1	Decolonising Parasitology: The Case of Trypanosoma
2	brucei rhodesiense
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17	Abstract
18	Trypanosoma brucei rhodesiense was named after Rhodesia which in turn was named
19	after the British imperialist and white supremacist Cecil Rhodes. In the light of the
20	Black Lives Matter movement and contemporary consciousness of post-colonial

legacy, it seems opportune to reconsider the subspecies name. Pros and cons of

renaming *T. b. rhodesiense* are discussed.

Many tropical parasite species were discovered and described at the turn of the 20th 24 century. Most of these research efforts were driven by colonial powers (but not 25 exclusively, e.g. the pioneering work of Oswaldo Cruz (1879-1934) and Carlos Chagas 26 (1872-1917) on Chagas disease in Brazil). Cures for diseases caused by parasites 27 were sought in order to protect military and civilian personnel working in the colonies, 28 and to sustain the native people who provided the labour and local administration 29 needed to maintain a colonial economy and to maximise the economic return. Most of 30 the investigations were carried out by colonial military personnel and by Europe-based 31 scientists. A frequent practice in naming newly discovered parasite species was (and 32 still is) to name them after their appearance, disease characteristics, geographical 33 regions where they were first discovered, discoverers, or in honour of distinguished 34 personalities. 35

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37 The Discovery of Trypanosoma (brucei) rhodesiense

The identification of the causative agent of human African trypanosomiasis or sleeping 38 sickness is a story of inter-colonial and international collaboration but also a story of 39 serendipity and rivalry [1]. The species Trypanosoma brucei was discovered as a 40 causative agent of nagana disease or cattle trypanosomiasis by the Scottish 41 pathologist and microbiologist David Bruce (1855–1931) in 1895 while working for the 42 Army Medical Service [2]. The first unequivocal report of trypanosomes in the blood of 43 a human came from the British Colonial surgeon Robert Michael Forde (1861–1948) 44 in 1901 when examining a steamboat captain in The Gambia [2]. However, he thought 45 it was a worm. It was the English physician Joseph Everett Dutton (1874–1905) who 46 identified the organisms as trypanosomes a few months later and proposed the name 47 *Trypanosoma gambiense* in 1902 [3]. In 1910, the British parasitologists John William 48 Watson Stephens (1865–1946) and Harold Benjamin Fantham (1876–1937) 49 described a trypanosome organism obtained from an English sleeping sickness 50 patient who got infected in the Luangwa Valley (between Mzaza and Feira) in former 51 52 North-Eastern Rhodesia in September 1909 (Figure 1) [4]. At that time the Luangwa

Valley was known to be heavily infested with *Glossina morsitans*, a tsetse fly species 53 unknown in transmitting sleeping sickness in those days [4]. Stephens and Fantham 54 distinguished this newly discovered trypanosome organism from T. gambiense on the 55 basis of morphological differences [4]. As they thought they were dealing with a new 56 trypanosome species, they proposed the name *Trypanosoma rhodesiense*.[4]. Today, 57 T. gambiense and T. rhodesiense are referred to as subspecies of T. brucei and have 58 been termed T. brucei gambiense and T. brucei rhodesiense, although the 59 subspecies/species status of the latter is controversial (see **Box 1** and below) [2,5]. 60

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62 *T. b. rhodesiense* and Its Eponymous Association with Cecil Rhodes

The naming of T. b. rhodesiense followed that of T. b. gambiense, which was named 63 after the region where the first case of human infection with this parasite occurred, The 64 Gambia [3]. Accordingly, as the first reported case of sleeping sickness caused by T. 65 *b. rhodesiense* was from Rhodesia, the species was named after this historical region 66 in southern Africa [4]. This territory was demarcated by the British South Africa 67 which comprised three protectorates, Company, North-Eastern Rhodesia, 68 Barotseland-North-Western Rhodesia, and Southern Rhodesia (Figure 1). North-69 Eastern Rhodesia and Barotseland-North-Western Rhodesia were amalgamated in 70 1911 to form Northern Rhodesia, which has been Zambia since 1964. The name 71 Rhodesia was first used by white settlers in the 1890s who informally called their new 72 home after the British mining magnate and politician Cecil John Rhodes (1853-1902), 73 the founder and managing director of the British South Africa Company. In 1895 the 74 British South Africa Company adopted the name Rhodesia for this southern African 75 territory and the British government followed suite officially in 1898. 76

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Cecil Rhodes was an ardent imperialist and believed in the supremacy of the "*English race*". This racial attitude is confirmed in a letter of 1877 which Rhodes wrote when he was about 22 years old [6]. He also advocated vigorous settler colonialism and was an integral participant in southern African and British imperial policy [7,8]. Cecil

Rhodes was always a controversial figure, but since the 1950s, opposition to his 82 memorials has been escalating. Particularly in South Africa, protesters have 83 demanded his monuments be taken down. In recent years, the Black Lives Matter 84 movement has given fresh impetus to the demand to remove memorials of Cecil 85 Rhodes. For example, in June 2020, the governing body of Oxford Oriel College 86 bowed to student pressure and voted to remove a Rhodes statue from the facade of 87 the college^{*i*}. In response to the Black Lives Matter protests against institutional racism, 88 the Rhodes Art Complex in Bishop's Stortford, Hertfordshire, England, was changed 89 to South Mill Arts in August 2020 ⁱⁱ. In this context, it seems timely to re-evaluate the 90 subspecies name T. b. rhodesiense. 91

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93 Renaming *T. b. rhodesiense*: What Are the Options?

There are three options to deal with the controversial subspecies epithet "*rhodesiense*". The first option is to leave it as it is. The second option is to revoke the subspecies status of *T. b. rhodesiense* while the third option is to give *T. b. rhodesiense* a different subspecies epithet. In the following paragraphs, the pros and cons of the second and third options are discussed.

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The second option would be in agreement with previous molecular and genetic studies 100 (reviewed in [9,10]) and phylogenetic relationship analysis [11], which all concluded 101 that T. b. rhodesiense is only a phenotypic variant of T. b. brucei while T. b. gambiense 102 type 1 constitutes a valid (sub)species of T. brucei. The only characteristic that 103 distinguishes T. b. rhodesiense from T. b. brucei is the ability of the former to infect 104 humans. The human infectivity of T. b. rhodesiense is associated with a single gene, 105 the serum resistance associated (SRA) gene [12,13]. Indeed, it has been 106 demonstrated that transferring the SRA gene into T. b. brucei was alone sufficient to 107 confer resistance to human serum [13]. An accidental laboratory infection with a T. b. 108 *brucei* strain expressing the SRA gene proved unequivocally that human infectivity of 109 *T. b. rhodesiense* is solely based on this gene [14]. In addition, it has been shown that 110

it is possible to experimentally cross T. b. brucei and T. b. rhodesiense in the laboratory 111 (reviewed in [15]). Some of the hybrid clones acquired copies of the SRA gene and 112 were resistant to lysis by human serum, indicating that they inherited the human 113 infectivity phenotype [15]. Furthermore, population genetics studies have evidenced 114 that there is gene flow between T. b. brucei and T. b. rhodesiense [15]. These studies 115 all strongly suggest that T. b. rhodesiense is not a valid subspecies but just a host 116 range variant of T. b. brucei. Revoking the subspecies status of T. b. rhodesiense 117 leaves the phylogenetically distinct T. (b.) brucei and T. (b.) gambiense as the two 118 aetiologic agents of human African trypanosomiasis and ameliorates the need for a 119 subspecies nomenclature. The major consequence of doing so is that human infective 120 isolates from surveillance of livestock and tsetse will not be differentiated for human 121 infectivity by their nomenclature unless a suffix (such as SRA+/-) is also adopted. It 122 has the advantage that as well as being more socially acceptable *T. brucei* SRA+ is 123 also more accurate scientifically than T. b. rhodesiense. Abolishing the subspecies 124 status of *T. b. rhodesiense* may also cause practical problems in everyday laboratory 125 handling of this parasite. For example, the risk of a mix-up between non-human 126 pathogenic and human pathogenic variants of *T. b. brucei* due to labelling errors may 127 increase. In addition, T. b. brucei and T. b. rhodesiense are usually assigned to 128 different hazard groups iii, with T. b. rhodesiense being classified as a biological agent 129 that can cause serious disease in humans. The classification of a biological agent in 130 two different hazard groups may cause its own problems with respect to risk 131 assessment and application of appropriate control measures. However, the 132 assignment of a biological agent that includes both pathogenic and non-pathogenic 133 strains in different hazard groups is not unprecedented. For example, the non-134 pathogenic Escherichia coli laboratory strain K-12 is usually assigned to Hazard Group 135 1 while the verocytotoxigenic *E. coli* strain O157:H7 is classified into Hazard Group 3 136 iii. Revoking the subspecies status of *T. b. rhodesiense* could also raise concerns by 137 clinicians, particularly if the "brucei" epithets were retained as it is crucial for them to 138 know with which human-pathogenic *T. brucei* subspecies a patient is infected. This 139

knowledge is important as different drug regimens are used for the treatment of East 140 African and West African sleeping sickness caused by T. b. rhodesiense (then T. 141 brucei) and T. b. gambiense, respectively, so it may be wise to readopt the use of T. 142 gambiense. However, in either case, revoking the subspecies status of T. b. 143 rhodesiense may not cause any problems for the treatment of sleeping sickness 144 patients as both forms of the disease have distinct geographical distribution with 145 Uganda being the only country in which both forms of sleeping sickness co-occur but 146 in different regions without overlapping. 147

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For the third option one would need to find a suitable replacement for the subspecies 149 epithet "rhodesiense". This should follow the previous naming of the human 150 pathogenic subspecies of *T. brucei*, which was according to the locations of the first 151 reported cases of infection. As the first documented case of *T. b. rhodesiense* infection 152 was from North-East Rhodesia (Figure 1) [4], which since 1964 is part of present-day 153 Zambia, the logical choice would be to rename the subspecies as *T. b. zambiense*. As 154 the three British South Africa Company protectorates, North-Eastern Rhodesia, 155 Barotseland-North-Western Rhodesia, and Southern Rhodesia were initially 156 collectively known as Zambesia ^{iv}, *T. b. rhodesiense* could alternatively also be 157 renamed as T. b. zambesiense. An obstacle to the renaming is the law of priority, 158 which is a basic principle of the International Code of Zoological Nomenclature ^v. This 159 law states that "the valid name of a taxon is the oldest available name applied to it" v. 160 However, a name can be invalidated by any ruling of the International Commission on 161 Zoological Nomenclature ^v. 162

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Whatever the decision may be, the major obstacle of all remains the more than 100
years of literature published on *T. b. rhodesiense* and East African sleeping sickness.
It is easy to remove a statue from public view but it is impossible to erase a species
name from the scientific record.

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- 172

173 **Resources**

- ⁱ https://www.bbc.co.uk/news/education-53487991
- ⁱⁱ https://www.independent.co.uk/news/uk/home-news/cecil-rhodes-arts-complex-
- theatre-name-change-black-lives-matter-protests-a9655221.html
- ¹⁷⁷ ⁱⁱⁱ https://www.hse.gov.uk/pubns/misc208.pdf
- ¹⁷⁸ ^{iv} https://www.britannica.com/place/Zimbabwe/The-British-South-African-Company
- 179 vhttps://iczn.org/the-code/the-international-code-of-zoological-nomeclature/
- 180 ^{vi} <u>https://commons.wikipedia.org/wiki/File:Rhodesia_map_EB1911.png</u>
- 181 vii https://commons.wikipedia.org/wiki/File:Zanbia_Zimbabwe_Locator.png
- 182

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Box 1. How the three subspecies nomenclature became adopted

In the early years of parasitological research, species discovery and identification 224 relied mainly on the description of morphological and biological features. Accordingly, 225 the discovery of sleeping sickness trypanosomes was solely based on morphological 226 description [3,4]. However, when *T. rhodesiense* was discovered, it was thought by 227 some (the so-called "British" school including Bruce himself) that this trypanosome 228 was identical to T. brucei, because the two species were morphological 229 indistinguishable, and both were transmitted by the same group of tsetse flies (G. 230 morsitans group) and showed equal virulence to animals [9,10]. In contrast, others 231 thought (the so-called "German" school) that T. rhodesiense and T. brucei were 232 distinct species mainly based on the fact that isolates of *T. brucei* were non-infectious 233 to humans [9,10]. Further evidence for *T. rhodesiense* being a distinct species came 234 from the Tinde experiment, which showed that a strain of *T. rhodesiense* did not lose 235 its human infectivity after prolonged serial cyclical passages (23 years) through tsetse 236 flies and sheep [9,10]. However, human infectivity of *T. rhodesiense* and non-human 237 infectivity of *T. brucei* were shown not to be absolute characteristics of the two species 238 [9]. Eventually, the British protozoologist and parasitologist Cecil Arthur Hoare (1892-239 1984) demoted both sleeping sickness trypanosome species to subspecies of T. 240 brucei [9,10]. 241

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Figure 1. Map of historical Rhodesia. The protectorates North-Eastern Rhodesia and North-Western Rhodesia, which were amalgamated into Northern Rhodesia in 1911 (Zambia since 1964), are shown in green. The protectorate Southern Rhodesia (Zimbabwe since 1980) is shown in orange. The region where the first documented case of a human *T. b. rhodesiense* infection presumably occurred is highlighted in red. The map has been created by merging a 1911 Encyclopædia Britannica illustration ^{vi} and a map indicating the location of Zambia and Zimbabwe ^{vii}.

