Hormonal therapy for ovarian dysfunctions in high-productive cows

Liliya Khamitova*, Roman Rudakov, Mariya Knyazeva, and Anastasiya Metlyakova
Izhevsk State Agricultural Academy, Izhevsk 426069, Russia

Abstract. The analysis was made of culling data in the farm of the Udmurt Republic. As a result, it was found that the animals are eliminated due to gynecological and limb diseases. They make up 19–24 % and 18–31 % respectively. Analyzing the results of gynecological clinical examination of cows, we found a predominance of ovarian diseases 32–36 %, as well as endometritis 28–30 %. The ovarian dysfunction progresses mostly. This can be attributed to an increase in the productivity of cows in the Udmurt Republic over the past 5 years from 5601 to 6250 kg per 1 cow. In connection with an increase in the incidence of ovaries, the calf yield per 100 cows decreases from 88 to 85 %. Causes of ovarian disease can be diseases of the birth and postpartum period. For the treatment of ovarian hypofunction, 3 schemes were used for anestral and reactive reproductive cycles. Often, the cause of ovarian hypofunction is uterine pathology; drugs that have a therapeutic effect on the uterus are included in the scheme.

1 Introduction

High productivity is accompanied by a violation of the reproductive function of cattle. Now this is the main problem of increasing animal productivity and the profitability of animal husbandry. From the barren cows of the economy, no offspring has been received; premature culling of cows occurs due to massive diseases of the uterus and ovaries. A large number of young cows are removed from the herd before cost-effective use [1, 2].

The ovaries are a paired organ with generative and endocrine functions [3]. The ovary is a dynamic organ, and its structure in the reproductive period undergoes constant cyclic changes [4].

Ovarian hypofunction (Hypofunctio ovarii) is a disease characterized by a decrease in the hormonal activity of the ovaries, in which inferior reproductive cycles (areactive, anestral, alibid, anovulatory) or anaphrodisia are observed [5].

Research purpose: to develop a therapy system for ovarian hypofunction at the Rico-Agro farm in the Uvinsky district of the Udmurt Republic.

To achieve this goal, the following tasks have been developed:

• To study the prevalence of ovarian dysfunction in cows in a particular farm (clinical and instrumental research, ultrasound);
• To study the therapeutic efficacy of hormonal drugs for the treatment of ovarian hypofunction in cows;
• Develop treatment regimens.

2 Materials

The work was carried out on the basis of the farm of Rico-Agro, in the Udmurt Republic, Uvinsky district, the village of Porshur-Tuklya. The farm is a breeding factory and is engaged in the cultivation of cattle of black and motley breed.

The object of research is cattle, cows of black-motley breed, unborn cows with a service period of more than 60 days and a productivity of more than 6000 kg for 305 days of lactation.

Cows were selected by the method of pair-analogues according to the results of gynecological medical examination. They were selected according to the following characteristics: absence of reproductive cycles, violation of cycles, ovarian pathology.

To study the condition of animals on the farm, they used: retrospective, anamnestic, clinical research, including rectal research, ultrasound of the genital organs.

Animals for the experiment were selected for 10 animals with 2 and 3 calving and productivity of over 6000 kg for 305 days of lactation, as well as 5 heifers with a productivity of over 20 liters per day. The live weight of cows is over 550 kg, and the first-calf is over 500 kg.

3 Results

The farm contains 1646 heads of cattle of black and motley breed. Of these, 637 cows are cows. The average milk yield of cows for 305 days of lactation is 6250 kg. The yield of live calves is 85 %. This indicator over 5 years amounted to more than 88 %.

The Rico-Agro enterprise raises cattle of the dairy direction of a black and motley breed. They use artificial
insemination of cows and heifers in a rectocervical way
and sperm of outstanding bulls for milk productivity,
mass fraction of fat and protein, which have categories
for milk yield and mass fraction of fat. Sperm is acquired
at specialized breeding enterprises.

Animals are kept in clean, dry, warm cowsheds with
good lighting and ventilation. Cows have predominantly
average fatness.

A stall-walking system is applied for keeping
animals. Cattle are kept in rooms where both harness and
free placement are used. Milking of animals takes place
automatically in the milk pipeline on the ground and in
the milking parlor on the installation. Manure is cleaned
mechanically with a scraper conveyor.

From the main herd, cows are transferred to the
maternity ward 10 to 20 days before calving. In the
room, a leash is used, except for calving boxes, where
animals are kept freely. Straw, hay are used as litter in
the boxes. Boxes are disinfected after each calving, as
are the stalls where new-born animals are located.
Disinfection is carried out chemically after mechanical
cleaning, during a passive exercise of animals daily.

Calves are kept for up to two days in calving with
their mother, then transferred to a dispensary, and cows
– to a group of animals after calving. Milking occurs
using a machine in a milk line. Manure is removed using
a scraper conveyor.

Preventive clinics in winter are located in a building
divided into two parts – a barn and a calf. In summer,
calves are kept in individual houses with walking yards.
Animals are kept in this room for 20–30 days, then
transferred to another calf, where animals are kept for up
to 6 months. Milk calves are fed from a nipple drinker,
and calves over 10 days of age – from a bucket. In the
winter period, calves are kept in group cages of 5-6
animals each. Cleaning of litter and manure is carried out
manually every day.

Animals up to 6 months old are kept in cages, in
groups of 10-15 animals each. Feeding is carried out
from group feeders twice a day – a substitute for whole
milk, in free access are concentrates, hay, chalk; from 3-
4 months old, they begin to add silage to the diet. Near
the calf, there is a walking yard where passive exercise is
carried out. Manure is cleaned manually and
mechanically using a scraper conveyor.

The exercise is present in the main livestock, calves
up to 6 months old, but in the winter-stall period they do
not receive a full-fledged daily exercise, because the area
of walking areas does not allow to drive out all animals
every day. Newly born animals receive a daily passive
exercise.

Silage-silage type of feeding is applied, feed is
distributed using a mixer. Also, additives are used –
protein-mineral vitamin premix, selenium and kayod
preparations, since the Udmurt Republic is located in the
biogeochemical zone with the insufficient content of
selenium and iodine in soils and water. Feeding in the
winter-stall period is carried out twice a day. The total
nutritional value of the diet of lactating cows averages
21.7 feed units. In the summer-pasture period, the diet of
lactating cows includes: green mass, concentrates, salt
and additives. For each liter of milk, 0.2 kg of
concentrates are added to cows. Individual drinking
bowls are used for drinking. In the chemical study of
feed, the following results were obtained: hay and silage
of the 1st – 2nd class, haylage - the 2nd class.

| Causes                  | 2014 | 2015 | 2016 | 2017 | 2018 |
|-------------------------|------|------|------|------|------|
| Low productivity        | 22.2 | 19.6 | 20.7 | 21.0 | 15.6 |
| Gynecological diseases  | 20.4 | 19.6 | 19.0 | 21.0 | 23.9 |
| Diseases of the extremities | 25.9 | 25.4 | 18.5 | 30.9 | 28.1 |
| Trauma                  | 7.4  | 9.8  | 6.9  | 4.9  | 6.2  |
| Udder diseases          | 20.4 | 19.6 | 19.0 | 14.8 | 17.7 |
| other diseases          | 3.7  | 5.9  | 8.6  | 7.4  | 8.3  |

Cows are often rejected on the farm due to limb
diseases. Laminitis, arthritis and sprain of limbs were
revealed. Cows with such diseases often have ovarian
hypofunction and uterine atony, and as a result the
absence of reproductive cycles. This results from the
inactivity of the animal.

For various reasons, gynecological diseases arise that
prevent estrus, ovulation and conception in the first
phase of development. The drugs used in the treatment
of ovarian hypofunction are necessary for the full
restoration of their function.

Surfagon is a hormonal drug, which contains the
active substance lutiberin acetate. It acts on the cells of
the anterior pituitary gland, quickly reaching the
maximum concentration in the blood. It is necessary to
accelerate the maturation of follicles in the ovaries.

Fertagon is a hormonal drug containing the
composition such active substances as dalargin and
alarelin acetate. Dalargin weakens the effect of stress on
the body, and meanwhile, alarelin acetate stimulates
follicular maturation and ovulation.

Estrofan is a prostaglandin that has a luteolytic effect
on the corpus luteum of the ovaries, thereby reducing the
amount of progesterone, which inhibits the growth of
follicles.

Drugs exert their effect directly on the growth and
development of follicles, while the effect on the uterus is
very low or absent altogether. Due to the fact that
postpartum complications, endometritis, atony and
uterine subinvolution are a common cause of ovarian
hypofunction, it is necessary to prevent these causes of
the disease. Analysis of table 1 confirms the presence of
these diseases.

There are problems on the farm: retention of the
placenta, endometritis, subinvolution, atony of the
uterus. On average, the retention of the placenta in a cow
lasts 5–7 days and then when the cervix is opened, it is
removed and treatment of endometritis begins.

In order to prevent retention of the placenta 6 hours
after calving, cows are injected, depending on body
weight, 2–3 ml of estrofan. To improve the recovery
period, cows are fed energy shakes. In addition, in the
first days after calving, cows are fortified with tetranavit in a dose of 10 ml.

Immediately after removal of the placenta, the cows are given an intruterine foam-forming tablet called «Enroflon», they are repeated every other day, and then treatment of the catarrhal-purulent endometritis is started by introducing the drug «Floxavet» into the uterine cavity, this procedure is repeated 3 times every other day.

Thus, the cow that has undergone this course of treatment in full by the 20th day after calving, recovered in 80 % of cases. The remaining animals in the amount of 20 % more often go into the chronic form of the course of endometritis. Of the obvious forms of endometritis, purulent-catarrhal prevails. It manifests itself with abundant purulent efflux from the genitals.

For the treatment of latent forms of uterine diseases, drugs are needed that provide a therapeutic effect, as well as increase its tone and contribute to the full passage of all stages of the sexual cycle.

For this experiment, drugs acting on the myometrium were selected: «Metrostim» and «Metrilong».

Metrostim, which includes carbacholine, briefly enhances the contraction of the smooth muscles of the uterus. The metrilong containing propanolol in the composition promotes the manifestation of the activity of endogenous oxytocin, thereby exerting a contracting effect on the smooth muscles of the uterus. Chitosan and glycercin, which are part of it, have a prolonging effect of propanolol.

Obstetric and gynecological medical examination is carried out on 15, 30 and 45 days after calving, in order to timely detect diseases of the reproductive system and conduct fruitful insemination the first time. Insemination of cows is carried out no earlier than 60 days after calving. The heifers are inseminated at the age of 15 months, upon reaching a live weight of not less than 360 kg.

There is no exercise on the farm, so the restoration of cows after calving forces the use of uterine-reducing drugs for a long time. At the same time, there is no high-quality feed base, with year-round feeding of silos and haylage.

In the stall, sexual hunting is determined by changing the behaviour of the animal and the presence of outflows from the genitals. Before insemination, the spotted cow is checked rectally with an ultrasound scanner or by palpation for the presence and size of the follicle. The insemination technician conducts artificial insemination of cows in the rectocervical method, in the presence of a mature follicle. Cows are inseminated with sperm after thawing, as well as with a sexed seed, to increase the breeding stock of cows.

Thus, when developing these treatment regimens, we took into account the need for a combination of drugs for an integrated approach to treating ovarian hypofunction and increasing the reproductive ability of cows on the farm.

The production indicators of animals in the experiment are presented in table 2.

In connection with increasing the productivity of animals, problems with the reproductive ability of cows begin. This is evidenced by the annual decrease in calf yield. It is also worth noting a sharp jump in the disposal of cows from 10–12 to 15 %.

Gynecological diseases are one of the common causes of cows dropping out of the herd for several years, they occupy almost a third of all causes.

A significant number of cows are prone to ovarian disease. Sick animals are of different ages.

Table 2. Degree of damage to animals by gynecological diseases

| Groups of animals | service period (days), productivity (kg) |
|-------------------|------------------------------------------|
|                   | 3 lactation | 2 lactation | 1 lactation |
| 1                 | 110, 6815  | 180, 7358  | 235, 6102  |
| 2                 | 112, 7436  | 212, 7108  | 214, 7234  |
| 3                 | 128, 7218  | 167, 7954  | 208, 6418  |
| the control       | 114, 7087  | 187, 6984  | 228, 7123  |

Table 3. Degree of damage to animals by gynecological diseases

| Indicators          | Years             |
|---------------------|-------------------|
|                     | 2014 | 2015 | 2016 | 2017 | 2018 |
| gynecologically sick, total; heads (%) | 277 (100) | 298 (100) | 302 (100) | 372 (100) | 390 (100) |
| of which: endometritis heads (%) | 79 (28) | 85 (29) | 91 (30) | 107 (29) | 108 (28) |
| retention of membranes fetal heads (%) | 71 (26) | 74 (25) | 79 (26) | 98 (26) | 91 (23) |
| ovarian disease, heads (%) | 94 (34) | 99 (33) | 95 (32) | 124 (33) | 142 (36) |
| other gynecological diseases, heads (%) | 33 (12) | 40 (13) | 37 (12) | 43 (12) | 49 (13) |

Table 3 shows the annual growth of ovarian disease. They occupy a third of all other gynecological diseases. The most common diseases of the ovaries are hypofunction of the ovaries and follicular cysts (Table 4).

The main etiological factors that provoked the development of ovarian hypofunction are the lack of exercise in the postpartum period, and its incomplete provision, especially in the winter – stall period, as well as intensive exploitation of animals [6,7,8].

The exercise has a positive effect on the cardiovascular, respiratory, digestive, sexual and musculoskeletal systems of cows [9]. Under conditions of physical inactivity, most animals after childbirth remain with impaired sexual function and infertile [10].

Table 4. Degree of damage to cows by ovarian disease

| Ovarian Diseases / unit of measurement | 2014 | 2015 | 2016 | 2017 | 2018 |
|----------------------------------------|------|------|------|------|------|
| follicular cyst heads                  | 28   | 31   | 29   | 37   | 40   |
| luteal cyst heads                      | 13   | 16   | 17   | 20   | 21   |
| Hypofunction, heads                    | 26   | 25   | 26   | 38   | 46   |
| Atrophy of the ovaries, heads          | 2    | 1    | 4    | 3    | 3    |
| persistent corpus luteum, heads        | 21   | 26   | 19   | 26   | 32   |

3
According to the results of an obstetric-gynecological study, 24 animals with a presumptive diagnosis of ovarian hypofunction were selected. Two animals of the first, second and third lactation were selected for the study.

Ovarian therapy in the household was previously carried out with standard drugs. For the treatment of follicular cysts, surfacton was used in a dose of 5 ml for 3 days, and for the treatment of hypofunction of the ovaries, the follimage was administered once in a dose of 1000 IU, and on day 14, estrofan in a dose of 2 ml. But, due to the fact that the follimage does not give the expected results, it is necessary to develop a new treatment regimen for ovarian hypofunction.

The developed treatment regimens for ovarian hypofunction are presented in table 5. These regimens are used for different forms of ovarian hypofunction. Scheme No. 1 and No. 2 are with anestral or arrhythmic sexual cycle; scheme No. 3 is with a full and rhythmic sexual cycle. The old treatment regime was used as a control.

| Scheme №№ | 7 day cycle | 14 day cycle | 18 day cycle | 21 day cycle | lack of a hunting stage | 18 day cycle |
|------------|-------------|--------------|--------------|--------------|------------------------|--------------|
| 1          | metrostim, 2 ml, 3 times in one day | Surfagon 10 ml 3 days | estrofan 2 ml once | insemination | estrofan 2 ml once |
| 2          | metrilong, 10 ml once | Fertagon 10 ml 3 days | estrofan 2 ml once | insemination | estrofan 2 ml once |
| 3          | metrostim, 2 ml, 3 times in one day | Surfagon 10 ml 3 days | – | Surfagon, 10 ml 1 hour before insemination | – |
| the control | follimag, 1000 IU, once | estrofan 2 ml once | – | – | – |

Table 6. Ultrasound Results

| Scheme №№ | 38 days after the first insemination, heads | 38 days after the second insemination, heads | Not pregnant 60 days after insemination, heads | no sex hunt, heads |
|------------|---------------------------------|---------------------------------|---------------------------------|-----------------|
| 1          | 5                               | 1                               | –                               | 1               |
| 2          | 2                               | 2                               | 2                               | 1               |
| 3          | 5                               | –                               | 1                               | –               |

Ultrasound, in table 4, shows the positive dynamics of therapy according to schemes No. 1 and No. 3. 80 % of the tested animals according to these schemes became fruitful after the first insemination.

Gynecological diseases are very common in the reproduction of a herd of black-motley cattle. The farm "Rico-Agro" is no exception and has very high rates of incidence of cows in gynecology, and accordingly, the rates of reproduction are reduced. So from 2014 to 2018, culling for gynecological diseases progressed from 20.4 % to 23.9 %, and ovarian diseases increased from 33–34 to 36 % over the past year. Based on this, the calf yield per 100 cows decreased from 88 to 85 %.

The main reasons for the decrease in reproductive ability are inflammatory processes of the genitals, lack of exercise, high productivity, lack of trace elements, violation of animal health standards, unbalanced diets, which violates metabolism, lack of energy, and a long stall period.

Thus, using treatment regimens No. 1 and 3, which include drugs such as Metrostim, Surfagon and Estrofan, we found high rates of cow fertilization from the first insemination. This indicator was 80 %. The use of new treatment regimens will help the farm to bring reproduction to a new level, with an intensive increase in indicators for artificial insemination of cows, a reduction in the service period and the insemination index.

4 Conclusion

High rates of milk productivity are increasingly accompanied by a malfunction of the reproductive system of cattle. This problem reduces the economic performance of production. Diseases of the ovaries and uterus are massive. Young cows are heavily rejected for cost-effective use. Ovarian hypofunction (Hypofunctio ovarii) is a dysfunctional disease characterized by a decrease in the hormonal activity of the ovaries, in which inferior reproductive cycles are observed, including the reactive and anestral forms. Complex therapy in the initial stage of the disease against the background of balanced feeding and improvement of the living conditions has a beneficial therapeutic effect. As a result of the experiment, positive dynamics with identical results were obtained using scheme No. 1 in the reactive cycle and scheme No. 3 in the anestral cycle. In general, fertilization occurred during the first insemination in 5 out of 6 cows. The preparation Metrostim (the active substance is carbacholine) has a restoring effect on the myometrium, and Surfagon, a synthetic analogue of GnRH, has a gonadotrophic effect.

References

1. L.V. Madoz, M.J. Giuliodori, A.L. Migliorisi, M. Jaureguiberry, R.L. de la Sota, Endometrial cytology, biopsy, and bacteriology for the diagnosis of subclinical endometritis in grazing dairy cows, J. Dairy Sci., 97(1), 195–201 (2014)
2. J. Stevenson, Eleven truths about ovarian cysts, Hoard’s Dairyman, The National Dairy Farm Magazine, 21 (January 2012)
3. A.M. Batista, D.M. Silva, M.J. Rego, F.L. Silva, E.C. Silva, *The expression and localization of leptin and its receptor in goat ovarian follicles*, Anim. Reprod. Sci., **141**, 142–147 (2013)

4. D. Scheetz, K. Joseph Forger, G.W. Smith, *Granulosa cells are refractory to FSH action in individuals with a low antral follicle count*, Reprod. Fertil. and Developm., **24**, 327–336 (2012) DOI: 10.1071/RD11020

5. F.O. Ahmed, A.S. Elsheikh, *Uterine bacterial infection during postpartum delays the recrudescence of the reproductive traits in dairy cows*, J. of Americ. Sci., **9**(6), 596 (2013)

6. N. Forde, M.E. Beltman, P. Lonergan, M. Diskin, J.F. Roche, *Oestrous cycles in Bos Taurus cattle*, Anim. Reprod. Sci., **124**, 163–169 (2011)

7. D. Vuković, B. Stančić, A. Božić, *Review of the dairy cows herd efficiency based on reproductive parameters*, in Proc. of 23rd Int. Symp. “New technologies in contemporary animal production,” 69–71 (Novi Sad, Serbia, 19–21 June 2013)

8. I.M. Sheldon, E.J. Williams, A.N.A. Miller, D. M. Nash, S. Herath, *Uterine diseases in cattle after parturition*, The Veter. J., **176**(1), 115–121 (2008)

9. W. Senosy, M. Uchiza, N. Tameoka, Y. Izaike, T. Osawa, *Impact of ovarian and uterine conditions on some diagnostic tests output of endometritis in postpartum high–yielding dairy cows*, Reprod. in domestic anim., **46**, 805 (2011)

10. F. Selami, G. Selami, *Preliminary survey of the administration of PGF2Α in postpartum dairy cows*, The Alban. J. of agricult. sci., **11**(4), 229 (2012)