

**PATIENT PACKAGE INSERT
IN ACCORDANCE WITH THE
PHARMACISTS' REGULATIONS
(PREPARATIONS) – 1986**
**The medicine is dispensed with a
doctor's prescription only**

Stalevo®
50/12.5/200 mg
Film-coated tablets

Stalevo®
75/18.75/200 mg
Film-coated tablets

Stalevo®
100/25/200 mg
Film-coated tablets

Stalevo®
125/31.25/200 mg
Film-coated tablets

Stalevo®
150/37.5/200 mg
Film-coated tablets

Stalevo®
200/50/200 mg
Film-coated tablets

Active ingredients
Stalevo 50/12.5/200 mg
Each tablet contains levodopa/
carbidopa/entacapone
50/12.5/200 mg

Stalevo 75/18.75/200 mg
Each tablet contains levodopa/
carbidopa/entacapone
75/18.75/200 mg

Stalevo 100/25/200 mg
Each tablet contains levodopa/
carbidopa/entacapone
100/25/200 mg

Stalevo 125/31.25/200 mg
Each tablet contains levodopa/
carbidopa/entacapone
125/31.25/200 mg

Stalevo 150/37.5/200 mg
Each tablet contains levodopa/
carbidopa/entacapone
150/37.5/200 mg

Stalevo 200/50/200 mg
Each tablet contains levodopa/
carbidopa/entacapone
200/50/200 mg

Inactive ingredients
See section 6 'Further Information' and
section 2 'Before using the medicine'
in section 'Important information
about some of the ingredients of the
medicine'.

**Read the leaflet carefully in its
entirety before using the medicine.**
This leaflet contains concise information
about the medicine. If you have further
questions, refer to the doctor or
pharmacist.

This medicine has been prescribed
to treat your ailment. Do not pass it
on to others. It may harm them even
if it seems to you that their ailment is
similar.

1. WHAT IS THE MEDICINE INTENDED FOR?

For treatment of Parkinson's disease.

Therapeutic group: Anti-Parkinson
medicines, dopa and dopa derivatives.
Stalevo contains 3 active ingredients
(levodopa, carbidopa and entacapone)
in one film-coated tablet. Parkinson's
disease is caused by low brain levels
of a substance called dopamine.
Levodopa increases the amount
of dopamine, thereby reducing the
symptoms of Parkinson's disease.
Carbidopa and entacapone improve the
anti-Parkinsonian effects of levodopa.

2. BEFORE USING THE MEDICINE

Do not use the medicine if:

- You are allergic (hypersensitive) to
levodopa, carbidopa, entacapone or
to any of the additional ingredients
of the medicine, listed in section 6
'Further Information'.
- You have narrow-angle glaucoma (a
type of eye disease).
- You have a tumor of the adrenal
gland (pheochromocytoma).
- You are taking certain medicines
to treat depression (a combination
of selective MAO-A and MAO-B
inhibitors, or non-selective MAO
inhibitors).
- You have ever had neuroleptic
malignant syndrome (NMS; this is
a rare reaction to medicines used
to treat severe mental disorders).
- You have ever had non-traumatic
rhabdomyolysis (a rare muscle
disorder).
- You have a severe liver disease.
- You are breastfeeding.

**Special warnings regarding use of
the medicine:**

**Before taking Stalevo, talk to the
doctor or pharmacist if you have or
have ever had:**

- A heart attack, or any other heart
disease, including heart rhythm or
blood vessel disturbances
- Asthma or any other lung disease
- A liver problem, because the dosage
may have to be adjusted
- A kidney or hormone-related disease
- Stomach ulcers or convulsions
- If you are experiencing prolonged
diarrhea, consult with your doctor as
it may be a sign of inflammation of the
bowel
- Any type of severe mental disorder,
such as psychosis
- Chronic wide-angle glaucoma,
as it may be necessary to adjust
your dosage and to monitor your
intraocular pressure

**Consult your doctor if you are
currently taking:**

- Antipsychotics (medicines used to
treat psychosis)
- A medicine which may cause low
blood pressure when rising from a
chair or bed. You should be aware
that Stalevo may make these effects
worse.

**Consult the doctor if during the
treatment with Stalevo you:**

- Notice that your muscles get very
rigid or jerk forcefully, or if you get
tremors, shaking, confusion, fever,
rapid pulse, or wide fluctuations in
your blood pressure. If any of these
happen to you, **contact the doctor
immediately**
- Feel depressed, have suicidal
thoughts, or notice unusual changes
in your behavior

- Find yourself suddenly falling
asleep, or if you feel very sleepy. If
this happens, do not drive and do
not use any tools or machinery (see
also section 'Driving and operating
machinery')
- Notice that uncontrolled movements
begin or get worse after you started
to take Stalevo. If this happens,
the doctor may need to change
the dosage of your anti-Parkinson
medicines
- You have diarrhea: monitoring of your
weight is recommended in order to
avoid potentially excessive weight
loss
- You experience advanced anorexia,
weakness, exhaustion and weight
decrease within a relatively short
period of time. If this happens, a
general medical evaluation, including
liver function, should be considered
- Feel the need to stop using Stalevo;
see section 'Stopping treatment'.

Tell the doctor if you or your family or
your carer notices you are developing
addiction-like symptoms leading to
craving for large doses of Stalevo
and other medicines used to treat
Parkinson's disease.

Tell the doctor if you or your family or
your carer notices you are developing
urges or cravings to behave in ways that
are unusual for you or you cannot resist
the impulse, drive or temptation to carry
out certain activities that could harm you
or others. These behaviors are called
impulse-control disorders and can
include addictive gambling, excessive
eating or spending, an abnormally high
sex drive or a preoccupation with an
increase in sexual thoughts or feelings.
The doctor may need to review your
treatments.

If you are due to undergo surgery,
please tell your doctor that you are
using Stalevo.

Stalevo is not recommended to be
used for treatment of extrapyramidal
symptoms (e.g., involuntary
movements, shaking, muscle rigidity
and muscle contractions) caused by
other medicines.

Tests and follow-up

The doctor may perform some routine
laboratory tests during long-term
treatment with Stalevo.

Periodic evaluation of hepatic,
hematopoietic, cardiovascular and
renal function is recommended during
extended treatment with Stalevo.

All patients treated with Stalevo
should be monitored carefully for
the development of mental changes
(e.g., hallucinations and psychoses),
depression with suicidal tendencies,
and serious antisocial behavior. In case
any of these changes occur, refer to the
doctor immediately.

Children and adolescents

Experience with use of Stalevo in
children and adolescents under 18
years of age is limited; therefore,
the medicine is not intended for this
population.

Elderly patients

Dosage adjustment is not necessary
for the elderly.

Drug interactions

**If you are taking, or have recently
taken, other medicines, including
non-prescription medicines and
nutritional supplements, tell the
doctor or pharmacist.** Especially if
you are taking:

Do not take Stalevo if you are
taking certain medicines for treating
depression (combination of selective
MAO-A and MAO-B inhibitors, or
non-selective MAO inhibitors).

Stalevo may increase the effects and
side effects of certain medicines,
including:

- Medicines used to treat depression
such as moclobemide, amitriptyline,
desipramine, maprotiline, venlafaxine
and paroxetine
 - Rimiterol and isoprenaline, used to
treat lung diseases
 - Adrenaline, used to treat severe
allergic reactions
 - Noradrenaline, dopamine and
dobutamine, used to treat heart
diseases and low blood pressure
 - Alpha-methyldopa, used to treat high
blood pressure
 - Apomorphine, used to treat
Parkinson's disease
- The effects of Stalevo may be weakened
by certain medicines. These include:
- Dopamine antagonists used to treat
mental disturbances, nausea and
vomiting
 - Phenytoin, used to prevent convulsions
 - Papaverine, used to relax the muscles.

Stalevo may make it harder to digest
iron. Therefore, do not take Stalevo and
iron supplements at the same time.
After taking one of them, wait at least
two to three hours before taking the
other.

Use of Stalevo and food

Stalevo may be taken with or without
food.

For some patients, Stalevo may not
be well absorbed if it is taken with, or
shortly, after a protein-rich meal (such
as meat, fish, dairy products, seeds and
nuts). Consult the doctor if you think this
applies to you.

Pregnancy

If you are pregnant or breastfeeding,
if you think you are pregnant, or are
planning to have a baby, consult the
doctor or pharmacist before taking the
medicine.

Do not use Stalevo during pregnancy,
unless the benefit to the mother
outweighs the possible risks to the
unborn baby.

Breastfeeding

Do not breastfeed during treatment
with Stalevo. Levodopa is secreted into
breast milk.

Driving and operating machinery

Stalevo may lower your blood pressure,
which may make you feel dizzy.
Therefore, be particularly careful when
you drive or when you operate any tools
or machinery.

If you feel very sleepy, or if you
sometimes find yourself suddenly
falling asleep, you must avoid driving
or engaging in any other activity that
requires alertness (operating tools or
machinery). Otherwise, you put yourself
and others at risk of serious injury or
death.

Wait until you feel fully alert before
driving or engaging in anything that
requires alertness.

Important information about some of the ingredients of the medicine

The tablets contain sucrose.

If the doctor has told you that you have
an intolerance to certain sugars, refer to
the doctor before taking Stalevo.

3. HOW SHOULD YOU USE THE MEDICINE?

Always use the preparation according
to the doctor's instructions. Check with
the doctor or pharmacist if you are
uncertain regarding the dosage and
treatment regimen of the preparation.

Dosage

The dosage and treatment regimen will
be determined by the doctor only.

Adults and elderly:

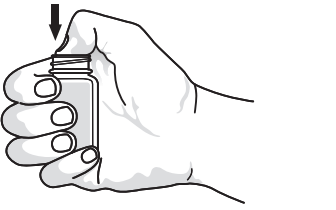
- The doctor will tell you exactly how
many Stalevo tablets to take each day.
- Do not split or break the tablet into
smaller pieces.
- Take only one tablet each time.
- Depending on how you respond to
treatment, the doctor will suggest a
higher or lower dosage.
- At dosages of 50/12.5/200 mg,
75/18.75/200 mg, 100/25/200 mg,
125/31.25/200 mg, 150/37.5/200 mg:
do not take more than 10 tablets each
day.
- At a dosage of 200/50/200 mg: do not
take more than 7 tablets each day.

**Do not exceed the recommended
dose.**

Consult the doctor or pharmacist if you
think the effect of the medicine is too
strong or too weak, or if you experience
side effects.

To open the bottle for the first time:
open the cover, and then press with
your thumb on the seal until it breaks
(see picture 1).

Picture 1



Taking more Stalevo than required

If you took an overdose or if a child has
accidentally swallowed the medicine,
immediately refer to a doctor or proceed
to a hospital emergency room and bring
the package of the medicine with you.

In case of an overdose, you may feel
confused or agitated, your heart rate
may be slower or faster than normal or
the color of your skin, tongue, eyes or
urine may change.

If you forget to take Stalevo

Do not take a double dose to
compensate for a forgotten tablet.

**If it is more than 1 hour until the next
dose:** Take one tablet as soon as you
remember, and the next tablet at the
usual time.

**If it is less than 1 hour until your next
dose:** Take one tablet as soon as you
remember, wait one hour, and then take
another tablet. After that, return to the
usual times.

To avoid possible side effects, always
wait at least one hour between taking
one Stalevo tablet and the next.

Adhere to the treatment regimen as
recommended by the doctor.

Stopping treatment

Even if there is an improvement in your
health condition, do not stop treatment
with the medicine without consulting
the doctor.

In case of discontinuation of treatment,
the doctor may need to readjust your
other anti-Parkinson medicines,
especially levodopa, so there will be
sufficient control of your symptoms.
Sudden discontinuation of Stalevo and
other anti-Parkinson medicines may
result in unwanted side effects.

**Do not take medicines in the dark!
Check the label and the dose
each time you take a medicine.**
Wear glasses if you need them.
**If you have further questions
regarding use of the medicine,
consult the doctor or pharmacist.**

4. SIDE EFFECTS

As with any medicine, use of Stalevo
may cause side effects in some users.
Do not be alarmed when reading the list
of side effects. You may not suffer from
any of them.

Many of the side effects can be relieved
by adjusting the dosage.

If during the treatment with Stalevo you
experience the following symptoms,
refer to a doctor immediately:

- Your muscles get very rigid or jerk
forcefully, you get tremors, shaking,
confusion, fever, rapid pulse or wide
fluctuations in your blood pressure.
These can be symptoms of neuroleptic
malignant syndrome (NMS, a rare
severe reaction to medicines used
to treat disturbances of the central
nervous system) or rhabdomyolysis
(a rare severe muscle disorder).
- Allergic reaction; the signs may
include urticaria, itching, rash,
swelling of the face, lips, tongue or
throat. This may cause difficulties in
breathing or swallowing.

Very common (may affect more than 1 in 10 patients)

- Involuntary movements (dyskinesias)
- Nausea
- Harmless reddish-brown discoloration
of urine
- Muscle pain
- Diarrhea

Common (may affect up to 1 in 10 patients)

- Light-headedness or fainting due
to low blood pressure; high blood
pressure
- Worsening of Parkinson's symptoms,
dizziness; drowsiness
- Vomiting; abdominal pain and
discomfort; heartburn; dry mouth;
constipation
- Inability to sleep; hallucinations;
confusion; abnormal dreams
(including nightmares); tiredness
- Mental changes – including problems
with memory; anxiety and depression
(possibly with suicidal thoughts)
- Heart or artery disease events (e.g.,
chest pain); irregular heart rhythm
- More frequent falling
- Shortness of breath
- Increased sweating; rashes
- Muscle cramps; swelling of legs
- Blurred vision
- Anemia
- Decreased appetite; decreased
weight
- Headache; joint pain
- Urinary tract infection

Uncommon (may affect up to 1 in 100 patients)

- Heart attack
- Bleeding in the gut
- Changes in the blood cell count which
may result in bleeding; abnormal liver
function test results

- Restlessness
- Psychotic symptoms
- Inflammation of the colon (colitis)
- Discolorations other than urine (e.g.,
skin, nails, hair, and sweat)
- Swallowing difficulties
- Inability to urinate

**Certain side effects may be rare or
very rare**

- Convulsions

Unknown frequency (cannot be estimated from the available data)

Craving for large dosages of Stalevo
beyond that required to control motor
symptoms, known as dopamine
dysregulation syndrome (DDS). Some
patients experience severe abnormal
involuntary movements (dyskinesias),
mood swings or other side effects after
taking a large dosage of Stalevo.

**The following side effects have also
been reported:**

- Hepatitis (inflammation of the liver)
- Itching

You may experience the following side
effects:

- Inability to resist the impulse to
perform an action that could be
harmful, which may include:
 - Strong impulse to gamble
excessively, despite serious
personal or family consequences
 - Altered or increased sexual interest
and behavior of significant concern
to you or to others, for example, an
increased sexual drive
 - Uncontrollable excessive shopping
or spending
 - Binge eating (eating large amounts
of food in a short time period) or
compulsive eating (eating more
food than normal and more than is
needed to satisfy your hunger).

Refer to a doctor if you experience
any of these behaviors; the doctor will
advise you as to how to manage or
reduce the symptoms.

**If a side effect occurs, if one of the
side effects worsens, or if you suffer
from a side effect not mentioned in
the leaflet, consult with the doctor.**

Side effects can be reported to the
Ministry of Health by clicking on the
link "Report Side Effects of Drug
Treatment" found on the Ministry of
Health homepage (www.health.gov.il)
that directs you to the online form for
reporting side effects, or by entering
the link:
<https://sideeffects.health.gov.il>

5. HOW SHOULD THE MEDICINE BE STORED?

Avoid poisoning! This medicine, and
any other medicine, should be kept in
a safe place out of the reach and sight
of children and/or infants in order to
avoid poisoning. Do not induce vomiting
unless explicitly instructed to do so by
the doctor.

Do not use the medicine after the expiry
date (exp. date) that appears on the
package. The expiry date refers to the
last day of that month.

Storage conditions:

Store below 25°C.

After first opening the bottle, use within
6 months.

Do not discard medicines in wastewater
or a household waste bin. Ask your
pharmacist how to dispose of medicines
that you no longer use. These measures
will help protect the environment.

6. FURTHER INFORMATION

In addition to the active ingredients, the
medicine also contains:

Tablet:

Mannitol, maize starch, povidone,
croscarmellose sodium, magnesium
stearate

Tablet coating:

Hypromellose, titanium dioxide,
sucrose, magnesium stearate, glycerol
85%, red iron oxide, polysorbate 80,
yellow iron oxide

Yellow iron oxide is not found in tablets
of the following dosages:

75/18.75/200 mg, 125/31.25/200 mg,
200/50/200 mg

The amount of sucrose (in mg) per tablet:

200/50/200 – 2.26 mg
150/37.5/200 – 1.91 mg
125/31.25/200 – 1.62 mg
100/25/200 – 1.55 mg
75/18.75/200 – 1.35 mg
50/12.5/200 – 1.18 mg

**What the medicine looks like and the
contents of the package**

Stalevo 50/12.5/200 mg:

A brownish-red to grayish-red colored,
round, convex, unscored tablet, marked
with LCE 50 on one side.

Stalevo 75/18.75/200 mg:

A light brownish-red colored, oval,
unscored tablet, marked with LCE 75
on one side.

Stalevo 100/25/200 mg:

A brownish-red to grayish-red colored,
oval, unscored tablet, marked with LCE
100 on one side.

Stalevo 125/31.25/200 mg:

A light brownish-red colored, oval,
unscored tablet, marked with LCE 125
on one side.

Stalevo 150/37.5/200 mg:

A brownish-red to grayish-red colored,
elongated-ellipse, unscored tablet,
marked with LCE 150 on one side.

Stalevo 200/50/200 mg:

A dark brownish-red colored, oval,
unscored tablet, marked with LCE 200
on one side.

Package sizes: A bottle that contains 10
or 30 or 100 tablets.

Not all package sizes are marketed.

License holder and address:
Inovamed Pharma Ltd., P.O.B. 2349,
Bnei Brak 5112202.

Manufacturer and address: Orion
Corporation, Espoo, Finland.

Edited in May 2020.

Registration number of the medicine
in the National Drug Registry of the
Ministry of Health:

Stalevo 50/12.5/200 mg:
132 34 31105

Stalevo 75/18.75/200 mg:
144 30 32962

Stalevo 100/25/200 mg:
132 35 31106

Stalevo 125/31.25/200 mg:
144 31 32963

Stalevo 150/37.5/200 mg:
132 36 31107

Stalevo 200/50/200 mg:
141 94 31768