by the ionophore A23187 and inhibiting the release of Ca²⁺ from preloaded liposomes (30).

The hypoglycemic effect is attributed to two factors: a substance labeled charantin, a crystalline fraction obtained from an alcoholic extract of the fruit, and a polypeptide (p-insulin, or v-insulin, for vegetable-insulin) obtained from the fruit, seed and tissue (31,27,4). Charantin, administered at 50 mg/kg doses reduces hyperglycemia in rabbits by 42%. Charantin possesses pancreatic and extra-pancreatic action, and has a slight antispasmodic and anticholinergic effect. Charantin helps to control diabetic patients (5), but should not be viewed as curative.

The p-insulin is structurally and pharmacologically comparable to bovine insulin. The purification of p-insulin can be carried out in a manner similar to the purification of insulin from the pancreas of animals (31). Amino acid analysis of p-insulin indicates that it has 166 residues composed of 17 different amino acids and a molecular weight of approximately 11,000. The p-insulin is found to be composed of two polypeptide chains bound together by disulfide bonds (31). Although similar in many aspects, immunoassays did not indicate any cross-reactivity between p-insulin and bovine insulin (27). Subcutaneous and intramuscular administration of the polypeptide, p-insulin isolated from the fruit, produced hypoglycemic activity in diabetic gerbils, langurs and humans. In juvenile diabetic patients, the peak effect was observed after 4-8 hours as compared to 2 hours for bovine insulin (31,27,4). Hypersensitivity reactions to p-insulin are absent in patients previously treated with animal insulin, but studies on juvenile diabetic patients, the peak effect was observed after 4-8 hours as compared to 2 hours for bovine insulin (31,27,4). Hypersensitivity reactions to p-insulin are absent in patients previously treated with animal insulin, but studies on juvenile diabetic patients, the peak effect was observed after 4-8 hours as compared to 2 hours for bovine insulin (31,27,4). Hypersensitivity reactions to p-insulin are absent in patients previously treated with animal insulin, but studies on juvenile diabetic patients, the peak effect was observed after 4-8 hours as compared to 2 hours for bovine insulin (31,27,4). Hypersensitivity reactions to p-insulin are absent in patients previously treated with animal insulin, but studies on juvenile diabetic patients, the peak effect was observed after 4-8 hours as compared to 2 hours for bovine insulin (31,27,4).

**Other biologically active compounds**

The immature fruit of *Momordica charantia* tastes bitter, due to the cucurbitacins found in it. Cucurbitacins comprise a group of triterpenes distributed in the cucurbitaceous plants. Five non-bitter cucurbitacins have been isolated from the seeds and are designated momordicosides A-E (32,33). The immature fruit of this plant has been the source of two bitter glycosides, momordicosides K and L (34) and four non-bitter glycosides, momordicosides F₁, F₂, G and I (35). The leaves and vines contain three bitter principles which are glycosides named momordicins I, II, and III. These momordicins are very similar in structure to momordicosides K and L (35).

Two seed storage proteins have been isolated from the seeds of *M. charantia*. These proteins contain high concentrations of glutamic acid, glutamine and arginine. The proteins appear to consist of four sub-units of 5,500 daltons (36).

The seeds are also the source of vicine, a toxin, which is a glycoside of a non-protein nitrogenous base. This toxin induces fawnism, an acute condition characterized by fever, headaches, abdominal pain, severe anemia, prostration and coma (37).

Tomita et al. (38) originally purified galactose binding lectins from *M. charantia* seeds by affinity chromatography using a Sepharose 4B column. Linn et al (39), isolated and partially characterized a toxic and non-toxic lectin from the seeds of the bitter melon. The purified toxic lectin, momordin, has a molecular weight of 24,000 and inhibits protein biosynthesis in Ehrlich ascites tumor cells. The non-toxic lectin, momordica agglutinin, has a molecular weight of 32,000 and is able to agglutinate the human O type red blood cells at a concentration of 0.5 g/ml. The hemagglutination is inhibited by galactose and its derivatives as reported by Tomita et al. (38). The amino acid compositions of momordin and momordica agglutinin are different (39).

Further studies (40,41) of the hemagglutinating lectin indicate that it is a glyco-protein containing 10% hexoses and 2.5% hexosamine. The lectin has a molecular weight of 120,000 and is tetrameric in nature, containing two pairs of non-identical sub-units of molecular weights 28,000 and 30,000. The lectin has two binding sites per molecule which bind galactose and related sugars independently of each other. There do not appear to be any additional low affinity receptors (40). Chemical modification studies found that tryptophan and tyrosine residues are important for the carbohydrate binding activity (41).

An ethyl ether extract of *M. charantia* seeds is found to contain two inhibitors of protein synthesis in a cell-free system (a lysate of rabbit reticulocytes) (42,43). The inhibitors appear to act through an inhibition of peptide chain-elongation rather than through an inhibition of initiation as the extent of protein inhibition is the same when the extract is added before or well after starting the reaction (44). One inhibitor of protein synthesis is also a hemagglutinating lectin. The lectin has a molecular weight of 115,000 and consists of four sub-units (43). The lectin is non-mitogenic to human lymphocytes (45). The second protein, a basic glycoprotein (pI >8) (46), has a molecular weight of 31,000 (46) and has no hemagglutinating properties (43). This second protein is called *Momordica charantia* inhibitor (MCI) and is a ribosome-inactivating protein (47). Several plants contain inhibitors of protein synthesis which inhibit the 60 S sub-unit of eukaryotic ribosomes (45,44) and inhibit the replication of plant viruses (45). These ribosome inactivating proteins affect mammalian ribosomes more than plant ribosomes and have no effect on homologous ribosomes (45). Abrin and ricin, two such inhibitors, are toxic to animals or cells in vitro, while many other inhibitors have low or no toxicity (44).

Both the *M. charantia* lectin and the inhibitor are capable of inhibiting protein and subsequently DNA synthesis in mitogen-stimulated, normal, human, peripheral blood lymphocytes and human leukemic lymphocytes (47). The inhibitor has a low toxicity (rats tolerated 100 mg/100 g body weight), while the hemagglutinating lectin has a more rapid and pronounced effect on cellular protein synthesis. This difference is probably due to better penetration of the lectin into cells (42,43,47). Although the *M. charantia* inhibitor contains sugar residues (mannose, glucose, xylose, fucose and glucosamine), these residues may not be necessary for the biological activity of the protein. The pokeweed antiviral protein isolated from the seeds of the pokeweed contained no detectable sugars and yet is a potent inhibitor of protein synthesis (48,46). The inhibitor contains some thiol groups which would allow it to be linked to antibodies for possible use as an immunotoxin (46).

The 31,000 dalton protein synthesis inhibitor from *M. charantia* exhibits antiviral activity in virally infected cells similar to other ribosome-inactivating proteins such as pokeweed antiviral protein (PAP), wheat germ inhibitors, gelonin and dianthins. Treatment of virally infected cells with 200 g/ml of *M. charantia* inhibitor reduced the viral yield of polio virus infected cells by 98%. This reduction in virus is not due to inhibition of virus penetration in cells, but due to inhibition of protein synthesis in cells. The inhibitor decreased herpes simplex virus-1 plaque forming efficiency (the plaques were fewer and smaller). Protein synthesis was found to be inhibited more in virally infected cells than in uninected cells (44).

Recently the 31,000 dalton protein, called MAP 30, has been purified and a partial sequence obtained. This protein has a sequence similar to ricin A chain but is not identical. This protein exhibited a dose-dependent inhibition of HIV-1 infection of H9 cells (50).

A crude aqueous extract of the fruit of the bitter melon contains many proteins and many biochemical activities. The ripe fruit and leaves of the bitter melon are the source of a guanylate cyclase inhibitor (51,52). This inhibitor is not a lipid as the inhibitory activity remained in the aqueous phase after a double extraction with ether: ethanol (4:1) followed by an extraction with chloroform:methanol (2:1) (52). The guanylate cyclase inhibitor is acid stable and heat labile. This partially purified inhibitor has the ability to decrease basal levels of guanylate cyclase and to impair chemical carcinogen-induced increases in guanylate cyclase activity. The guanylate cyclase inhibitor has no effect on adenylate cyclase (52).

As guanylate cyclase activity is found to be increased in many tumor cells and is associated with concanavalin A (Con A) stimulated lymphocytes (53), the guanylate cyclase inhibitor has been tested against tumor cells and mitogen-treated lymphocytes. The aqueous extract of the bitter melon ripe fruit and leaves is found to block the growth of rat prostatic adenocarcinoma in vivo, and inhibit the incorporation of 3H-thymidine into DNA of Con A-stimulated cells. DNA
histograms indicate that the extractinhibitsatthe\text{6}^+ M phase of the cell cycle (54,55). In low amounts, this material also inhibits the self associating reaction of bovine tubulin and prevents the polymerization of chick tubulin in a manner similar to colchicine (56,55). The crude extract of the fruit of the bitter melon is preferentially cytotoxic for leukemic cells versus normal lymphocytes. This cytotoxicity correlates with the guanylate cyclase inhibitory activity (54,55,51). Testing of nine other plant extracts that are cytotoxic to cells revealed that only an extract from the Lawson’s cypress, \textit{Chamaecyparis laevisianna}, exhibits comparable cytotoxicity and guanylate cyclase inhibition levels (51).

Partial purification of a guanylate cyclase inhibitor finds that it co-elutes as a single, trypsin sensitive peak with amolecular weight of 70,000. This purified protein exhibits--preferential cytotoxicity to several human lymphoblastoid cell lines (57).

A second substance (58) can be purified with a molecular weight of 40,000 that is cytostatic to BHK-21 cells but has no effect on cellular cyclic GMP. The 40K molecule is sensitive to trypsin and to boiling and is able to inhibit RNA and protein synthesis in intact tissue culture cells and cell free systems. This cytostatic molecule also exhibits non-specific antiviral activity.

Another cytostatic molecule (59) which preferentially inhibits RNA synthesis in intact cells can also be purified from the crude extract. This molecule has an apparent molecular weight of 11,000 and is not sensitive to boiling or to treatments with trypsin ribonuclease, or deoxyribonuclease. This molecule has no effect on guanylate cyclase nor phosphodiesterase.

As stated above, a crude extract of the fruit of the bitter melon can preferentially inhibit \textit{in vitro} tumor cell growth (54,47,57,51). Jilka et al. (60) have shown that this crude aqueous extract also exhibits \textit{in vivo} antitumor activity. The extract is able to inhibit tumor formation in CBA/H mice injected with L1210 or CBA/DI tumor cells and tumor formation in DBA/2 mice injected with L1210 or P388 tumor cells. This \textit{in vivo} antitumor activity requires both the prior exposure of tumor cells to the extract \textit{in vitro} and intraperitoneal injection of 8 g protein extract twice a week.

Characterization of this anti-lymphoma moiety indicates that it is sensitive to treatment with trypsin and boiling, but insensitive to DNase or RNase. On SDS-PAGE, this moiety has a molecular weight of 40,000 (Takemoto, unpublished data).

Splenocytes of animals injected twice a week with 8mg protein of the bitter melon extract exhibited an enhanced mixed lymphocyte reaction, while thymocytes of these animals exhibited an enhanced incorpora-tion of \textit{H}--thymidine into trichloroacetic acid-precipitable material after 48 hours of exposure to Con A (60). Resistance to L1210 lymphoma formation in DBA/2 mice can be adoptively transferred by injection of peritoneal exudate cells, twice a week, from DBA12 mice which have been injected with the bitter melon antitumor factorcrakemoto, unpublished data). Therefore, some of the antitumor activity of the bitter melon extract may be due to immunopotentiataion (59).

Sprafico et al. (61) studied some of the immunomodulatory characteristics of the 30,000 dalton \textit{Momordica charantia} inhibitor. They found that \textit{in vitro} incubation of MCI with splenocytes decreased the re-sponsiveness to the T cell mitogens, PHA and Con A, but not the B cell mitogen, LPS. \textit{In vitro or in vivo} treatment of splenic or peritoneal macrophages with MCI is able to augment their antigenic response (as measured by a plaque forming cell assay) to T dependent antigens while not affecting the T independent antigens. This group concluded that the MCI has no effect on the B cell arm of the immune response, but may exert its effect by inhibition of T-regulatory cells (61).

Recently, we have identified the affected immune cells as NK cells (62). These studies suggested that an \textit{in vivo} anti-tumor effect is not due entirely to ribosomal inactivating activity. Rather, there are endogenous activators of the host’s immune system.

It should be noted that the bitter melon extract is not toxic. In view of the many reports of its antitumor and anti-HIV activity, this plant should be further studied (62).

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Studies on the constituents of Momordica charantia L. Isolation and obtained from plant source

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- Licastro, F., C. Franceschi, L. Barbieri, and F. Stirpe. 1980.