# CBD F/1: Unwind™

Healthy Sleep Response\* 30ml Full Spectrum CBD Product Information

**Unwind** genetically aligned CBD formulation oil is administered orally and sublingually (under the tongue). Formulated to genetically align with individual DNA profiles, **Unwind** consists of a proprietary formulation of cannabidiol (CBD) derived from hemp, terpenes and essential oils in specific ratios to promote a healthy sleep response and sense of well-being when taken regularly.\*

## PRODUCT DESCRIPTION Primary Ingredients

# Hemp-derived cannabidiol (CBD)

Cannabidiol or CBD, is one of the many cannabinoids, or chemical compounds, found in marijuana and hemp. Unlike THC, CBD is non-intoxicating but still psychoactive. In other words, CBD interacts with your endocannabinoid receptors (psychoactive), and proteins and cells in the brain, but will not cause impairment or "get you stoned" (non-intoxicating).

CBD has been shown in a number of phase I and phase II research, and phase III clinical trials, to either directly or indirectly affect symptoms of restless sleep and sleeplessness[1], anxiousness and stress[2], depressed mood[3], PTSD[4], Multiple Sclerosis[5, 6], epilepsy[7], psychopathy[8], pain[9], inflammation[10], ADHD[11], cognitive impairment and memory function[12,13], and nausea[14].

According to a 2013 study published in the British Journal of Clinical Pharmacology, CBD benefits including acting in some experimental models as an anti-inflammatory, anticonvulsant, antioxidant, antiemetic, anxiolytic and antipsychotic agent, and is therefore a potential medicine for the treatment of neuroinflammation, epilepsy, oxidative injury, vomiting and nausea, anxiety and schizophrenia.[15]

CBD can also boost levels of naturally occurring endocannabinoids in your brain such as anandamide. Over time, anandamide can stimulate neurogenesis in certain parts of your brain. This has been linked to improvements in anxiety and stress resilience.[16]

CBD has been shown to activate the 5-HT1A serotonin receptor[<u>17</u>], which may help with anxiety, addiction, appetite, sleep, nausea, vomiting. It also binds to TRPV1 receptors[<u>18</u>], which has been shown to moderate pain and inflammation. CBD activates peroxisome proliferator activated receptors (PPARs), which has been shown to produce an anti-cancer effect and help with Alzheimer's[<u>19</u>].

In addition cannabinoids and terpenes have been shown to balance endocannabinoid deficiencies[20] which may help promote a homeostatic condition necessary to support overall good health and wellness[21].

# **INDICATIONS FOR USE**

Unwind is intended as a dietary supplement to support healthy sleep, and promote a sense of well-being and relaxation.\*

Unwind formulation uses a proprietary blend of full spectrum cannabidiol (CBD), terpenes and essential oils. All ingredients are derived from natural, botanical sources and processed to ensure a 95% potency. This unique process allows for formulation consistency, and ensures that each tincture provides consistent health outcomes associated with the formulation.\*

|                      | Indications   | Mechanism of Action  | Cautions  |
|----------------------|---|--|---|
| Cannabidiol<br>(CBD) | Research suggests that the<br>endocannabinoid system plays a<br>role in maintaining certain body<br>functions, such as mood,<br>appetite, sleep, and regulating<br>circadian rhythms [22].<br>Additionally, cannabidiol (CBD)<br>has been shown to decrease<br>anxiety and pain, both of which<br>have been shown to interfere with<br>restful sleep. By reducing certain<br>symptoms, it's also possible that<br>sleep may improve[1]. | Within the endocannabinoid system is a network of<br>cannabinoid receptors in the brain and central nervous<br>system. The two primary receptors identified are CB1 and<br>CB2. Cannabinoids attach to these cells, which can produce<br>a variety of biological effects[9,16,21]<br>Cannabinoids act on multiple systems and it is understood<br>that cannabinoids interact with many neurotransmitters and<br>neuromodulator systems. It is important to note that<br>cannabinoids have the ability to interact with all kinds of<br>cellular pathways implicated in a range of diseases.<br>Cannabinoids act as ligands (a small molecule able to dock<br>onto the binding site of a protein) conferring their ability to<br>modulate a receptor's behavior and consequently their<br>downstream biological pathways. Although the<br>cannabinoids may have similar structures, they display a<br>remarkably wide array of actions.<br>Some research indicates that CBD may interact with specific<br>receptors, potentially affecting the sleep/wake cycle [1,23]. | <ul> <li>Unwind is contraindicated for anyone:</li> <li>With a known or suspected allergy to hemp, other cannabinoids, Medium Chain Triglyceride (MCT) carrier oil and coconut.</li> <li>With significant hepatic or renal impairment</li> <li>With serious cardiovascular disease, such as ischaemic heart disease, arrhythmias, poorly controlled hypertension or severe heart failure</li> <li>With a history of schizophrenia or any other psychotic disorder</li> <li>Under 18 years of age</li> <li>Who is pregnant or nursing</li> </ul> |

# Interactions:

Food may affect the absorption and blood levels of cannabidiol. To avoid significant fluctuations in blood levels, cannabidiol should be taken about the same time each day consistently, either with or without food. Consumption of grapefruit and grapefruit juice should be limited, as it may increase the blood levels of cannabidiol. Patients who consume grapefruit or grapefruit juice may require a lower dosage of the medication based on blood levels. Talk to your doctor if you have any questions or concerns.

It is important to tell your doctor about all other medications you use, including vitamins and herbs. Do not stop using any medications without first talking to your doctor.

For more information on specific drug to drug interaction visit <u>www.mydna.live</u> for a free drug to drug interaction tool.

### Terpenes

The hemp plant includes a wide variety of chemicals and compounds. About 140 of these belong to a large class of aromatic organic hydrocarbons known as terpenes. Terpenes may play a role in boosting the therapeutic effect of cannabidiol. Terpenes are believed to act on receptors and neurotransmitters[24], which

means they may play a role in providing therapeutic value in achieving health sleep patterns. Some terpenes appear to act as serotonin uptake inhibitors (similar to antidepressants like Prozac) and have been shown to enhance norepinephrine activity (similar to tricyclic antidepressants like Elavil)[25]. As well, some terpenes have been shown to increase dopamine activity[24], and seem to augment GABA[26] (the "downer" neurotransmitter that counters glutamate, the "upper") which is important to achieve healthy sleep. More specific research is needed for improved accuracy in describing and predicting how terpenes in hemp can be used for specific health concerns.

The FDA, as well as other international health and safety agencies, have recognized terpenes as safe.

# Unwind formulation uses a proprietary blend of terpenes derived from natural, botanical sources and refined to ensure a 95% potency. This process allows for formulation consistency to ensure that each tincture provides the health outcomes you expect, every time.\*

|             | Indications  | Mechanism of Action  | Cautions                       |
|-------------|--|--|--------------------------------|
| Myrcene     | Sedation and supports prolonged sleep<br>time[27]. Muscle relaxing effects[28].<br>Analgesic[24,29] and anti-inflammatory<br>properties[30]<br>Shown to block carcinogenic effects of<br>aflatoxin[31] in the liver.   | Appears to enhance activity of the inhibitory GABAA<br>receptor[32]. This mechanism is shared with<br>benzodiazepines, which are sedative drugs used for<br>anxiety and sleep.<br>Shown to lower resistance across the blood to brain<br>barrier[33], supporting a more rapid onset of the<br>cannabinoid effect. Shown to increase the maximum<br>saturation level of the CB1 receptor, allowing for a<br>greater maximum psychoactive effect.  | Known allergy to the compound. |
| Terpinolene | Terpinolene has been found to be a central<br>nervous system depressant[34] and may<br>induce drowsiness or sleep.<br>Has been used to reduce hyperactivity and<br>anxiety. Terpinolene was found to markedly<br>reduce the protein expression of AKT1 in K562<br>cells and inhibited cell proliferation involved in<br>a variety of human cancers[35].                                    | Terpinolene has been shown to provide a calming effect<br>in persons exhibiting hyperactive behaviour[36]. In mice,<br>terpinolene has been shown to reduce locomotor<br>activity. Like some of the other cannabis terpenes,<br>terpinolene can enhance activity of the inhibitory<br>GABAA receptor[37], which could present as a calming<br>and relaxing of the mind and body. Terpinolene also<br>possesses pain reducing properties, although no<br>studies have yet confirmed this in humans. | Known allergy to the compound. |
| Linalool    | Linalool is a non-cyclic monoterpenoid and<br>may promote calming, relaxing effects [38].<br>Linalool inhibited acetylcholine release and<br>alters ion channel function at the<br>neuromuscular junction [39,40].<br>Studies also suggest that linalool may act to<br>boost the immune system as it directly<br>activates immune cells through specific<br>receptors and/or pathways[41]. | Linalool may work in several ways: First, it is an<br>antagonist of the NMDA receptor[44, 45]. This means<br>that it can reduce activation of this receptor by<br>glutamate, the main excitatory neurotransmitter of the<br>brain. Other NMDA antagonists also have strong<br>sedative properties. Second, it enhances GABA<br>activity[46], although it does this in a different way than<br>myrcene since it does not bind the same receptor site.   | Known allergy to the compound. |

| It has been shown to significantly reduce lung inflammation[ <u>42</u> ].   |  |
|---|--|
| Linalool may be useful in the treatment of<br>Alzheimer's disease by reversing the<br>histopathological (the microscopic<br>examination of biological tissues to observe<br>the appearance of diseased cells and tissues in<br>very fine detail) hallmarks of Alzheimer's<br>Disease and could restore cognitive and<br>emotional functions via an anti-inflammatory<br>effect[43]. |  |

# **Essential Oils**

# Unwind genetically aligned formulation uses only the highest quality essential oils, blended from naturally derived botanical sources to a minimum 95% purity.

|              | Indications   | Mechanism of Action  | Cautions                   |
|--------------|---|--|----------------------------|
| Lavender Oil | There is growing evidence suggesting that lavender<br>oil may be an effective medication in treatment of<br>several neurological disorders, including depression,<br>anxiety, insomnia and Alzheimer's disease. Several<br>animal and human investigations suggest anxiolytic,<br>mood stabilizer, sedative, analgesic, and<br>anticonvulsant and neuroprotective properties for<br>lavender[47]. | Several investigations were performed to clarify<br>the mechanism of action of lavender in neuronal<br>tissues. Lavender inhibited<br>lipopolysaccharide-induced inflammatory reaction<br>in human monocyte THP-1 cells effect, which<br>might be associated with the expression of HSP70<br>[48]. Antioxidant and relatively weak cholinergic<br>inhibition was reported for lavender [49,50] and<br>linalool.[51,52,53]<br>These findings indicate that several targets<br>relevant to treatment of Alzheimer's disease. The<br>neuroprotective effect of lavender oil against<br>cerebral ischemia/reperfusion injury is suggested<br>to be attributed to its antioxidant effects[54].<br>Lavender oil may also modulate GABAergic<br>neurotransmission, especially on GABAA<br>receptors, and enhance the inhibitory tone of the<br>nervous system.[55,56,57] The cholinergic system<br>(a grouping of organized nerve cells in the brain) is<br>suggested to play a role in lavender analgesic, | Known allergy to Lavender. |

|                   |   | antianxiety, anti- depression, and anticonvulsant effects. [58, 59, 60]  |  |
|-------------------|---|--|--|
| Chamomile<br>Oil  | Traditionally, chamomile has been used to treat<br>insomnia and to induce sedation (calming effects).<br>Chamomile is widely regarded as a mild tranquilizers<br>and sleep-inducer[61].<br>Chamomile extracts exhibit benzodiazepine-like<br>hypnotic activity. Studies in preclinical models have<br>shown anticonvulsant and CNS depressant effects<br>respectively.<br>Compounds, other than apigenin, present in extracts<br>of chamomile can also bind BDZ and GABA<br>receptors in the brain and might be responsible for<br>some sedative effect; however, many of these<br>compounds are as yet unidentified[62].   | For more information on mechanism of action and<br>possible side effects:<br><u>https://www.foundationalmedicinereview.com/wp-</u><br><u>content/uploads/2019/02/v13-1-58.pdf</u>  | Persons with known sensitivity<br>Chamomile.   |
| Sandalwood<br>Oil | Intraperitoneal administration of alpha- and<br>beta-santalols in mice increased<br>hexobarbital-induced sleeping time. Oral,<br>intraperitoneal, and intracerebroventricular<br>administration of alpha-santalol reduced rectal<br>temperature and spontaneous motor activity more<br>effectively than beta-santalol. However, beta-santalol<br>was found to decrease acetic acid-induced writhing<br>more effectively than alpha-santalo[63].<br>Though not statistically significant, leg and foot<br>massage with sandalwood oil reduced anxiety in<br>patients[64]. Topical administration of sandalwood<br>oil produced "harmonizing" effects (ie, a reduction in<br>the level of autonomic nervous system arousal but<br>no behavioral level deactivation) in healthy<br>volunteers, whereas alpha-santalol had<br>relaxing/sedative effects[65] | Known to mediate inflammation in vitro through<br>multiple mechanism: the oil inhibits the oxidative<br>enzyme 5-lipoxygenase and has DPPH radical<br>scavenging activity and, in vivo, was able to<br>protect mouse livers from damage resulting from<br>oxidative stress and the formation of reactive<br>oxygen specie[66]. | Contraindication and<br>interactions: None well<br>documented[66]. Information<br>regarding safety and efficacy in<br>pregnancy and lactation is<br>lacking. Sandalwood oil can<br>cause dermatitis, although it is<br>generally considered to be no-<br>irritating to human skin. |

# PHYSICAL DESCRIPTION

Unwind is an amber-coloured oil preserved in a dark violet Miron glass tincture bottle.

Miron glass and product preservation.

Miron glass[67] is vastly different from the typical glass or plastic CBD tincture bottles available. The dark violet glass protects and preserves, allowing only ultraviolet, violet and infrared spectrum to penetrate to the liquid within. It works as a natural filter blocking the complete spectral range of visible light, with the exception of violet light. Other glass types allow all or most of the visible light spectrum to pass through. This makes them less capable of protecting the sensitive properties of the compounds, like micronutrients, cannabinoids, terpenes and essential oils.

Miron glass relies on a little-known theory called biophotonics. Biophotons are thought to be essential to intercellular communication and the promotion and balance of energy [68]. This unique dark violet glass has been shown to have outstanding properties of preservation over extended periods of time[69]. Water, for example, has been kept fresh in violet Miron glass for over three years without any artificial preservation techniques.

Miron glass offers unique protection from other light frequencies and is believed to enhance the durability and potency of products like CBD oil, without the addition of any preservatives.

#### DOSAGE AND ADMINISTRATION

| SUPPLEMENT FACTS   |                     |      |  |
|--|---------------------|------|--|
| Serving size: 0.7ml (~12 drops) Serving per container: 42**  |                     |      |  |
|  | Amount per serving: | %DV  |  |
| Calories   | 4                   |      |  |
| Total Fat  | Less than 1g        | <1%† |  |
| Hemp Extract (Aerial parts)<br>(88% CBD Distillate) (<0.3% THC)  | 23.81mg             | +    |  |
| †Percent Daily Values based on a 2000 calorie diet<br>‡Daily Value Not Established   |                     |      |  |
| Other ingredients: Medium Chain Triglyceride (MCT) carrier oil (CONTAINS COCONUT), Lavender Oil, Chamomile Oil and Sandalwood Oil. |                     |      |  |
| **DOSAGE PER DROP IS 1.96MG TO 2.04MG AND VARIES BASED ON DROP SIZE.   |                     |      |  |

#### Suggested use:

Shake well. Take 0.7ml under the tongue and hold for 1 minute. Swallow. Use 2 times per day or as directed by a physician. Individual results may vary.

### Warning:

Intended for adult use only. Consult a doctor before use if you are pregnant, nursing or taking any medications. Keep out the reach of children. Store in a cool, dry place.

\*These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure, or prevent any disease. US Patent PCT/US2018/058199

Manufactured in accordance with GMP and ISO 9001 quality assurance standards for: Effica Wellness, Thousand Oaks, CA, USA

For a complete Certificate of Analysis (COA), more information and links to references for this formulation visit <u>www.efficawellness.com</u>/unwind.