

## Role Of Krishna Dhattura (Datura Metel Linn.) As An Ayurvedic Contraceptive

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### ABSTRACT

**Background:** Krishna Dhattura (*Datura metel* Linn., Solanaceae), known in Sanskrit as Dhustura and colloquially as Black Datura or Devil's Trumpet, is one of the most pharmacologically potent plants described in the Ayurvedic Materia Medica. Classical texts classify it under Upavisha (sub-toxic substances) and document its applications in diverse therapeutic contexts including neurological disorders, pain management, skin diseases, and reproductive medicine. Among its traditional uses, its role as a contraceptive agent — termed Garbha Nirodha or Santati Niyamana in Ayurvedic literature — has been documented in regional folk medicine and select classical references, though it remains underexplored in systematic biomedical review.

**Objective:** To conduct a comprehensive review of classical Ayurvedic and ethnobotanical references to Krishna Dhattura as a contraceptive agent; to analyze its phytochemical constituents with documented antifertility activity; to correlate these with modern reproductive pharmacology; and to critically appraise its safety, toxicity, and potential for future drug development.

**Methods:** Classical Ayurvedic texts (Charaka Samhita, Sushruta Samhita, Ashtanga Hridayam, Dhanvantari Nighantu, Raja Nighantu, Bhavaprakasha Nighantu, and Sharangdhara Samhita) were reviewed alongside ethnobotanical databases and peer-reviewed literature from PubMed, Scopus, and Web of Science (1970-2025) on *Datura metel* reproductive pharmacology, alkaloid biochemistry, and antifertility studies.

**Results:** *Datura metel* contains a well-characterized alkaloid profile — predominantly scopolamine (hyoscyne), atropine (dl-hyoscyamine), hyoscyamine, and daturine — with demonstrated effects on the hypothalamic-pituitary-gonadal axis, ovulation, sperm motility, implantation, and early embryonic development in animal models. Classical Ayurvedic formulations using Dhustura seeds and root in micro-doses describe both temporary and potentially permanent antifertility effects, corresponding with modern pharmacological evidence.

**Conclusion:** Krishna Dhattura presents a biologically plausible mechanism for contraceptive action through its tropane alkaloid constituents. However, the narrow therapeutic index, significant anticholinergic toxicity, and irreversibility concerns make it unsuitable for unsupervised use. Standardized, sub-toxic formulations merit controlled experimental and clinical investigation within an ethical biomedical framework.

**Keywords:** Krishna Dhattura, *Datura metel*, Dhustura, Ayurvedic contraception, Garbha Nirodha, Tropane alkaloids, Scopolamine, Antifertility, Phytochemistry, Reproductive pharmacology.

### INTRODUCTION

Contraception — the deliberate prevention of conception — has been practiced across civilizations for millennia, and Ayurveda is no exception. The classical Ayurvedic literature, particularly the Nighantu texts (pharmacopoeial compilations) and regional folk medicine traditions, documents a rich repertoire of plant-based contraceptive agents termed Garbha Nirodha Dravya (substances that prevent conception) or Santati Niyamana Aushadhi (medicines for reproductive regulation). These range from mild emmenagogues to potent anti-implantation agents and include substances applied locally, administered orally, or used in Nasya (nasal therapy) preparations.

Among these, Krishna Dhattura — the dark-flowered or black variety of *Datura metel* Linn. (Family: Solanaceae), known in Sanskrit as Dhustura, Unmatta (one that causes intoxication), Shivapriya (dear to Shiva), and Kanaka (golden) — occupies a unique and complex position. On one hand, it is classified as an Upavisha (sub-poison or secondary toxic substance) in texts like Ashtanga Hridayam and Sharangdhara Samhita, with documented toxicity including hallucinations, mydriasis, tachycardia, and even death at higher doses. On the other hand, in precise micro-dosing formulations supervised by qualified Vaidyas, it has been used therapeutically for conditions ranging from asthma and neurological disorders to reproductive regulation.

The plant itself is easily identifiable: a robust annual or perennial herb growing to 1 meter in height, with large ovate leaves, characteristic trumpet-shaped flowers that are violet-purple in the Krishna variety, and spiny capsular fruits containing numerous reniform seeds. It is widely distributed across tropical and subtropical regions of Asia, Africa, and the Americas, with the Indian subcontinent being a primary centre of both botanical diversity and traditional utilization.

This review aims to comprehensively examine the botanical identity, classical Ayurvedic descriptions, phytochemical profile, and reproductive pharmacological evidence for Krishna Dhatura as a contraceptive agent, while critically appraising its safety and potential for future pharmaceutical development..

## 2. BOTANICAL IDENTITY AND CLASSICAL NOMENCLATURE

### 2.1 Taxonomy and Botanical Description

Taxonomic Rank	Classification
Kingdom	Plantae
Order	Solanales
Family	Solanaceae
Genus	Datura
Species	Datura metel Linn. (syn. D. fastuosa, D. alba)
Sanskrit Name	Dhustura, Unmatta, Shivapriya, Kanaka, Matula, Tutura
Hindi Names	Dhattura, Dhatura, Kala Dhatura (black variety)
Regional Names	Ummatta (Kannada/Telugu), Vellai Oommathai (Tamil), Dhaturu (Gujarati)
English Names	Devil's Trumpet, Black Datura, Horn of Plenty, Downy Thorn Apple
Parts Used	Seeds (primary), Leaves, Root bark, Flowers, Fruit rind

The genus *Datura* comprises approximately 9-12 species globally, of which *Datura metel* (syn. *D. fastuosa*) and *Datura stramonium* are the most pharmacologically and toxicologically studied. The Krishna (black/dark purple-flowered) variety of *Datura metel* is specifically distinguished in Ayurvedic texts as having more potent properties compared to the white-flowered (Shveta Dhustura) variety. This distinction likely reflects real phytochemical differences in alkaloid concentration and ratios between the two varieties, though systematic comparative studies remain limited.

### 2.2 Classical Ayurvedic Description

The earliest detailed description of Dhustura appears in Charaka Samhita (Chikitsa 23/85-87) in the context of anti-poisoning formulations and neurological treatments. Sushruta Samhita (Kalpa 7/8-12) provides detailed descriptions of its toxic properties and therapeutic applications in Panchakarma adjuvant procedures. Ashtanga Hridayam (Uttarasthana 35) classifies it among Upavisha dravyas.

The Nighantu texts provide the most comprehensive pharmacognostic descriptions. Bhavaprakasha Nighantu (Guduchyadi Varga) describes Dhustura as Tikta (bitter) and Kashaya (astringent) in taste, Ushna (hot) in potency (Virya), Laghu (light) and Ruksha (dry) in qualities (Guna), with Vata and Kapha pacifying effects and a tendency to aggravate Pitta in excess. Its Prabhava (special action) is described as Unmadakara (intoxicating) and Vedanasthapana (pain-relieving). Dhanvantari Nighantu specifically mentions its use in Garbha Nirodha (contraceptive) applications in seed form.

## 3. PHYTOCHEMICAL COMPOSITION

### 3.1 Major Alkaloid Profile

The pharmacological activity of *Datura metel* is primarily attributable to its tropane alkaloid content, which varies significantly by plant part, geographic origin, season of harvest, and variety. The major alkaloids identified

include:

Alkaloid	Chemical Class	Concentration (Seeds)	Primary Pharmacological Action
Scopolamine (Hyoscine)	Tropane	0.2-0.45%	Anticholinergic, CNS depressant, antiemetic, antisecretory
Hyoscyamine (l-form)	Tropane	0.1-0.3%	Muscarinic antagonist, antispasmodic, mydriatic
Atropine (dl-Hyoscyamine)	Tropane	0.05-0.2%	Anticholinergic, increases heart rate, mydriatic
Daturine	Tropane	Minor	Anticholinergic, sedative
Meteloidine	Tropane	Minor	Estrogenic-like activity reported
Noratropine	Tropane	Trace	Anticholinergic
Withanolides (root)	Steroidal lactone	Variable	Adaptogenic, anti-inflammatory, potential hormonal modulation

### 3.2 Non-Alkaloid Constituents

Beyond tropane alkaloids, *Datura metel* contains a range of secondary metabolites with documented biological activity. Flavonoids including luteolin, apigenin, and quercetin derivatives have been isolated from leaves and flowers. Withanolides — steroidal lactones structurally related to those in *Withania somnifera* — have been identified particularly in the root and demonstrate immunomodulatory and potential endocrine-disrupting effects. The seeds additionally contain fixed oils (predominantly oleic, linoleic, and stearic acid esters), saponins, and tannins. The steroidal content of *Datura metel* root has been proposed as a basis for its estrogenic-like effects observed in some animal studies.

The seeds of Krishna Dhatura (*Datura metel* var. *fastuosa*) have been reported to contain higher concentrations of scopolamine relative to atropine compared to the white-flowered variety, a distinction with practical pharmacological significance given scopolamine's documented effects on reproductive function at sub-toxic doses.

## 4. CLASSICAL AYURVEDIC REFERENCES TO CONTRACEPTIVE USE

### 4.1 Textual Evidence

Direct references to Dhustura as a contraceptive agent are found across several classical and medieval Ayurvedic sources, though the descriptions are often brief, encoded in symbolism, or embedded within broader therapeutic contexts. The following references merit detailed examination:

Charaka Samhita (Chikitsa 30/95-100) contains a section on Santati Niyamana (reproductive regulation) that lists several plant-based agents. Among oral preparations, Dhustura seed powder processed with specific anupanas (vehicle substances) is mentioned as producing temporary cessation of ovarian function (Artava Kshaya) when used in precise doses under physician supervision.

Sharangdhara Samhita (Uttara Khanda 10/15-19) provides the most specific antifertility formulation involving Dhustura. A preparation described as a combination of Dhustura seed (Krishna variety) powder with Tankana (borax), Palasha (*Butea monosperma*) seed, and Pippali (*Piper longum*) root, administered in a dose of one Ratti (approximately 125 mg of the combined preparation) per day on a specific schedule relative to the menstrual cycle, is described as producing contraceptive effect lasting one year. The text explicitly warns against use in pregnancy or lactation and specifies that the preparation should be discontinued if pregnancy is desired.

Raja Nighantu (Shatapushpadi Varga) lists Dhustura among Garbha Nashaka dravyas (substances causing abortion/preventing implantation) and specifically notes the seeds of the Krishna variety as more potent. This classification as Garbha Nashaka rather than Garbha Nirodhaka (preventive of conception) suggests that some classical authorities considered its primary action to be at the implantation or early embryonic stage rather than at ovulation or sperm function.

Bhaisajya Ratnavali and several regional folk medicine compilations from Maharashtra, Rajasthan, and Bengal document the use of Krishna Dhatura seed in medicated butter (Navanita) preparations for temporary infertility, with the folk tradition specifying that one seed per month administered at a specific phase of the cycle produces contraception without permanent sterility.

#### 4.2 Route of Administration

Classical references describe both oral (with milk, ghee, or specific herbal decoctions as anupana) and local (intravaginal, in medicated oil preparations) routes of administration. The local route preparations — described in Visha Chikitsa chapters of Sushruta Samhita as using Dhustura leaf extract in sesame oil — may correspond to spermicidal or cervical mucus-altering mechanisms rather than systemic hormonal effects. The oral route preparations, in contrast, are more likely to produce systemic anticholinergic and potential hormonal effects.

Notably, Ashtanga Hridayam (Uttarasthana 35/49-52) cautions that even the smoke from burning Dhustura leaves, when inhaled, can produce Garbha Pata (abortion) in pregnant women — a warning that aligns with modern knowledge of scopolamine's abortifacient potential at higher exposures and provides an additional biological plausibility dimension.

### 5. PHARMACOLOGICAL MECHANISMS OF CONTRACEPTIVE ACTION

#### 5.1 Effects on the Hypothalamic-Pituitary-Gonadal (HPG) Axis

The tropane alkaloids of *Datura metel* — particularly scopolamine and hyoscyamine — are potent antagonists of muscarinic acetylcholine receptors (mAChRs). Muscarinic receptors (M1-M5 subtypes) are expressed throughout the hypothalamus, anterior pituitary, and gonadal tissues. Muscarinic signaling plays a well-established role in regulating GnRH (Gonadotropin-Releasing Hormone) pulsatility from the hypothalamic pulse generator.

Studies using scopolamine in animal models have demonstrated dose-dependent suppression of GnRH pulse frequency, with consequent reduction in LH (Luteinizing Hormone) and FSH (Follicle-Stimulating Hormone) secretion. In female rats, chronic administration of scopolamine at sub-toxic doses produced anovulatory cycles, atrophic changes in the ovarian follicles, and elevated prolactin levels — a hormonal milieu consistent with contraceptive effect. The GnRH pulse generator in the arcuate nucleus of the hypothalamus has dense cholinergic innervation, and M1/M3 receptor blockade by tropane alkaloids provides a plausible central mechanism for HPG axis suppression.

#### 5.2 Direct Ovarian Effects

Several animal studies have demonstrated direct effects of *Datura metel* extracts on ovarian function independent of central HPG modulation. Aqueous and ethanolic seed extracts administered to female mice and rats produced dose-dependent inhibition of follicular development, reduction in antral follicle count, and decreased serum estradiol levels. Histological examination revealed atretic follicles and disruption of granulosa cell integrity — changes consistent with anti-gonadotropic and/or direct cytotoxic effects on ovarian tissue.

The withanolides present in the *Datura* root fraction have been proposed as mediators of these direct ovarian effects, given the well-documented ovarian-protective and hormonal-modulatory effects of withanolides from related species (*Withania somnifera*). Conversely, the estrogenic-like effects of meteloidine — a minor tropane alkaloid — may contribute to negative feedback suppression of FSH through estrogen receptor-mediated pathways.

#### 5.3 Anti-Implantation and Post-Coital Effects

Multiple rodent studies have evaluated *Datura metel* seed extracts for post-coital antifertility activity — a mechanism corresponding to the anti-implantation (rather than contraceptive) category. In studies using Wistar rats and Swiss albino mice, oral administration of petroleum ether, chloroform, and aqueous seed extracts at doses of 200-400 mg/kg body weight on days 1-7 post-coitum produced significant reduction in implantation sites (50-90% reduction in different studies) and increased resorption rates.

The proposed mechanisms include: (1) Disruption of the uterine receptivity window (days 4-5 in rodents) through anticholinergic inhibition of uterine muscarinic receptors that regulate endometrial stromal cell decidualization; (2) Anti-estrogenic or anti-progestogenic effects on the endometrium preventing the specific molecular cascade required for blastocyst implantation (including LIF, HOXA10, integrin  $\alpha$ v $\beta$ 3 expression); and (3) Direct embryotoxic effects on the pre-implantation blastocyst.

#### 5.4 Effects on Male Reproductive Function

The antifertility effects of *Datura metel* are not limited to female reproductive physiology. Studies in male rodents have demonstrated that both the whole plant extract and isolated alkaloid fractions produce dose-dependent impairment of spermatogenesis. Specifically, oral administration of *Datura metel* seed extract for 60 days in male rats produced significant reductions in testicular sperm count and epididymal sperm motility, increased sperm

morphological abnormalities, and histological evidence of seminiferous tubule disruption including vacuolation of Sertoli cells and arrest of spermatogenesis at the primary spermatocyte stage.

The mechanisms proposed include: (1) Anticholinergic suppression of cholinergic innervation of the testis and epididymis, which is important for sperm maturation and motility; (2) Oxidative stress induction in testicular tissue; and (3) Disruption of the blood-testis barrier. Critically, at higher doses, these effects appeared irreversible in some studies — a toxicity concern that must inform any clinical application.

### 5.5 Spermicidal Activity

In vitro studies have demonstrated direct spermicidal activity of *Datura metel* leaf and seed extracts. Aqueous extracts at concentrations of 1–5 mg/mL produced complete motility arrest and morphological disruption of human spermatozoa within 30–60 seconds of exposure in in vitro assays. The alkaloid fraction showed the highest spermicidal potency, with scopolamine demonstrating direct membrane-disrupting effects on sperm plasma membranes. These findings provide pharmacological basis for the classical Ayurvedic local formulations using Dhustura-based medicated oils as vaginal preparations.

Scopolamine has additionally been shown to inhibit the acrosome reaction — the essential step in sperm penetration of the zona pellucida — through its anticholinergic effects on sperm M1 receptors, which regulate intracellular calcium signaling in the acrosomal pathway. This mechanism would prevent fertilization even in the presence of viable sperm reaching the secondary oocyte.

## 6. TOXICOLOGY AND SAFETY PROFILE

### 6.1 Acute Toxicity

*Datura metel* is classified as a Schedule E1 poison under the Drugs and Cosmetics Act in India, reflecting its significant toxicity potential. The toxic syndrome associated with *Datura* alkaloids is classically described as anticholinergic toxidrome: dry mouth (xerostomia), blurred vision (cycloplegia and mydriasis), tachycardia, hyperthermia, urinary retention, decreased bowel sounds, flushed skin, and — most dangerously — CNS effects ranging from agitation and confusion to hallucinations, seizures, and coma.

The LD50 values reported for *Datura metel* seed extract in rodent models range from 400–1000 mg/kg (oral) depending on the extraction solvent and species. The therapeutic-to-toxic ratio is extremely narrow, with antifertility effects documented at 200–400 mg/kg in rodents and toxic effects beginning at similar or only marginally higher doses. This narrow therapeutic index represents the most significant obstacle to clinical application of *Datura*-based contraceptive formulations.

### 6.2 Chronic Toxicity

Subchronic and chronic toxicity studies in rodents have documented hepatotoxicity, nephrotoxicity, and gonadotoxicity at doses sufficient to produce antifertility effects. Liver histopathology revealed periportal inflammation and hepatocyte vacuolation; renal histopathology showed tubular epithelial degeneration. These organ toxicity findings present serious safety concerns for any proposed clinical use. The reversibility of antifertility effects — a critical parameter for any contraceptive agent — has been inconsistently reported across studies, with some demonstrating recovery of spermatogenesis and ovarian function after discontinuation and others showing persistent changes.

### 6.3 Teratogenicity and Embryotoxicity

*Datura metel* alkaloids have demonstrated significant embryotoxic and teratogenic potential in animal studies. Administration during organogenesis in rats produced neural tube defects, limb malformations, and increased resorption rates. These teratogenic effects are particularly concerning given that the plant's anti-implantation effects may be incomplete, and exposure of a continuing pregnancy to *Datura* alkaloids would carry substantial risk of fetal malformation. Any contraceptive application must therefore ensure reliable prevention of continuation of pregnancy, or be explicitly categorized as abortifacient rather than contraceptive.

### 6.4 The Upavisha (Sub-Poison) Framework in Ayurveda

The Ayurvedic classification of Dhustura as Upavisha is itself a sophisticated pharmacological framework that deserves recognition. Unlike Maha Visha (major poisons) where therapeutic use is essentially impossible, Upavisha substances are recognized as having genuine therapeutic value at carefully titrated sub-toxic doses following appropriate Shodhana (purification) processing. The Shodhana of Dhustura seeds — described in Sharangdhara Samhita and Rasashastra texts — involves processing with cow's milk (Dugdha Shodhana) or Triphala decoction to reduce alkaloid content while retaining therapeutic activity. The processed (Shodhita) seeds are considered substantially safer than unprocessed (Ashodhita) seeds, though rigorous chemical quantification of alkaloid reduction post-Shodhana is lacking in the literature.

7. CORRELATION: CLASSICAL CLAIMS AND MODERN EVIDENCE

Classical Claim	Proposed Mechanism (Modern)	Experimental Evidence	Evidence Level
<b>Garbha Nirodhana (prevents conception)</b>	HPG axis suppression via mAChR blockade; spermicidal activity; anti-implantation	Animal studies (rodents); in vitro spermicidal assays	Level III-IV (Animal/In vitro)
<b>Artava Kshaya (ovarian suppression)</b>	Reduced GnRH pulsatility; direct follicular atresia	Rodent studies showing anovulation, follicular atresia	Level III (Animal)
<b>Garbha Nashana (anti-implantation)</b>	Disruption of uterine receptivity; anti-decidualization	Post-coital rodent studies: 50-90% implantation reduction	Level III (Animal)
<b>Shukra Dushti (sperm impairment)</b>	Direct spermicidal; inhibition of acrosome reaction; cholinergic impairment of epididymal maturation	In vitro spermicidal assays; chronic dosing rodent spermatogenesis studies	Level III-IV (Animal/In vitro)
<b>Reversible effect with proper dose</b>	Dose-dependent and time-dependent recovery of reproductive function	Inconsistent; some studies show recovery, others do not	Level IV (Conflicting animal data)
<b>Shodhana reduces toxicity</b>	Alkaloid reduction through heat/milk processing	Limited chemical analyses of processed vs. unprocessed seeds	Level V (Preclinical/Descriptive)

8. COMPARATIVE ANALYSIS WITH OTHER AYURVEDIC CONTRACEPTIVE PLANTS

Krishna Dhattura does not exist in isolation within the Ayurvedic antifertility pharmacopoeia. Several other plants described in classical texts as Garbha Nirodha dravya have also been subjected to modern pharmacological investigation, and a comparative analysis contextualizes Datura metel's profile:

Plant (Sanskrit)	Botanical Name	Primary Antifertility Mechanism	Evidence Level	Toxicity Profile
<b>Dhustura (Krishna)</b>	Datura metel	HPG suppression, anti-implantation, spermicidal	Animal/In vitro	HIGH — narrow therapeutic index, teratogenic
<b>Apamarga</b>	Achyranthes aspera	Anti-implantation, abortifacient (saponins)	Animal/In vitro	Moderate — saponin toxicity
<b>Chitraka</b>	Plumbago zeylanica	Plumbagin: anti-implantation, embryotoxic	Animal	Moderate-High — hepatotoxic
<b>Palasha</b>	Butea monosperma	Phytoestrogenic anti-implantation; anti-spermatogenic	Animal/In vitro	Low-Moderate — reversible effects
<b>Indravriksha</b>	Wrightia tinctoria	Androgen disruption,	Animal	Moderate

Plant (Sanskrit)	Botanical Name	Primary Antifertility Mechanism	Evidence Level	Toxicity Profile
<b>Nimba</b>	Azadirachta indica	Spermicidal (nimbodin), anti-implantation	Animal/Clinical (limited)	LOW — reversible, well-tolerated
<b>Shalmali</b>	Bombax ceiba	Estrogenic, anti-implantation	Animal	Low-Moderate

This comparative analysis reveals that while Krishna Dhatura demonstrates among the most potent antifertility activity of the classical Ayurvedic contraceptive plants, it also carries the highest toxicity risk. Neem (*Azadirachta indica*) and Palasha (*Butea monosperma*) represent far more favorable safety profiles relative to their contraceptive efficacy, and these plants have accordingly received more sustained modern research attention. The development path for *Datura metel*-based contraceptives would require substantially more work on standardization, dose optimization, and safety characterization before any clinical application could be ethically considered.

## 9. ETHNOBOTANICAL AND FOLK MEDICINE PERSPECTIVES

Ethnobotanical surveys conducted across India have documented widespread folk use of *Datura metel* for reproductive regulation, particularly in rural communities with limited access to modern contraceptive services. Survey data from Rajasthan, Madhya Pradesh, Uttar Pradesh, and Maharashtra have recorded consistent reports of Krishna Dhatura seed use for contraception and abortion among traditional communities, typically administered by local Vaidyas or dai (midwives) using preparations transmitted through oral tradition.

A notable ethnobotanical survey conducted in the Vindhya region of Madhya Pradesh documented that seeds of *Datura metel* were used in a specific preparation: one seed per lunar month (approximately equivalent to 125-150 mg of seed per month) mixed with jaggery and administered on the day after menstruation ceases. Informants claimed this produced contraception for the duration of use without permanent sterility. In a separate survey from tribal communities of Jharkhand, the root bark of Krishna Dhatura processed with cow's milk was described as a male contraceptive, administered for 30 days to produce temporary infertility in men without loss of libido.

These ethnobotanical records, while not constituting clinical evidence, serve several important research functions: they corroborate the classical textual references, suggest specific dose ranges for pharmacological investigation, provide information on traditionally safe preparation methods (Shodhana equivalents), and indicate that both male and female antifertility applications were documented in practice. The consistency across geographically separated traditional communities suggests a genuine pharmacological basis for these observations rather than purely cultural mythology.

## 10. PROSPECTS FOR PHARMACEUTICAL DEVELOPMENT

### 10.1 Isolated Compound Approach

The identification of scopolamine and hyoscyamine as the primary bioactive constituents of *Datura metel* opens the possibility of developing isolated or semi-synthetic derivatives with enhanced contraceptive selectivity and improved safety profiles. Scopolamine is already a WHO Essential Medicine (for motion sickness and postoperative nausea) with an established human pharmacokinetic and safety profile at anti-emetic doses. Sub-emetic doses of scopolamine — substantially below anti-emetic doses — may achieve the hypothalamic GnRH suppression required for contraceptive effect, warranting investigation.

Structure-activity relationship (SAR) studies on tropane alkaloid modifications could potentially yield derivatives with preferential affinity for the specific muscarinic receptor subtypes involved in GnRH regulation (primarily M1 in hypothalamic interneurons) over the M3 receptors responsible for peripheral anticholinergic side effects. Such receptor subtype selectivity would dramatically improve the therapeutic index and could make a *Datura*-derived contraceptive pharmacologically viable.

### 10.2 Standardized Plant Extract Approach

An alternative development pathway involves the preparation of standardized, quality-controlled extracts of Shodhita (processed) *Datura metel* seeds with defined alkaloid content, subjected to rigorous preclinical

toxicology studies. The traditional Shodhana processing — involving repeated trituration with cow's milk or Triphala decoction — may reduce total alkaloid content by 30-60% while preferentially retaining certain fractions. Quantitative analysis of Shodhita versus Ashodhita seed alkaloid profiles using HPLC-MS would clarify whether the traditional processing achieves meaningful reduction in the most toxic alkaloids (atropine > hyoscyamine) relative to the potentially more selectively reproductive-active fractions.

### 10.3 Research Gaps and Future Directions

Systematic dose-finding studies in established animal models using Shodhita seed extracts with quantified alkaloid content.

Comparative toxicokinetics of Shodhita versus Ashodhita preparations to validate traditional processing methods.

Mechanistic studies on HPG axis effects using ex vivo hypothalamic preparations to characterize the specific GnRH regulatory role of mAChR subtypes.

Evaluation of reversibility of antifertility effects at the lowest effective dose in long-term animal studies.

In vitro estrogenic/androgenic receptor binding assays for withanolide and meteloidine fractions.

Ethnopharmacological documentation of traditional processing methods and dose schedules for systematic pharmacological validation.

## 11. ETHICAL AND LEGAL CONSIDERATIONS

Any discussion of *Datura metel* as a contraceptive must be accompanied by rigorous ethical and legal analysis. The plant is classified under Schedule E1 of the Drugs and Cosmetics Act in India, which restricts its sale, distribution, and use to licensed medical practitioners. Promotion of unsupervised self-use of *Datura* preparations for contraception would be both legally problematic and medically dangerous.

The documented cases of *Datura* poisoning in India — where the plant is involved in a significant proportion of plant-poisoning hospitalizations, and where it has historically been used criminally to incapacitate victims — underscore the real-world risks of popularizing its use. Any research publications on *Datura*'s antifertility properties must include clear, prominent safety warnings that these findings relate to research-grade preparations under controlled conditions and that self-administration of any *Datura* preparation constitutes a significant health risk.

From a bioethical standpoint, the development of plant-based contraceptives from traditionally used herbs like *Datura metel* serves legitimate goals in expanding contraceptive options, particularly for populations with limited access to pharmaceutical contraceptives or preferences for plant-based medicines. However, this development must occur through the standard regulatory pathway of preclinical safety and efficacy studies, followed by Phase I-III clinical trials, with full informed consent and robust adverse event monitoring. The traditional knowledge base represented by Ayurvedic texts and ethnobotanical records should be respected as prior art and compensated appropriately under frameworks addressing bioprospecting and traditional knowledge protection.

## 12. DISCUSSION

This review has assembled a comprehensive body of evidence supporting a biologically plausible basis for the Ayurvedic use of Krishna Dhatura (*Datura metel*) as a contraceptive agent. The convergence of classical textual references across multiple independent Ayurvedic traditions, ethnobotanical corroboration from diverse geographic regions, and pharmacological evidence from animal and in vitro studies collectively suggests that this is not merely cultural belief but reflects genuine empirical observation of reproductive effects accumulated over centuries of documented use.

The primary mechanism of contraceptive action appears to be multi-modal: central HPG axis suppression through muscarinic receptor blockade of GnRH pulsatility, direct ovarian follicular effects, disruption of uterine implantation receptivity, and spermicidal activity. This multi-modal profile, while pharmacologically interesting, also complicates dose optimization, as different mechanisms may operate at different dose thresholds with different toxicity profiles.

The critical limitation — and the factor that most significantly constrains translational potential — is the narrow therapeutic index inherent to tropane alkaloid pharmacology. The doses required to produce demonstrable antifertility effects in rodents (200-400 mg/kg) are uncomfortably close to doses producing overt anticholinergic toxicity (400-1000 mg/kg), leaving limited margin for safe clinical application of whole-plant preparations. This is precisely the problem that motivated the traditional Shodhana processing requirement described in classical texts — the recognition that therapeutic benefit was achievable only within a narrow dose window that required both expert preparation and individualized dosing by a qualified Vaidya.

Modern pharmaceutical science has the tools to address this challenge: standardized extraction, alkaloid

quantification, receptor subtype-selective modification, and controlled-release delivery systems could collectively expand the therapeutic window to clinically useful proportions. The isolated compound approach — developing scopolamine-derived or hyoscyamine-derived analogues with selective GnRH-regulatory receptor affinity — represents the most scientifically tractable development pathway.

### 13. CONCLUSION

Krishna Dhatura (*Datura metel* Linn.) presents a pharmacologically compelling case as a plant-based contraceptive agent with historical documentation spanning two millennia of Ayurvedic and folk medicine practice. Its tropane alkaloid constituents — principally scopolamine and hyoscyamine — demonstrate multi-modal antifertility activity through central HPG axis modulation, direct gonadal effects, anti-implantation mechanisms, and spermicidal activity across multiple experimental models.

However, the significant toxicity profile, narrow therapeutic index, potential irreversibility of antifertility effects at higher doses, and teratogenic risk collectively make unsupervised clinical use of *Datura metel* preparations for contraception inadvisable in the current state of knowledge. The classical Ayurvedic framework — which recognized these risks through the Upavisha classification and mandatory Shodhana processing — embedded appropriate safeguards that modern application must likewise adopt and rigorously validate.

The most productive path forward is through systematic pharmaceutical research: isolation and structural modification of bioactive alkaloids for receptor subtype selectivity, rigorous preclinical characterization of Shodhita preparations, and eventually, ethically conducted clinical trials under regulatory oversight. The traditional knowledge encoded in Ayurvedic texts and ethnobotanical records provides an invaluable compass for this research, pointing toward specific dose ranges, preparation methods, and clinical applications that have withstood empirical testing across generations of traditional practice.

Krishna Dhatura remains, for now, a plant of remarkable pharmacological promise and equally remarkable danger — a fitting embodiment of the Ayurvedic aphorism that there is no substance in nature that is not a medicine, and no medicine that is not a potential poison. The wisdom lies entirely in the dose, the preparation, and the practitioner.

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