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# Phytochemical, ethnopharmacological and medicinal importance of *Catharanthus roseus* (Apocyanaceae): A mini review

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## ABSTRACT

From the ancient period, nature has always blessed human beings with a variety of medicinal plants having no or less side effects. *Catharanthus roseus* (Apocyanaceae) is a well-known ayurvedic herb that can be found in vast quantities all around the world. The common name for *C. roseus* are periwinkle, pink periwinkle, Madagascar periwinkle, old maid, graveyard plant, cape periwinkle, and bright eyes. The present research has aimed to review the presence of phytochemicals, ethnopharmacological, traditional, and important medicinal information on *C.roseus*.This plant has traditionally been used to treat muscle discomfort, central nervous system depression, and wasp stings. Recent research has revealed that *C. roseus*contains more than 70 different types of alkaloids (indole alkaloids), saponins, flavonoids, carbohydrates, and chemotherapeutic agents that are effective in treating life-threatening diseases like cancers. The phytoconstituents present in *C. roseus* exhibited antidiabetic, anticancer, antiulcer, antioxidant, antibacterial, and other pharmacological effects in different *invitro* and *in vivo* studies. Vinca alkaloids (vincristine and vinblastine) are one of the prominent phytoconstituents present in this plant which highly contribute to cancer chemotherapy. This minireview will provide valid and crucial information about *C. roseus*to future research in natural drug discovery processes and also for clinical researchers to find vital and effective medication to treat life-threatening conditions.

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*Keywords: Catharanthus roseus; phytoconstituents; medicinal use; pharmacological activity; Vinblastine; Vincristine.*

15 **1. INTRODUCTION**

16

17 **The medicinal plant** has great significance in humans' lives as they are composed of  
18 chemical compounds **that combat sufferings** from dawn to civilization [1]. Traditional  
19 medicines have been serving a large portion of the population for millennia. They  
20 have an abundant impact on developing most of the modern synthetic medicines. A  
21 lot of traditional medicines are now under scientific assessment due to their  
22 bioactive efficacy and safety [2]. In the initial stages, medicinal herbs were mostly  
23 preferred for treatment rather than disease prevention. Recently immense research  
24 has been done on herbal plants and their bioactive constituents that can be  
25 employed as alternative therapeutic instruments to prevent or treat a variety of  
26 infectious diseases. *Catharanthus* is a genus like a vinca, commonly known as  
27 periwinkle. Among eight known species, seven are endemic to Madagascar.  
28 Scottish botanist George Don determined the botanical nomenclature of  
29 ***Catharanthus roseus* (L)(*C. roseus*).** *C. roseus* (L.), an essential medicinal plant in  
30 the Apocynaceae family, is used to treat a wide range of fatal conditions. The plant  
31 has a long history of usage in Ayurvedic medicine and traditional Chinese medicine,  
32 but it was only in the twentieth century that Western medical science began to  
33 investigate it. It is an evergreen, bushy flowering plant, erect annual herb, native  
34 species to the Indian Ocean Island of Madagascar. It is also found in India,  
35 Australia, Southern Europe, Africa, and southern parts of the United States in  
36 California, Texas, Georgia, Florida, Mississippi, Louisiana, South, and North  
37 Carolina [3]. The word *Catharanthus* came from the Greek language which meant  
38 "pure flower" and *roseus* means rose. Thus, the Madagascan periwinkle has been  
39 given the name the "rosy" periwinkle. *C. roseus* is mainly used as an ornamental  
40 plant that **measures** 80 cm in height. There are different varieties of *C. roseus* by  
41 flower coloration, prevailing those of white color, white with a red center, purple,  
42 white with dispersing center, or with dark center bordered with red [4]. Its leaves are  
43 commonly oval to oblong, 2.5-3cm long, and 3.5cm broad, glossy green hairless  
44 with a pale midrib and a slender petiole around 1-1.8 cm long, grouped in opposing  
45 pairs [5]. Generally, plants contain two types of phytochemical constituents which  
46 contribute to the pharmacological effects of medicinal plants. These are primary  
47 metabolites and secondary metabolites. Primary plant metabolites are those that  
48 execute vital life functions found in large amounts such as carbohydrates, protein,  
49 fatty acids, nucleic acids, etc. Secondary metabolites, non-essential to life are the  
50 products of the metabolism of primary metabolites having biological effects at  
51 certain doses such as terpenes, phenolic compounds, alkaloids, etc. In the  
52 traditional and folk medicinal systems, secondary plant metabolites were used to  
53 cure several ailments. In modern medicine, they are being used as true lead  
54 compounds for drug development [6]. *C. roseus* is a medicinal plant containing a  
55 group of about 130 alkaloids, 70 of which are pharmacologically active. Different  
56 alkaloids have different medicinal effects against various diseased conditions such  
57 as hyperglycemia, **and microbial infections**, while others assist in lowering blood  
58 pressure, cancer treatment, and so on [7]. Vinblastine and vincristine are potent  
59 alkaloids present in this plant which exhibit strong anticancer properties and are now  
60 commercially used in anticancer chemotherapy. They also possess tumor inhibition  
61 activity for the treatment of leukemia, lymphosarcoma, lymphogranulomatosis, and  
62 other malignant tumors. Vinblastine alone is used to treat Hodgkin's disease,

63 advanced testicular tumors, choriocarcinoma, breast carcinoma, Kaposi  
64 carcinoma, Letterer-si disease, and leukemia in children including acute myeloid  
65 leukemia and acute lymphoblastic leukemia<sup>2</sup>. Furthermore, both vinblastine and  
66 vincristine are sold and marketed under the brand name, Velban and Oncovin [8].

67 Monoterpene indole alkaloids, ajmalicine, and serpentine have anti-inflammatory  
68 and anti-hypertensive properties. Yohimbine is effective in the treatment of erectile  
69 dysfunction[9]. Vindoline, indoline, and vindolinine showed an antidiabetic effect in  
70 vitro. Moreover, the promising fact is that recently isolated indole alkaloids of *C.*  
71 *roseus* such as catharoseumine, 17-deacetoxyvinamidine, 17-deacetoxyvinamidine,  
72 14',15'-didehydrocyclo-vinblastine exhibited *in vitro* human cancer cell  
73 line inhibition properties. These findings denote the illuminating prospects of further  
74 investigation of the *C. roseus* plant [10]. Furthermore, as there are a large number  
75 of alkaloids present in *C. roseus* and the amount of these active alkaloids is very  
76 meager, the isolation process is very costly and laborious. It has been measured  
77 that the plenty of dried periwinkle leaves (nearly 500 kg of dried leaves) is needed to  
78 isolate and purify 1g of vinblastine [11]. *C. roseus* possessed antioxidant properties  
79 due to the presence of phenolic compounds in different parts of it [12]. Aqueous  
80 extract of it has also been used for wound healing [13] as well as blood coagulation  
81 properties. It is also recognized for having properties like anti-microbial [5],  
82 antibacterial [5], and antidiabetic [7], antihypertensive [14]. So, owing to their  
83 notable pharmaceutical value and constituting a minor part of this complex mixture,  
84 *C. roseus* is considered a plant of enormous pharmaceutical interest to researchers.

85 This review paper aims to elucidate the medicinal, traditional, and pharmacological  
86 knowledge of this plant along with its new and developed treatment approach.

## 87 2. METHODOLOGY

88

89 The findings of the existing research were based on a literature review of the  
90 phytochemistry, pharmacological characteristics, and medicinal uses *C. roseus*,  
91 which was conducted utilizing data from numerous internet sources, Scopus,  
92 Google Scholar, PubMed as well as Science Direct databases. Other information  
93 sources included pre-electronic sources, journal articles, book chapters, and other  
94 scientific articles gathered from the library at the university. We used numerous  
95 keywords during the search for information from different databases. These  
96 keywords include *Catharanthus roseus* OR *Madagascar Periwinkle*, *Vinca rosea* OR  
97 traditional AND phytochemical..

## 98 3. RESULTS AND DISCUSSION

99

100 In this process of data acquisition, all the articles were assessed for eligibility to be  
101 included in this minireview. Some of the articles were removed due to duplication  
102 and others were removed for not fulfilling the inclusion criteria. However, those  
103 articles lack full text and irrelevant titles were excluded from our minireview. After  
104 downloading and organizing the 185 scientific publications for this review into a  
105 folder, we carried out the preliminary search and removed any papers that didn't  
106 include the information we needed. Following that, 67 articles were added for full-  
107 text screening. Ultimately, 73 publications that covered details about the plant's

108 traditional uses as well as phytochemical and pharmacological information were  
109 included for this study. The actions of *C.roseus*, which are described in depth in the  
110 following selections, have been used to characterize these several articles.

### 111 3.1 COMMON NAME

112  
113 *C. roseus* is commonly known as rose periwinkle, pink periwinkle, Madagascar  
114 periwinkle, old maid, graveyard plant, cape periwinkle, and bright eyes. It is known in  
115 different countries with different names such as Sadabahar in India, Nayantara in  
116 Bangladesh, Baahramaasephool in Nepal. It is also introduced by the name of  
117 Annual vinca, Running myrtle, Vinca rosea, Vinca roseus, Jasmine, Tahi ayam.

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119

120 **Fig.1. *Catharanthus roseus* (Flower)**

### 121 3.2 BOTANICAL DESCRIPTION

122 *C. roseus* is mainly used as an ornamental plant that measures 80 cm in height.  
123 There are different varieties of *C. roseus* by flower coloration, prevailing those of  
124 white color, white with a red center, purple, white with dispersing center, or with dark  
125 center bordered with red [4]. Its leaves are commonly oval to oblong, 2.5-3cm long,  
126 and 3.5cm broad, glossy green hairless with a pale midrib and a slender petiole  
127 around 1-1.8 cm long, grouped in opposing pairs [5].

### 128 3.3 ETHNO MEDICINAL KNOWLEDGE

129 *C. roseus* has been used in folk medicine to treat diabetes and high blood pressure  
130 [15], especially to treat diabetes for ages in Europe. The juice of its leaves was used  
131 to treat wasp stings in India. The herb was boiled in Hawaii to form a poultice to stop  
132 bleeding. It was used as an astringent, diuretic, and cough medicine in China. It was  
133 utilized as a home treatment for lung congestion and inflammation in Central and  
134 South America. An extract from the blossoms was used to prepare a treatment to  
135 cure eye irritation and infections throughout the Caribbean. When Western  
136 researchers learned about a tea that Jamaicans were drinking to treat diabetes in  
137 the 1950s, they became aware of this plant. They observed that it contained several  
138 alkaloids that were beneficial to humans (130 in all at last count). Some of them

139 were catharanthine, leurosinesulfate, lochnerine, tetrahydroalstonine, vindoline, and  
 140 vindolinine which reduced blood sugar levels. Besides, other alkaloids like  
 141 vincristine and vinblastine were recognized as haemostatics (stop bleeding), and  
 142 had anticancer properties. The alkaloids like reserpine and serpentine had been  
 143 recognized as potent tranquilizers, were also found in periwinkle [16]C. roseus had  
 144 been used to combat a wide variety of diseases from time immemorial which were  
 145 summarized in Table 1.

146

**Table 1. Traditional uses of C. roseus**

Preparation	Disease	Origin	Mode of administration
Hot water extract of dried leaves	Menorrhagia, Diabetes	Australia, South Africa	Oral [17, 18]
Root bark extract	Febrifuge	Australia	Oral [17, 19]
Hot water extract of dried whole plant	<i>Diabetes mellitus</i>	Brazil	Oral [17,20,21]
Hot water extract of aerial parts	Menstrual regulation	China	Oral [17, 22]
Decoction of dried leaves	Diabetes Hypertension Cancer	Cook island	Oral [17, 23]
Decoction of dried leaves	<i>Diabetes mellitus</i>	Europe	Oral [17, 24]
Hot water extract of whole plant	Anti galactagogue	France	Oral [17, 25]
a) Hot water extract of dried whole plant b) Hot water extract of dried leaves c) Root extract	Cancer Hodgkin's disease Menorrhagia	India	Oral [17, 26]
Hot water extract of dried leaves	Diabetes	Jamaica, Kenya	Oral [17, 25]
Infusion of entire plant	Stomach problem	Mexico	Oral [17, 27]
a) Hot water extract of leaves b) Root extract	Diabetes, rheumatism, Hypotensive, febrifuge	Mozambique	Oral [17, 28]
Hot water extract of dried entire plant	Cancer, Heart disease, Leishmaniasis	Peru	Oral [17, 29]
Hot water extract of dried ovules	Diabetes	Pakistan	Oral [17,30]
Hot water extract of roots	Abortion	Philippines	Oral[17, 31]
Decoction of dried whole plant	<i>Diabetes mellitus</i> , Liver disease	Taiwan	Oral [17, 32]
Hot water extract of dried plant	Diabetes	Thailand	Oral [17, 33]

Water extract of dried root	Venereal disease	Venda	Oral [17, 25]
Hot water extract of whole plant	Diabetes, Hypertension, Dysentery, Cancer	Vietnam	Oral [17, 34]
Hot water extract of leafy stems	Diabetes	West indies	Oral [17, 25]

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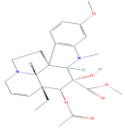
### 3.4PHYTOCHEMISTRY OF *Catharanthus roseus*:

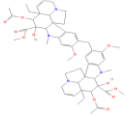
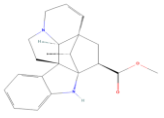
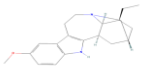
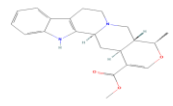
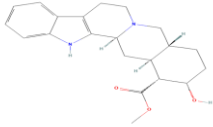
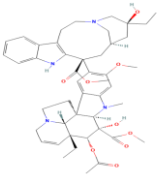
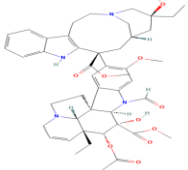
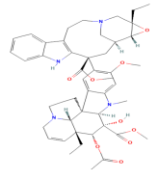
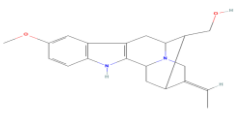
149 *C. roseus* contains a variety of alkaloids (Nitrogen-containing organic compounds  
150 other than amino acids, peptides, purines and derivatives, amino sugars, and  
151 antibiotics) [10]. These compounds had a nitrogen atom that was not found in any of  
152 the ring systems found in ephedrine, cathinone, or colchicine. This plant had been  
153 discovered to have a variety of key bioactive components that contributed  
154 significantly to herbal medicine production; nevertheless, their levels in the plant  
155 were generally insignificant. Plant secondary metabolite production was involved  
156 with the biotic and abiotic influences. Because of the plant's protective function,  
157 environmental disturbances have been discovered to induce the development of  
158 secondary metabolites, including alkaloids. As a result, more *in-vitro* and *in- vivo*  
159 investigations are focusing on manipulating environmental variables such as light,  
160 salinity, soil types and nutrients, drought, and metal stress to increase the levels of  
161 these chemicals. Citronellol (7.9%), geraniol (7.9%), (E,E)-2,4-hexadienal (7.7%),  
162 (Z,E)-pentadecanal (6.6%), and phytol (6.4%) were the primary volatile chemicals  
163 found from an Indian sample; tricosane(37.9%), heneicosane (20.8%), and  
164 tetracosane (6.1%) were the main volatile compounds discovered in the flower,  
165 palmitic acid (64.9%), methyl palmitate (7.7%), and myristic acid (6.6%) were found  
166 in high concentrations in the leaves, as well as palmitic acid (28.9%), methyl  
167 palmitate (13.1%), and geranylacetone (5.2%) were contained in the marcs-  
168 industrial plant residues [35]. Table 2 gives the structure of a few important chemical  
169 constituents.

170

**Table 2. Some crucial phytochemical structure of *C. roseus***

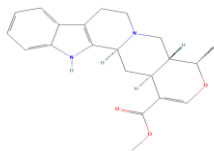
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Chemical constituents	Chemical structure	Isolated from
Catharanthine		Whole plant [10, 36]
Vindoline		Leaf, stem, and root [10, 36, 37]

Vindolicine		Leaf [10, 37]
Vindoline		Leaf [10, 37]
Ibogaine		Leaf [10, 38]
Raubasine		Leaf [10, 38]
Yohimbine		Leaf, stem, root [10, 39]
Vinblastine		Leaf, stem, root [10, 39]
Vincristine		Leaf, stem, root [10, 39]
Leurosine		Whole plant [10, 40]
Lochnerine		Whole plant [10, 41]

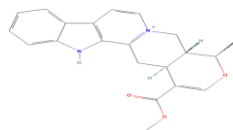
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Ajmalicine Leaf, stem, root  
[10, 39]



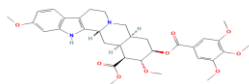
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Serpentine Root [10, 39]



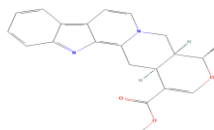
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Reserpine Leaf, stem, root  
[10, 39]



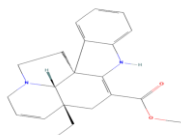
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Alstonine Root [10, 42]



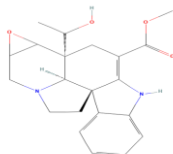
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Tabersonine Root [10, 43]



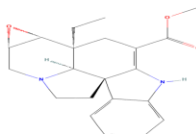
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Horhammericine Root [10, 44]



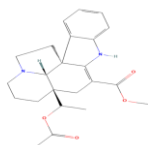
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Lochnericine Root [10, 45]



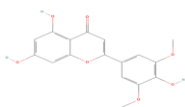
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Echitovenine Root [10, 46]



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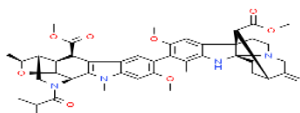
Tricin Flower [10, 45]



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Vingramine

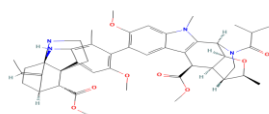
Seeds[10, 47]



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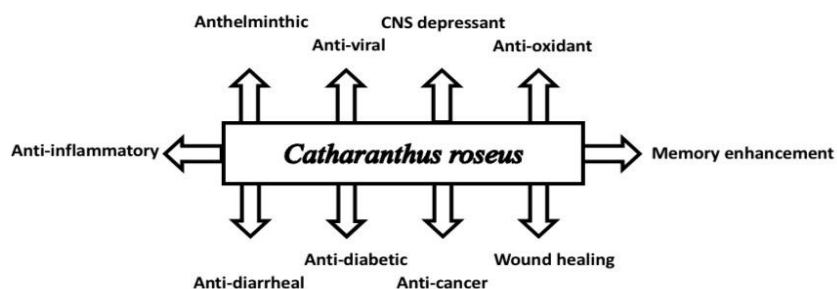
9Methylvingramine

Seeds [10, 47]



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### 3.5 Pharmacological activities



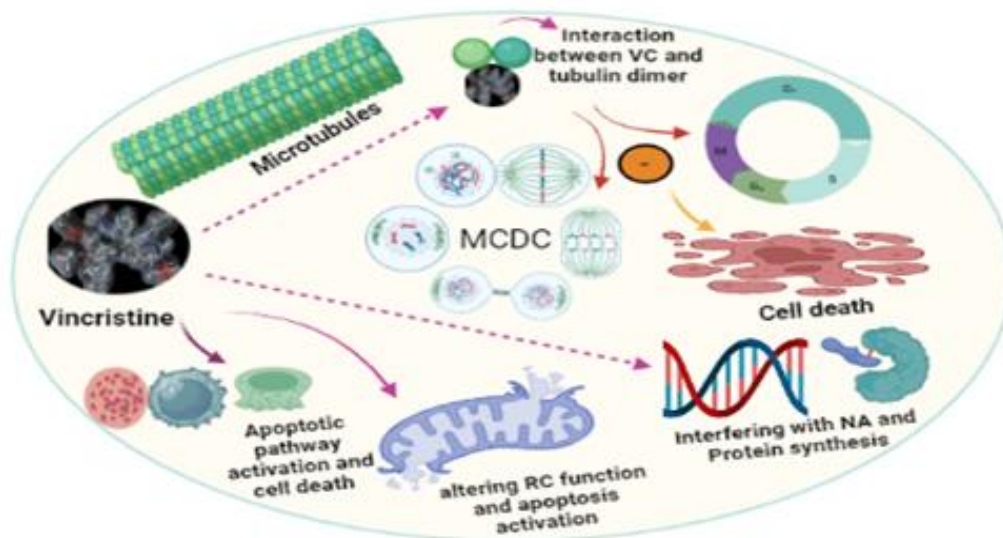
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179 **Fig.2. Pharmacological properties of *Catharanthusroseus***

180

#### 3.5.1 Anticancer activity:

181 Vinblastine and vincristine are anticancer alkaloids isolated from the stems and  
182 leaves of *C. roseus*. Some human cancers respond to these alkaloids by slowing  
183 down their growth. They block the formation of the mitotic spindle by preventing the  
184 assembly of tubulin dimers into microtubules[48]. Vinblastine is suggested for  
185 Hodgkins's disease and choriocarcinoma and is used experimentally to treat  
186 neoplasms. Another alkaloid, vincristine, is used to treat childhood leukemia. In  
187 vitro, different percentages of crude methanolic extracts of *Catharanthus* were  
188 discovered to have considerable anticancer activity against a variety of cell types,  
189 with the greatest activity against multidrug-resistant tumor types. The major  
190 mechanisms of the cytotoxicity of vinca alkaloids include interactions with tubulin  
191 and disruption of microtubule function, especially of microtubules that help to  
192 compensate the mitotic spindle apparatus, resulting in the metaphase arrest. They  
193 have a different mechanism of action and do not have cross-resistance with drugs  
194 that alkylate deoxyribonucleic acid (DNA)[49].



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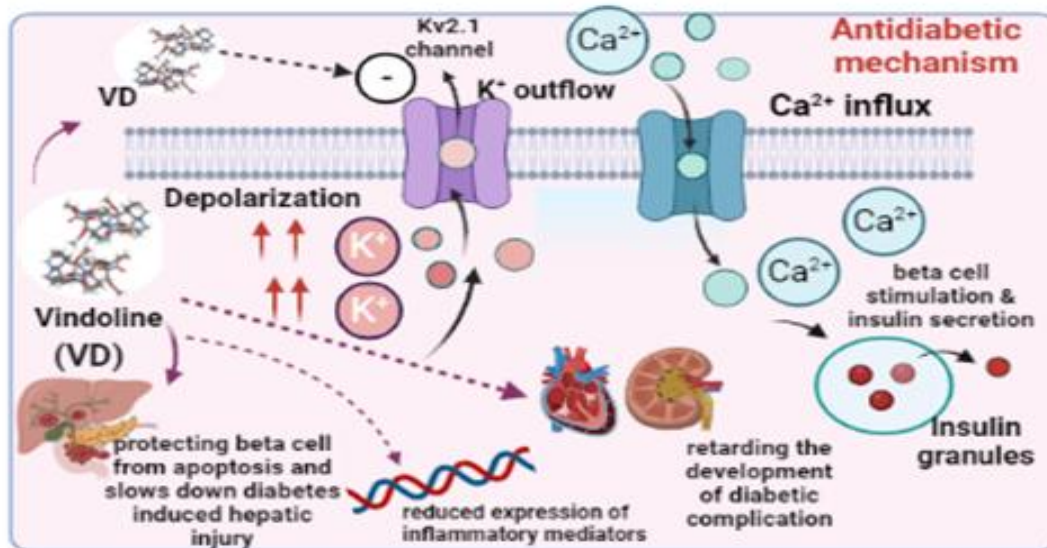
196 **Fig.3. Mechanism of action of vincristine** (Source references: Dhyani P et al,  
 197 2022; Thirumarar R et al, 2007; Starobova H and Vetter I, 2017; Lahare RP et al,  
 198 2020; Hoi HT, 2021; Saha A, 2022)

199 **The anticancer** mechanism of action of Vincristine leads to cancer cell arrest and  
 200 death. Vincristine (VC) causes the depolymerization of microtubules through binding  
 201 to tubulin leading to the mitotic cell division cycle (MDC) being arrested at  
 202 metaphase steps. This disrupted the formation & assembly of the mitotic spindle,  
 203 mainly during the S and M phases of the cell cycle. Therefore, destabilization &  
 204 interruption in microtubule dynamics leads to automated cell death or apoptosis. It  
 205 alters the function of the respiratory chain (RC) in mitochondria, followed by the  
 206 activation of apoptosis and cell death. It provides an effect contributing to the  
 207 degradation of cancer cells through the activation of the immune system and  
 208 apoptotic pathway. Another mechanism involves interfering with the synthesis of  
 209 protein & nucleic acid (NA) by blocking the utilization of glutamic acid[50-55].

### 210 **3.5.2 Anti-diabetic activity:**

211 Diabetes and related complications are major medical problems despite the  
 212 introduction of anti-diabetic agents. Patients with non-insulin-dependent diabetes  
 213 mellitus (NIDDM) are treated orally by folklore with various plant extracts since time  
 214 immemorial. Crude extracts of *C. roseus* were found to possess significant anti-  
 215 diabetic potential in different methods, including oral glucose tolerance test (OGTT)  
 216 on rodent model, alloxan, and streptozotocin-induced diabetic rats and rabbits at  
 217 different doses [56, 57]. Vindoline, vindolicine, vindolinine, vinculin, and  
 218 vindogentianine were the remarkable natural alkaloids isolated from the leaf extract  
 219 of *C. roseus* that might contribute to this potential effect [58]. Xin-gang Yao et al.  
 220 investigated that vindoline improved glucose homeostasis significantly in db/db mice  
 221 and streptozotocin combined with a high-fat diet (STZ/HFD) induced type 2 diabetes  
 222 in rats [59]. During 2019, Oluwafemi Omoniyi et al reported that vindoline provided a  
 223 prominent reduction in the blood glucose level in diabetic rats as well as reducing

224 different complications including diabetic cardiovascular and kidney disease [60]. It  
 225 provides antidiabetic action through different mechanisms including increasing  
 226 insulin secretion and insulin sensitivity, protecting the pancreatic-beta cell from  
 227 dysfunctions and apoptosis, inhibiting the expression of different inflammatory  
 228 mediators, and **reducing diabetic** complications [61]. Earlier a computational  
 229 background on  $\alpha$ -glucosidase inhibitors indicated that vindoline showed the highest  
 230 binding affinity for the target enzyme among 32 natural alkaloids isolated from  
 231 different plants [62]. Vindogentianine **provided the potential for the treatment** of type  
 232 2 diabetes (T2D) through the enhancement of glucose uptake and PTB-1B inhibition  
 233 [63].



234

235 **Fig.4. Mechanism of action of vindoline against type-2 diabetes.**(Source  
 236 references:Yao XG et al, 2013; Oguntibeju OO et al, 2019; Goboza M et al, 2019;  
 237 Tiong SH et al, 2015; Goboza M et al, 2020; Prabhakar PK and Doble M, 2011)

238 Vindoline provided antidiabetic activity by increasing the action potential of the  
 239 membrane through inhibition of Kv2.1 (voltage-gated K channel) leading to  
 240 enhanced Ca<sup>2+</sup> flow through an L-type voltage-dependent Ca<sup>2+</sup> channel. Increased  
 241 intracellular Ca<sup>2+</sup> caused stimulation of pancreatic beta cells and secretion of insulin  
 242 from beta cells through **the exocytosis** process. It also acted as an insulin sensitizer  
 243 and protected the pancreatic beta cell from apoptosis induced by cytokine through  
 244 Kv2.1 inhibition. It prevented diabetes-induced hepatic diseases by decreasing  
 245 serum alanine transferase (ALT), alkaline phosphatase, and aspartate  
 246 aminotransferase (AST) levels. It inhibited inflammatory gene expression (IL-1 $\beta$ , IL-  
 247 6, and TNF- $\alpha$ ) by nuclear factor kappa light chain enhancer of activated  $\beta$ -cells (NF-  
 248 kB) pathway. It also slowed down the diabetic cardiovascular complications  
 249 development through the reduction of total cholesterol (TC), low-density lipoprotein  
 250 (LDL), very low-density lipoprotein (VLDL), and triglycerides (TG), and diabetic  
 251 nephropathy by lowering the serum urea and creatinine level[59, 60, 61, 63, 64, 65].

252

253 **3.5.3 Antioxidant activity:**

254 In case of *C. roseus*, the presence of phenolic compounds had been reported. The  
255 presence of caffeoylquinic acids was found in parts of *C. roseus* in addition to other  
256 flavonoid derivatives. DPPH inhibitory activity of four alkaloids vindoline, vindolidine,  
257 vindolicine and vindolinine were evaluated and compared to ascorbic acid which  
258 was used as the reference standard. All four alkaloids showed free radical  
259 scavenging activities. DPPH inhibitory activity of vindoline, vindolidine, and  
260 vindolinine was better than that of the control [37/66]. Investigation concurred by  
261 Asheesh Kumar et al (2012) of the antioxidant activity of *C. roseus*, found that the  
262 habitat temperature affected the antioxidant activity of *C. roseus*. Previous research  
263 presented the results of several *in vitro* antioxidant assays in investigating the effect  
264 of solvent on the extraction and total antiradical potential of varieties extracts of *C.*  
265 *roseus*[38/67, 39/68].

266 **3.5.4 Anti-diarrheal activity:**

267 An investigation was conducted *in vivo* antidiarrheal activity of the ethanolic extract  
268 of *Catharanthus roseus* in Wistar rats[69]. Ethanolic leaf extract of *C. roseus* had  
269 exhibited antidiarrheal activity by reducing castor oil-induced diarrhea in Wistar rats  
270 and the investigation notified that the fractions of the ethanolic extract were  
271 responsible for the observed antidiarrheal effect of *C. roseus*[69].

272 **3.5.5 Anthelmintic activity:**

273 Infections with helminths cause chronic diseases in humans and livestock. The  
274 anthelmintic properties of *C. roseus* were assessed using *Pherethimaposthuma* as  
275 an experimental model and piperazine citrate as a standard reference. The  
276 ethanolic extract at 250 mg/ml showed significant anthelmintic action with a death  
277 time of 46.33 minutes, while the conventional medication at 50 mg/ml showed a  
278 death time of 40.67 minutes. The ethnomedical claims of *C. roseus* as an  
279 anthelmintic plant were supported by this study [70].

280 **3.5.6 Anti-viral activity:**

281 *Catharanthus* has an antiviral impact on the simplex herpes virus (type I) with a  
282 cytopathogenic effect at 0.8 µg/mL [4]. Catharoseumine, a monoterpene indole  
283 alkaloid with a unique peroxy bridge, was shown to be a potential inhibitor of  
284 falcipain-2 protozoa parasites (causes of malaria) having an IC<sub>50</sub> value of 4.06 µM.  
285 Vinblastine and vincristine had an antiparasitic effect on *Trypanosoma* where the  
286 parasite causing human trypanosomiasis, slowed down the mitosis and changed  
287 cellular structure in a dose-dependent way. Here, in comparison, 15 µM vinblastine  
288 and 50 µM vincristine decreased cellular division and cytokinesis as well as cellular  
289 morphology, whereas 3 µM vinblastine and 10 µM vincristine inhibited cytokinesis  
290 without affecting cell cycle progression [4].

291 **3.5.7 Anti-inflammatory activity:**

292 Various inflammatory diseases such as asthma, rheumatoid arthritis, hepatitis, and  
293 colitis **are alarming** causes of death and disability around the world. The solvent  
294 extracts of the whole plant of *C. roseus* had been found to possess the anti-  
295 inflammatory potential. The aqueous and ethanolic extracts of *C. roseus* had shown  
296 adequate anti-inflammatory **properties** during **in vivo carrageenan-induced**  
297 **antiinflammatory** study at doses of 250 and 300 mg/kg. During this assay, the  
298 ethanolic and aqueous solvent extracts of *C. roseus* plant reduced the carrageenan-  
299 induced edema in albino rats. **However**, the anti-inflammatory activities of ethanolic  
300 and aqueous extracts were found more **statistically significant** in comparison to non-  
301 polar extracts. These reduced inflammation in a dose-dependent manner.  
302 Vinpocetine, a synthetic ethyl ester of the apovincamine alkaloid, **was found**  
303 effective in **the reduction of** carrageenan-induced inflammation [71].

#### 304 **3.5.8 Wound healing activity:**

305 **In 2007, Nayak et al.** evaluated the wound healing activity of the ethanolic extract of  
306 *C. roseus* flower in Sprague Dawley rats when compared with placebo controls [13].  
307 The study revealed that there was a significant increase in tensile strength and  
308 wound contraction leading to an increase in hydroxyproline content which might be  
309 attributed to the phytoconstituents present in the extract. Recently **Satish et al**  
310 **(2021)** studied the wound healing effect of the extract of *C. roseus* flower in three  
311 cutaneous models of wound and sucralfate was used as a standard. This study  
312 revealed that the tissue breaking strength was remarkably increased in the incision  
313 model, epithelization and period to complete epithelization was decreased in the  
314 excision wound and the increase of hydroxyproline content accelerated the rate of  
315 wound contraction in the dead space model of adult Albino mice [72]. Further  
316 studies should be carried out to formulate a standard topical product of *C. roseus*  
317 extract commercially so that it can be used in wound management.

#### 318 **3.5.9 CNS depression activity:**

319 In experimental animal models, locomotor activity and pentobarbitone-induced  
320 sleeping time were used to examine the CNS depressing effect of *C. roseus* leaves.  
321 In the experiment, petroleum ether, chloroform, and ethanol extracts were prepared  
322 through fractionation sequentially. At the doses of 200 and 400mg/kg oral  
323 administration of petroleum ether extracts, substantial and dose-dependent CNS  
324 depressing action was observed. Diazepam at 4 mg/kg was used as a reference  
325 standard in this model. In the experimental animals, a dose of 100 mg/kg had no  
326 discernible effect. In qualitative chemical analysis, petroleum ether extract revealed  
327 the presence of steroids, tannins, flavonoids, glycosides, alkaloids, and phenolic  
328 substances. Petroleum ether extracts were high in steroid and phenol content,  
329 particularly flavonoids, which had been shown to have sensitive compounds in  
330 preclinical investigations for CNS depressive effect. The evaluated properties of *C.*  
331 *roseus* leaves could be due to the presence of steroids, glycosides, and phenolic  
332 compounds [73].

#### 333 **3.5.10 Memory enhancement activity:**

334 Vinpocetine, which was derived from the alkaloid vincamine, has been reported to  
335 be the most fascinating dietary substance because of its ability to increase brain  
336 function and memory. Therefore, it could be very helpful in the treatment of  
337 Alzheimer's disease. Vinpocetine at doses up to 60 mg/d was shown to be well  
338 tolerated in clinical trials for dementia and stroke, with no serious side effects  
339 reported. Blood thinners including warfarin, aspirin, and various dietary supplements  
340 like ginkgo, vitamin E, and garlic should not be used with vinpocetine [53].

#### 341 4. CONCLUSION

342 In this minireview, we have identified the presence of vincristine and vinblastine in  
343 *C. roseus*, well-known drugs used in cancer treatment. The bark, roots, leaves, and  
344 flowers of this plant also other major group of phytochemicals such as saponins,  
345 alkaloids, carbohydrates, flavonoids, and other substances. These compounds  
346 possess antibacterial, antiulcer, antioxidant, hypolipidemic, antidiabetic, and other  
347 properties. We hope this minireview can comprehensively contribute to identify the  
348 chemical components and pharmacological effects of *C. roseus*, intending to  
349 facilitate future research. In the future, this review paper will spread awareness about  
350 the medicinal properties of this plant to the scientific community.

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#### 353 COMPETING INTERESTS

354 The authors declare that they have no competing interests.

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