

1 2 **An Over View of Dermatophytosis in Rabbits**

3 **Abstract:**

4 Dermatophytosis is a fungal infection of the skin caused by dermatophytes-filamentous fungi
5 which have ability to invade the epidermis and keratinized structure derived from it such as hair
6 or nails. Rabbits are one of dermatophytes host; young rabbit below 12 months of age were more
7 frequently affected with the disease. *T. mentagrophyte*'s is the most common dermatophytes
8 isolated species. The disease can be diagnosed by direct examination, fungal culture, skin biopsy
9 sero and molecular diagnosis methods. This overview forecast more light of the different aspects
10 of this disease.

11 **Key words:**

12 Dermatophytosis,Rabbit,Clinical feature, Diagnosis.Treatment

13 **Introduction:**

14 Rabbits are calm by nature. They are prone to many bacterial, fungal or parasitic skin diseases if
15 proper care is not taken. Among them dermatophytosis is one of the most common diseases [1].
16 Dermatophytosis is a superficial cutaneous infection with one or more of the fungal species in
17 the keratinophilic genera *Microsporum*, *Trichophyton*, or *Epidermophyton* [2, 3]. Young or
18 immune compromised rabbits are most susceptible to the disease [4]. Dermatophytosis is a
19 zoonotic disease so it has important implications in public health [5]. Infection with
20 dermatophytosis can occurred in receptive hosts via arthrospores present on the hair coats of
21 infected animals or in the environment [6] .

22 23 **Epidemiology:**

24 The possibility of infection of dermatophytosis depends on fungal species, host age,
25 immunocompetence, condition of exposed skin surfaces, host grooming behavior, and nutritional
26 status [6,7]
27 Young below 12 months of age or immune compromised rabbits are thought to be most
28 susceptible. The susceptibility of young rabbit to dermatophytes could be as a result of immunity not fully
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31 developed [8]. However, differences in skin secretions, especially lower levels of fungistatic
32 fatty acids in sebum and lower levels of fungal inhibitory sphingosine, and the fast growth and
33 replacement of hair may also play a role in facilitating infection [9]. The presence of
34 ectoparasites, especially fleas and *Cheyletiella* mites, can also lead to spread of dermatophytosis
35 [9].

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37 **Risk factors:**

- 38 1- Young animal
39 2- Overcrowding
40 3- high humidity
41 4- poor sanitation
42 5- malnutrition
43 6- Immunosuppression (including immunosuppressive treatment)
44 7- Injury by ectoparasites or scratches due to pruritus
45 Reported by [10]

46 **Transmission**

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48 Dermatophytosis can be transmitted by direct or indirect contact with infected hair, scales or
49 materials. Infectious microconidia in the environment or on fomites can persist for many months.
50 The pathogenesis of dermatophytosis includes several stages; adhesion, germination, invasion,
51 penetration. Natural defences against dermatophytes depend on both immunological and non-
52 immunological mechanisms so infectious microconidia must first overcome a couple of local
53 defenses to be able to adhere the keratinized tissue, the stratum corneum [11,12].

54
55 [11].

56 **Clinical features:**

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59 Clinically, dermatophytes infect the epidermis and adhering structures, including hair follicles
60 and shafts [13, 14]. In rabbits dermatophytic infections may cause alopecia, redness scaly and
61 scurf localized mainly on the face, head, auricles, and dorsal area of the neck (fig. 1-3) [15-17].

62 This disease can also result in rabbit malnutrition, growth retardation, feed remuneration
63 reduction and even death.

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69 **Fig.1: Alopecia in rabbit mouth. [17]**
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Fig.2: Localized skin lesion of dermatophytes on rabbit leg [25]



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Fig.3: Numerous canary crusts and large scales were present on the head of rabbits with hair loss.[21]

82 **Etiology:**
83 *T. mentagrophyte*'s is the most common dermatophytes isolated from rabbits and some
84 researchers consider rabbits as asymptomatic carriers of this organism [18-22]. *M.gypseum* was
85 also isolated from rabbits [23]. *M.canis* was reported by [24]. *T.verrucosum*. *Arthroderma*
86 *benhamiae* were also recorded [25]. Rabbits are reported to be carrier for dermatophytes [26] so
87 isolated *T. mentagrophyte*'s, *M.gypseum*, *M.nanum* and *M.canis* from healthy rabbits [27]

88 **Diagnosis:**

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90 Hair and scrapings samples from rabbit suspected of dermatophytes infection are collected on the
91 basis of gross lesions. Samples can be kept in polyethylene bags. [28]

92 **Direct examination:**

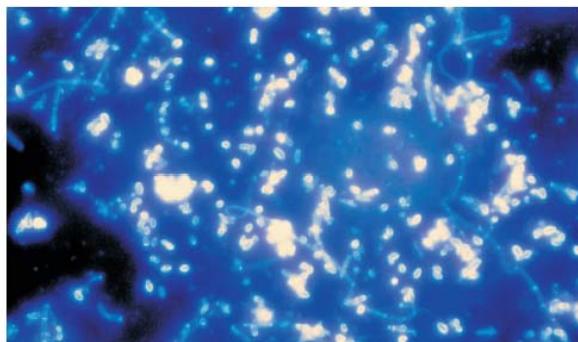
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94 Hairs and scraping samples can be mounted in potassium hydroxide (KOH) of varying
95 concentrations [29-31]. Infected hairs appear pale, wide and filamentous compared with normal
96 hairs when microscopically examined at x4 or x10 magnification. Arthrospores can be visible on
97 high magnification (x40) (fig. 4). Positive result of KOH direct test can lead to positive cultures,
98 which are considered as the gold standard. Calcofluor white (a textile brightener) can be used as
99 an alternative to KOH because it binds specifically to the fungal cell wall and fluoresces strongly
100 when viewed under a fluorescence microscope [27] as shown in figures 5and 6.



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104 **Fig. 4: Ectothrix arthrospores infection in hair**

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108 **Fig.5: Fluorescent microscopy (calcofluor white stain) of *Trichophyton mentagrophytes***

109 **complex hyphae and conidia isolated from healthy rabbits [27]**

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115 **Fig. 6: Fluorescent microscopy (calcofluor white stain) of *Microsporum gypseum* hyphae**

116 **and macroconidia isolated from healthy rabbits [27]**

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119 **Fungal culture:**

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121 Fungal culture is considered the ‘gold standard’ for diagnosis [32]. Sabouraud’s dextrose agar
122 (SDA) containing cycloheximide, penicillin and streptomycin were used in most diagnostic
123 laboratories. Plates should be incubated at 25°C for 5 weeks. Dermatophytes test media (DTM) is
124 recommended as the best media for isolation of dermatophytes because the presence of the red
125 color indicated positive result, this can help in early identification of highly suspected cultures
126 [33]. The isolates should be examined macroscopically and microscopically after staining with
127 lactophenol cotton blue using wet mount technique [34](fig. 7 - 14)

128 In addition to technique steps mentioned above, pigment production on corn meal agar, urease
129 activity on urea agar base (fig.15), growth at 37°C on SDA.

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132 **Fig.7: Culture of *T. mentagrophytes*: surface of colony show powder-like shape, white, loose**
133 **irregular mycelium on the edge.[21]**

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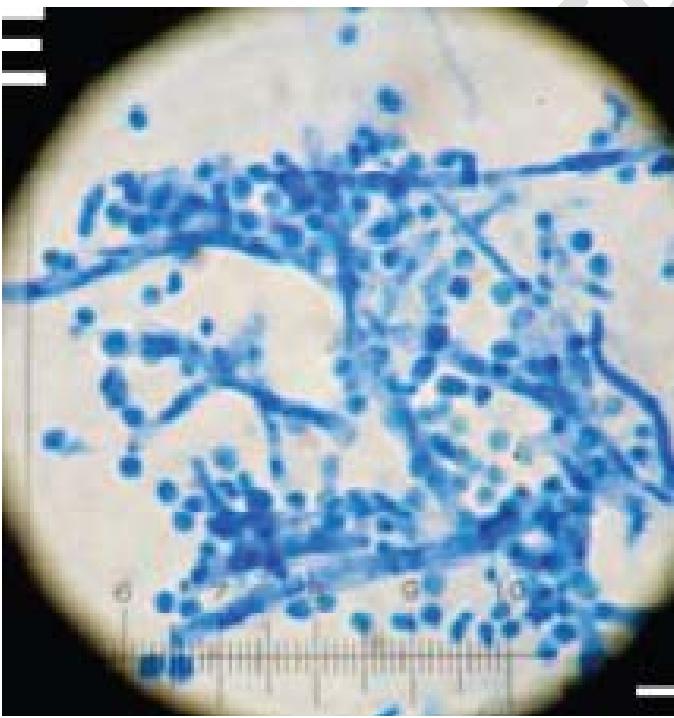
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141 **Fig.8: *Microsporum canis* culture, macroscopic colony[50]**

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Fig.9: *Microsporum gypseum* culture, macroscopic colony[50]



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Fig.10: Small conidia (size: 2-3×2-4 μm) and mycelium of *T. mentagrophytes* [21]



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150 Fig.11:Spiral hyphae of *T.mentogrophytes* var *mentogrophytes* Slide stained with
151 LPCB stain. [42]

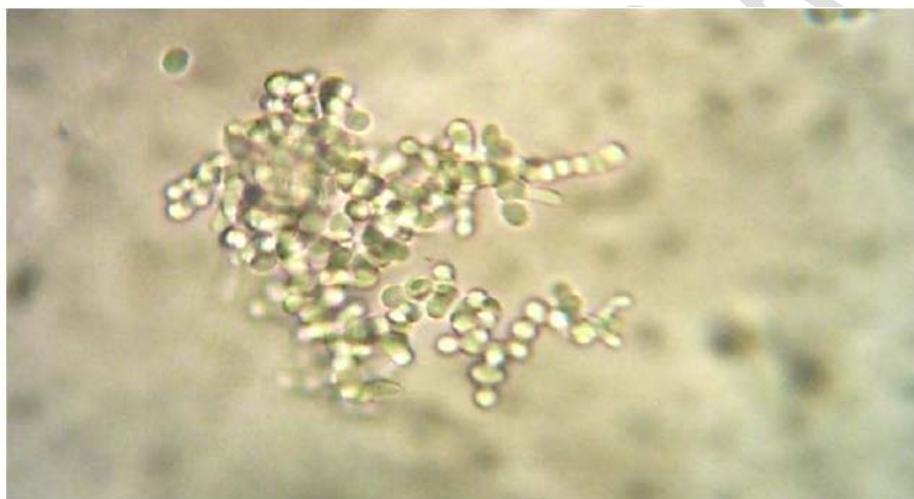
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157 Fig.12: *Microsporum canis* microscopic observation in lactophenol cotton blue[50]
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161 Fig.13: *Microsporum gypseum* microscopic observation in lactophenol cotton blue [50]
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164 Fig.14: Lactophenol cotton blue mount shows chains of chlamydospore of *Trichophyton*
165 *verrucosum* culture incubated at 37°C. [25]
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170 **Fig.15: Growth of *T.mentagrophytes* on urea agar after 4 days showing hydrolysis of the**
171 **urea. [42]**

172 **Molecular diagnosis:**

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174 Diagnosis with conventional methods is time-consuming because it might take up to 4 weeks or
175 longer to give the final results [35]. Furthermore, morphological identification may be confusing
176 due to polymorphism of dermatophytes [36]. During the last decade, a wide variety of molecular
177 techniques has become available as possible alternatives for routine identification of fungi in
178 clinical microbiology laboratories [37, 38]. *T. mentagrophytes* isolated from nine rabbits and
179 three farm staff were identified by using amplification of CHS-1 gene and ITS+ sequence. The
180 results of sequences of CHS-1 and ITS from different DNA samples revealed that they were
181 identical [21].

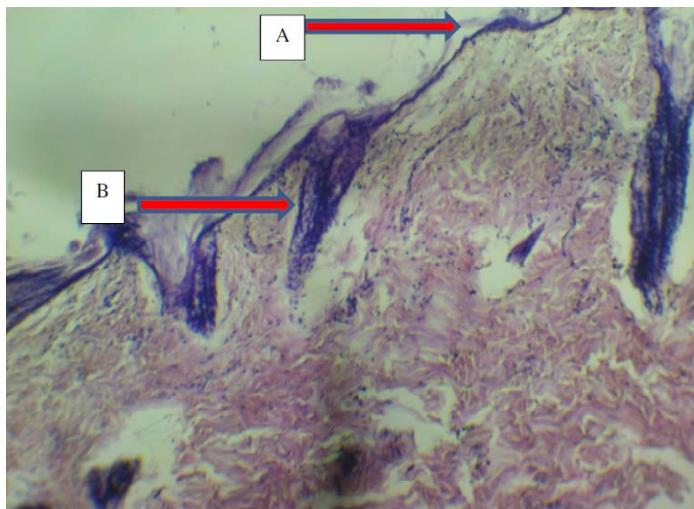
182 **Serodiagnosis:**

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184 Indirect ELISA tests developed to detect specific IgG in rabbits infected with *T. mentagrophytes*,
185 found that (ELISA-rabbits test) is highly sensitive (96.0 %) and highly specific (94.1 %) [39].

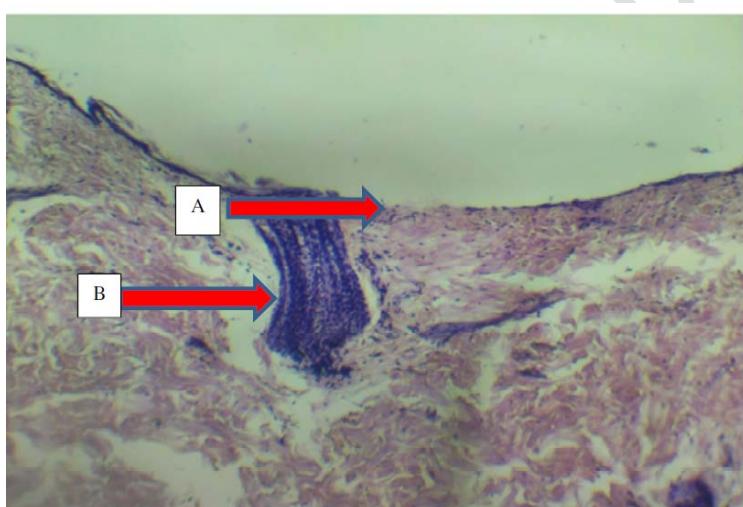
186 **Skin biopsy:**

187 Skin biopsy from rabbit infected with *T. mentagrophytes* showed pathological changes with
188 adherence of fungus to keratinocytes, through the stratum granulosum of the epidermis. In this
189 period of infection there was a hyperkeratosis , thickening of epidermis with hair follicle
190 plugging in addition to keratinized squamous epithelial lining with underlying moderate
191 periappendageal tissue and perivascular chronic inflammatory cells infiltration (lymphocytes)

192 In 8-10 days of induced infection there is keratinized squamous epithelial lining with focal area
193 of surface erosion and underlying moderate periappendageal tissue chronic inflammatory cells
194 infiltration (lymphocytes)



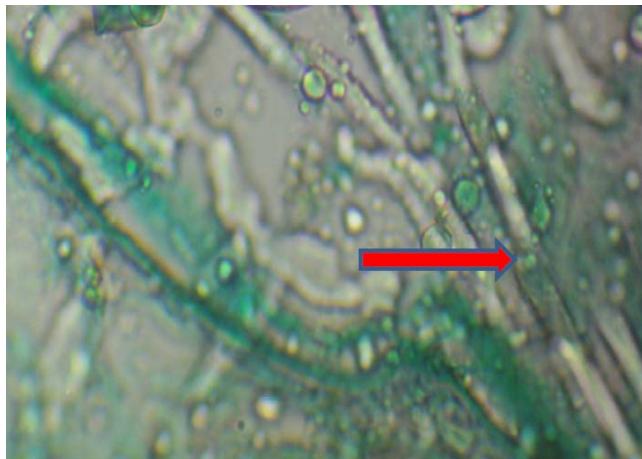
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196 **Fig.16: The bland looking (A) Hyperkeratosis , thickening of epidermis with (B) hair**
197 **follicle plugging in 4-5 days (stained with H&E,10 X)[40]**



199
200 **Fig.17: The bland looking with A- surface erosion and B- lymphocytes infiltration in**
201 **8-10 days (stained with H&E,10 X)[40]**

202 haematoxylin and eosin staining (H&E) may or may not identify dermatophytes and special
203 stains such as periodic acid Schiff (PAS) and Grocott methenamine silver (GMS) are needed. The
204 epidermis infiltrated with variable fungal septate hyphae in size in the surface of the squamous
205 epithelium [40] fig. 16-18.

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210 **Fig.18: Proliferation of septate hyphae of *T. mentagrophytes* in epidermis in 8-10 days**
211 **(stained with PAS,40X).[40]**

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213 **Treatment:**

214 Optimal therapy of dermatophytosis requires a combination of topical antifungal therapy,
215 concurrent systemic antifungal therapy and environmental decontamination. The treatment
216 should be continued until two consecutive negative cultures (at weekly or bi-weekly intervals)
217 are obtained [41]. Topical treatments speed resolution of clinical lesions and may help prevent
218 zoonotic contagion. Systemic therapies that have prolonged residual activity in the skin and hair
219 provide the most effective treatments.

220 **Topical Therapy:**

- 221 1. nystatin ointment for treatment of rabbit experimentally infected with *T. mentagrophytes* for 3
222 weeks [42].
223 2. Clotrimazole is well-documented antifungal agent for treatment of rabbits [43].
224 3. 0.12g of terbinaphine 1% cream, for 28 days [44].

225
226 **Systemic Therapy:**

- 227 1. Griseofulvin 25–30 mg/kg during 5–6 weeks. Avoid its use in pregnant animals [45,46].
228 2. Itraconazole 5-10 mg/kg daily, for 1 month [47].

229 **Environmental decontamination:**

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231 Enilconazole emulsifiable concentrate will be sprayed onto the walls and ceiling of rabbit house
232 (50 mg per m²) twice weekly for 23 weeks. Treated farm showed reduction of number of
233 clinically infected rabbits [48].

234 **Vaccination:**
235 It is a dried culture of an attenuated strain of *T. mentagrophytes*. It has a high immunogenic
236 activity for dermatophytosis in rabbits. The vaccine is non-reactogenic and is injected
237 intramuscularly. The vaccine has been recommended for practical use in USSR [49].

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239 **Conclusion:**
240 Dermatophytoses are the most common fungal infections in rabbits. Many studies were done
241 considering different aspects of the disease (eg. epidemiology, clinical presentation and
242 diagnosis, treatment, prevention, and control). As many rabbits share the environment with
243 owners as companion animal so they become a source of infection to human this can lead to
244 public health problem.

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246 Ethical: NA

247 Consent: NA

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UNDER PEER REVIEW