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Health-Related Quality of Life of Tuberculosis Patients During the COVID-19 Pandemic in Conakry, Guinea: A Mixed Methods Study

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Abstract: The COVID-19 pandemic has had a significant impact on all facets of life and has exacerbated many challenges faced by people living with tuberculosis (TB). This study aimed to assess the health-related quality of life (HRQoL) of TB patients in Guinea during the COVID-19 pandemic. A mixed-methods study was conducted using two validated psychometric tools to assess HRQoL and qualitative interviews among TB patients enrolled in treatment at three centers in Conakry, Guinea. Multinomial logistic regression was used to identify factors associated with the deterioration of HRQoL. We included 439 participants in the study, among whom 44% and 31% experienced pain and anxiety, respectively. We found that an increase in the number of household members and the distance from participants' residence to the health centre were significantly associated with lower HRQoL. Qualitative interviews highlighted nutritional and financial concerns which were exacerbated during COVID-19 pandemic and beliefs that the Guinean Government's assistance plan was insufficient. This study supports the implementation of specific relief plans for TB patients which includes nutritional and psychological support, especially those whose movements are limited by travel restrictions, impeding access to TB care, reducing work opportunities and exacerbating financial needs and stress.

Keywords: Tuberculosis; Health-related quality of life; COVID-19; Guinea

1. Introduction

Tuberculosis (TB) is one of the leading causes of death due to an infectious disease in low and middle-income countries [1]. The World Health Organization (WHO) estimates that 10 million people fell ill with TB in 2019, including 1.4 million deaths [1]. According to the Guinean National Program of Tuberculosis (NTP), the estimated number of notified cases was 122 in 2020 per 100,000 inhabitants, with a treatment success rate of 72% [2]. However, this notification rate is lower than WHO national estimates of 176 cases per 100,000 inhabitants, suggesting ongoing challenges with TB screening and detection in Guinea [2]. Such challenges are driven by factors including low coverage of TB services, human resources shortages, and TB notification among children and loss to follow up

among patients [1]. For those who are diagnosed, the successful management of TB patients in Guinea is constrained by the burden imposed by the principle of the directly observed treatment (DOT), requiring daily attendance at TB clinics, and the long duration of TB treatment [3,4], as well as patient characteristics such as experiences of stigma and psychological and emotional dysfunction, financial problems, side effects of anti-TB drugs, HIV co-infection and the overall quality of care available to TB patients [5–7]. The emergence of COVID-19 has exacerbated many of these factors, further complicating the management and prevention of TB. According to WHO, a 25 – 30% global reduction in the number of TB cases notified and treated over three months is estimated to result in an additional 0.2 to 0.4 million additional deaths [1].

Since the beginning of the COVID-19 pandemic in Guinea in March 2020, several mitigation actions have been taken by the Guinean authorities to control the pandemic, but no specific actions have been taken which are focused on people at high risk such as TB patients, who are more likely to experience more severe negative outcomes from COVID-19 compared to non-TB patients [8]. COVID-19 and TB share similar symptoms and clinical presentation including cough and fever [9]. However, unlike TB, the global COVID-19 response has been characterized by a high-level of attention from decision-makers leading to the establishment of a rapid system and a response plan proportional to the pandemic scale [9].

The first pillar of WHO's End TB strategy emphasizes patient-centered care and prevention [10]. Health-related Quality of Life (HRQoL) relates to a perception or response to physical, social, environmental and mental factors that contribute to a normal life, and plays an important role in optimizing treatment success among TB patients [11]. Existing literature suggests that low levels of HRQoL among TB patients is associated with negative health outcome such as physical pain, anxiety, and negative therapy-treatment outcome and psycho-social aspects [12,13]. HRQoL may be influenced by the clinical evolution of the TB, with an impaired mental well-being has been associated with an early month of treatment [14]. Additionally, emerging evidence has documented the negative impacts of the overall HRQoL during the COVID-19 pandemic, particularly among the elderly [15], people who were infected by COVID-19 [16], and those with pre-existing medical conditions [17,18].

Given the impact of both TB and COVID-19 on quality of life, we hypothesize that the risk of deteriorating HRQoL among TB patients may be worse during the COVID-19 era compared to before, particularly in a context such as Guinea where there is a marked lack of support for mitigating the impact of COVID-19 among TB patients. However, little is known about the HRQoL among TB patients in Guinea either before, and more importantly during the COVID-19 era. The goal of this study was to assess the psychosocial and traumatic impacts of COVID-19 on TB patients' HRQoL in Guinea, in order to inform appropriate, mitigate and protection measures.

2. Materials and Methods

Settings

This study was conducted in Conakry, the capital city of Guinea, with 2,039,725 estimated inhabitants [2]. This study took place among TB patients recruited from 11 TB diagnosis and treatment centers (DTCs) in three municipalities within the capital city of Conakry (Matoto, Ratoma and Dixinn), the current epicenter of the COVID-19 pandemic in Guinea and where 60% of national TB cases are notified [2]. TB notifications in Conakry ranged from 378 cases per 100,000 inhabitants in 2018 to 409 cases per 100,000 inhabitants in 2019 [2].

Study framework analysis

In this study, we hypothesize that the COVID-19 pandemic has led to reductions in the reported HRQoL of TB patients in Guinea. Figure 1 shows a hypothesized problem tree outlining this hypothesis. Three key factors influencing HRQoL emerge from this tree: 1) The level of information of the population, which was not suitable to explain the risk of infections for vulnerable people; 2) The weak commitment of the Ministry of Health and civil society actors in treating TB, resulting in a lack of funding which may exacerbate bottlenecks in the treatment of TB patients. Finally, the last level of this problem tree indicates a low level of information from community media on the potential risks of COVID-19 and a low level of community involvement, which may increase the level of fear and stigma. This whole chain of causation ultimately leads to a worsening of TB patients' quality of life (Figure 1).

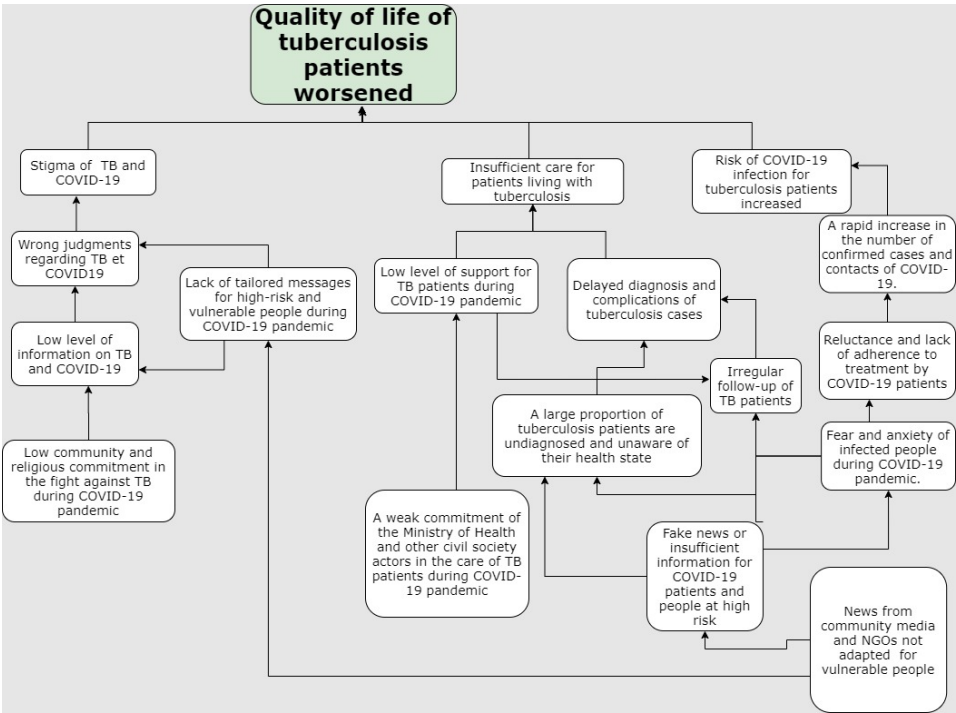


Figure 1: Hypothesized problem tree of HQoL of TB patients during the COVID-19 pandemic in Guinea.

Study design and population

We carried out a mixed methods study to explore the perception of the impact of COVID-19 and the resulting restrictive measures established by the Guinean authorities on HRQoL among TB patients. The mixed methods study comprised a quantitative, cross-sectional study and qualitative semi-structured interviews. Any patients (aged 10 to 80 years) currently receiving TB treatment at any of the 11 selected DTCs in Conakry during the study period (September 01 to September 18, 2020), and who were willing to participate in the study were eligible for inclusion. Those not consenting to participate in the study were excluded.

Study procedures and tools

Two, validated psychometric tools were used for the quantitative component. First, the Impact of Event-Revised Scale (IES-R) [19] a test of 22 difficulties that people sometimes experience following a traumatic event and is designed to measure the effect of life stress, daily trauma, and acute stress. We used the validated French version of IES-R [20]. Second, the descriptive system for HRQoL states in adults (EQ-5D-3L), designed to provide a generic and straightforward health measure for clinical and economic evaluation and measures five dimensions of health: mobility, self-care, usual activities, pain or discomfort and anxiety or depression [21]. In addition to these psychometric tools, a quantitative questionnaire was developed to collect the following patient information: age; sex (male/female); source of COVID-19 information; marital status (single, married); municipality of residence of the patients; education level (no formal education, primary, secondary, university, high school), occupation of the patient (unemployed, private employee, civil servant, freelance), number of people in the household, and the distance between the patient's home and the treatment site (in kilometres). The following TB clinical variables were also collected: date of diagnosis, type of TB (susceptible, resistant), localisation (pulmonary, extra-pulmonary), treatment history (new case, previously treated), method of diagnosis (bacteriologically confirmed, clinically diagnosed), treatment start date, type of treatment regimen (susceptible, multidrug-resistant), if multidrug-resistant, type of treatment regimen (short course, long course), duration of the appointments for anti-TB drugs supply (in days), HIV status (positive, negative), if HIV positive, antiretroviral treatment commenced (yes, no), and the length of the appointment for the supply of anti-TB drugs (in days).

For the qualitative component, a semi-structured interview guide was developed to assess patients' perceptions of COVID-19 and of the restrictive measures implemented. The interview guide explored the following themes: attitudes towards restrictive and measures, proximity to people affected by COVID-19, and the physical, psychological, social, and economic aspects (income, means of transportation). Interviews were conducted in the participant's preferred language. Investigators were trained in survey methodology, collection tools, and collection technique during a one-day training session which allowed the interviewers to master the context of the study and its objectives. All research tools were reviewed by the research team and suggestions for improvement were integrated to ensure clarity and improve the overall tool quality and coherency. A pre-test of the tools was organised to assess the feasibility of the field survey.

Sampling and recruitment

Consecutive sampling was used to recruit eligible patients into the quantitative study from the 11 DTCs study sites. Eligible patients were enrolled during their follow-up visits at each of these DTCs, with the number of patients to be enrolled at a DTC relatively proportional to the number of TB patients on treatment at the centre. The estimated sample size for the quantitative survey was based on a conservative assumption that 50% of TB patients would have a deterioration of HRQoL based on the experiences and professional opinions of the study team. With the desired accuracy of 5%, the minimum expected size was 384; because of non-response rates, this size was increased by 15% or 441 patients.

Participants for the qualitative component were purposely sampled from patients already recruited into the quantitative study. The study team aimed to recruit approximately thirty participants.

Data collection

For the quantitative component, patient follow-up agents (equivalent to community health workers) and other investigators were equipped with Android cell phones which were used to administer the questionnaires at the DTCs closest to the patient. Data were recorded through an Open Data Kit (ODK) application (XLSForm with integration into the ONA platform: <https://ona.io/home/>) and connected to a server deployed for this purpose. For the qualitative interviews, interviewers used android phones to record qualitative interviews accompanied by note-taking.

Data management and analysis

Quantitative data were entered directly into the ODK Platform and checked regularly by study team members. The setup of the variables assured good internal quality control and minimised errors and/or duplicate data. Data were carefully cleaned up before analysis. Descriptive statistics were used to summarise patient socio-demographic and clinical data captured by the questionnaires, while the analysis of psychosocial and traumatic events was carried out according to the tool used. For the EQ-5D-3L, traumatic events were calculated according to the three levels for each indicator (no problem, moderate problems, serious problems). Patients with scores corresponding to levels 2 and 3 (moderate and severe problems, respectively) were regarded as having a negative HRQOL. For the revised Event Impact Scale (IES-R), the proportions of patients reporting symptoms of mild post-traumatic stress disorder (PTSD), moderate PTSD, and severe PTSD were calculated based on sociodemographic and clinical features. All TB patients with a score equal to or less than 32 were considered to have low-stress levels, those with a score of 33-38 as moderate-stress levels and those with a score equal to or greater than 39 as severe-stress levels. Confidence intervals were built around these proportions. Chi-square test was used to compare the dependent variables of EQ-5D-3L and IES-R with sociodemographic and clinical variables. Multinomial logistic regression was used to identify significant associations between the EQ-5D-3L variables (pain/discomfort, personal care, and anxiety/depression) and the composite variable resulting from the recoding of the impact scale of the event with sociodemographic and clinical variables. A backward stepwise regression was used to build the model, using R. All the tests were considered significant, with a risk of $\alpha = 0.05$.

Recordings of the qualitative interviews were transcribed verbatim in French to facilitate data analysis. A thematic approach was used to develop a coding framework which was then applied to the transcripts. After a preliminary read of the data, two major themes emerged: the socioeconomic aspect of TB and the Government's response to COVID-19. The following sub-themes were applied to the transcripts: public perception /view of TB patients when they cough, assessment of physical and psychological health conditions, the current economic situation as well as restrictive, punitive and stimulative actions taken by the Government regarding TB patients during the COVID-19 pandemic. Coding was conducted using the package RQDA in R software.

3. Results

3.1 Quantitative findings

From August 29 to September 17, 2020 a total of 439 TB patients were included in the study, with a mean age of 35 years, and mostly male (61%). The majority of patients had drug sensitive TB (90%), while the 10% with MDR-TB were mostly on a short regimen. One quarter (24.4%) of participants were HIV positive, among whom 90% reported receiving ARV treatment. The mean distance from patients' homes to the DTC was 3km. The majority of patient's treatment started within ≤ 21 days (91%), and less than half had

completed more than three months of treatment (47.4%). Finally, the majority of our patient had more than 90 days for the supply of anti-TB drugs (Table 1).

Table 1. Socio-demographic and clinical characteristics of the study sample (n=439)

	Variable	n (%)
Sociodemographic characteristics		
Age	Children (< 18 years)	22 (5.0%)
	Young (19 – 39 years)	258 (59%)
	Adults (40 –59 years)	125 (28%)
	Elderly (> 59 years)	34 (7.7%)
Gender	Male	269 (61%)
	Female	170 (39%)
Marital status	Single	269 (61%)
	Married	170 (39%)
Residence	Matoto	168 (38%)
	Ratoma	97 (22%)
	Matam	95 (22%)
	Dixinn	15 (3.4%)
	Kaloum	32 (7.3%)
	Outside Conakry	32 (7.3%)
Level of education	None/ informal education only	133 (30%)
	Primary level (6 years)	86 (20%)
	Secondary level (10-13 years)	158 (36%)
	High school level (≥10) years	51 (12%)
	Tertiary/university level (≥14)	11 (2.5%)
Occupation type	None/ unemployed	133 (29.8%)
	Private employment	40 (9.1%)
	Civil servant/ public sector	18 (4.1%)
	Freelance/ self employed	248 (56%)
Average number of household members (range)		6 (4 – 8)
Clinical characteristics		
Tuberculosis type	Drug sensitive	394 (90%)
	Drug resistant	45 (10%)
TB disease localization	Pulmonary	354 (81%)
	Extrapulmonary	85 (19%)
TB treatment history		
	New case	390 (89%)

	Variable	n (%)
TB diagnosis type	Previously treated	49 (11%)
	Bacteriologically confirmed	379 (86%)
	Clinically diagnosed	60 (14%)
HIV status	Negative	317 (72%)
	Unknown	15 (3.4%)
	Positive	107 (24%)
	Started on ARV treatment	97 (90.7%)
Treatment initiation	Mean distance in Kilometres between residence and TB treatment center (range)	3 (2, 8)
Length of time for procurement of TB treatment drugs (in days)	≤14 days	363 (83%)
	15-30 days	45 (10%)
	>30 days	31 (7.1%)
Time between TB diagnosis and TB treatment initiation (in days)	≤30 days	110 (25%)
	31-90 days	121 (28%)
	>90 days	208 (47%)

Traumatic events reported by TB patients

Pain and discomfort, and anxiety and/or depression were the dominant traumatic events reported among the 439 participants surveyed, with 46.5 and 36.9% of participants reporting either a moderate or serious level of impairment, respectively (Table 2).

Table 2. Type and severity of traumatic events reported by TB patients using the EQ-5D-3L scale and IES-R (n=439)

Domain	Mean (SD ¹)	Level of impairment (%)		
		None	Moderate	Serious
Usual activities	1.31 (0.52)	72.2	21.1	2.7
Self-care	1.13 (0.38)	89.1	9.1	1.8
Pain and discomfort	1.49 (0.56)	53.5	43.5	3.0
Mobility	1.23 (0.45)	78.1	20.7	1.1

Anxiety and/or depression	1.43 (0.61)	63.1	30.5	6.4
IES-R	1.37 (0.65)	72.7	17.8	9.6

¹ SD: standard deviation.

In univariate analysis, the following variables household size, , disease localisation, TB diagnostic, distance to treatment site and length of time to drug procurement were significantly associated with stress; were significantly associated with anxiety/depression: level of education, household size, distance to treatment site, and length of time to drug procurement; age, household number size and whether or not antiretroviral treatment were significantly associated with pain/ discomfort; age and distance to treatment site were significantly associated with self care; age, level of education, disease localisation, treatment history, length of time to drug procurement and duration of treatment were significantly with associated usual activities, and; age, household size, , disease localisation and length of time to drug procurement were significantly associated with mobility (Table S1 in supplementary material).

Multivariate analysis identified numerous associations between the domains and the following socio-demographic and clinical variables. Difficulties performing usual activities was positively and significantly associated with elderly participants (OR = 5.42; 95% CI: 1.57 – 22.2; $p < 0.05$) and patients with drug-resistant TB (OR = 27.7; 95% CI: 6.02 – 132; $p < 0.005$). Those significantly less likely to report difficulties with performing usual activities included previously-treated patients (OR = 0.02, CI: 0.00–0.11), HIV-negative patients (OR = 0.52; 95% CI: 0.32–0.87; $p < 0.05$), and those who had been on treatment for more than 90 days (OR = 0.56; 95% CI: 0.32 – 0.98; $p < 0.05$) (Table 3).

Table 3: Multinomial logistic regression between the usual activities and sociodemographic and clinical characteristics.

Characteristic	AOR	95% CI	p-value
Age			
Children	—	—	
Young	1.34	0.46, 4.82	0.6
Adults	2.63	0.87, 9.72	0.11
Olds	5.42	1.57, 22.2	0.011
Tuberculosis type			
Sensitive	—	—	
Resistant	27.7	6.02, 132	<0.001
Treatment history			
New_case	—	—	
Already_treated	0.02	0.00, 0.11	<0.001
TB_diagnostic			
Bacteriological	—	—	
Clinical_examination	2.31	1.20, 4.42	0.011
HIV_status			
Positive	—	—	
Negative	0.52	0.32, 0.87	0.012
Distance between site and residence	1.02	1.00, 1.04	0.062
Duration of the appointment for drug supplies (in days)			
≤14	—	—	
15-30	0.50	0.19, 1.14	0.12

Characteristic	AOR	95% CI	p-value
>30	0.07	0.01, 0.32	0.003
Duration of treatment			
≤30	—	—	
31-90	0.92	0.51, 1.67	0.8
>90	0.56	0.32, 0.98	0.043
AOR = Adjusted Odds Ratio, CI = Confidence Interval			

Significant positive associations with pain and discomfort include household size (OR=1.05;CI: 1.00-1.11,p<0.05), and clinical diagnosis (OR=1.93,CI: 1.06-3.54,p=0.031), while patients whose time for drug procurement was between 15-30 days were 66% less likely to have pain/discomfort than those whose time for drug procurement was within 14 days (OR=0.34,CI: 0.16 - 0.67,p=0.003) (Table 4).

Table 4: Multinomial logistic regression pain /discomfort and sociodemographic and clinical characteristics.

Characteristic	AOR	95% CI	p-value
Age			
Children	—	—	
Young	0.67	0.27, 1.71	0.4
Adults	1.07	0.42, 2.83	0.9
Olds	1.71	0.56, 5.29	0.3
Household members number	1.05	1.00, 1.11	0.045
Tuberculosis _type			
Sensitive	—	—	
Resistant	1.83	0.96, 3.52	0.067
TB_diagnostic			
Bacteriological	—	—	
Clinical _examination	1.93	1.06, 3.54	0.031
RAV_treatment			
Yes	—	—	
No	3.76	0.97, 15.0	0.056
Duration of the appointment for drug supplies (in days)			
≤14	—	—	
15-30	0.34	0.16, 0.67	0.003
>30	0.50	0.21, 1.12	0.10
AOR = Adjusted Odds Ratio, CI = Confidence Interval			

For anxiety, we found that an increase of one unit in the household size was associations with a higher likelihood of reporting anxiety (OR=1.08,CI: 1.03-1.14, p=0.02), as was distance between patients' residence to DTCs (OR=1.03,CI: 1.01-1.05, p<0.001), while non TB/HIV coinfectd patients were 55% less likely to experience anxiety than those coinfectd (OR=0.45,CI: 0.29-0.71,p=0.02; Table 5).

Table 5: Multinomial logistic regression anxiety/depression and sociodemographic and clinical characteristics.

Characteristic	AOR	95% CI	p-value
Household_members number	1.08	1.03, 1.14	0.002
HIV_status			
Positive	—	—	
Negative	0.45	0.29, 0.71	<0.001
Distance between site and residence	1.03	1.01, 1.05	<0.001

Characteristic	AOR	95% CI	p-value
Duration of the appointment for drug supplies (in days)			
≤14	—	—	
15-30	0.35	0.14, 0.75	0.011
>30	0.06	0.00, 0.30	0.007
AOR = Odds Ratio, CI = Confidence Interval			

Regarding stress as measured by IES-R, each one unit increase in the household size, the odds of reporting stress was 1.10 times greater (OR=1.10, CI: 1.04-1.16, $p<0.001$). For an increase of one unit in the distance from patients' residence to DTCs, the odds of having anxiety was 1.03 times greater (OR=1.03, CI: 1.01-1.05, $p<0.001$). Similarly, patients whose duration of treatment was between 31- 90 days had 1.97 times more likely to be stressed than those whose duration of treatment was between 0 to 30 days (OR=1.97, CI: 1.07-3.70, $p=0.032$). Conversely, patients with extra pulmonary were TB had 59% less likely to experience stress than those with lung TB (OR=0.41, CI: 0.21-0.77, $p=0.008$; (Table 6).

Table 6: Multinomial logistic regression between stress and sociodemographic and clinical characteristics.

Characteristic	AOR ¹	95% CI ¹	p-value
Household_members number	1.10	1.04, 1.16	<0.001
HIV_status			
Positive	—	—	
Negative	0.57	0.26, 1.20	0.2
Distance between site and residence			
Duration of the appointment for drug supplies (in days)	—	—	
≤14	0.41	0.21, 0.77	0.008
15-30	1.03	1.01, 1.05	<0.001
>30			
AOR = Odds Ratio, CI = Confidence Interval			

3.2 Qualitative findings

In total, 36 TB patients, including 15 women, were interviewed about their perceptions regarding the COVID-19 pandemic. Thematic analysis was used to identify three predominate themes, and six sub-themes that emerged from the quantitative data:

Theme 1: Social and psychological aspect

Sub-theme: TB patients' perception about the stigma related to COVID-19.

Different opinions were expressed regarding the social and psychological impact of COVID-19. A commonly expressed concern was related to stigma, with the majority of participants accepting that stigma is commonly experienced by TB patients, and who also felt that the COVID-19 pandemic has increased the amount of stigma towards people with symptoms that may be suggestive of COVID-19. As one participant noted: "Once you cough or sneeze, people distance themselves from you; then you feel bad... they think you have COVID-19 if you cough once or twice. For those who do not know that you have TB, they are going to think that you have COVID-19 because we do not overthink TB now.". One participant shared how some people, without any knowledge on this person's health status, do not even hesitate to walk away from them with a contemptuous gaze; other TB patients

interviewed revealed how this practice has become more obvious during the COVID-19 pandemic, with people assuming symptoms such as coughs are an indication of COVID-19, rather than TB: "... 'It's so clear to me because my husband doesn't allow me to come near him since he learned that COVID-19 also causes coughs. Usually, I did everything with him. But now, everything has changed, he no longer approaches me like before coronavirus pandemic, we no longer talk too much, and even now we eat separately, you see...".

Sub-theme: description of the health status of TB patients during COVID-19 pandemic.

The majority of participants (n = 25) believed that their overall health conditions were better before the start of the COVID-19 pandemic. Some participants reported little hope for a full recovery of their health, due to the difficulties they faced in accessing effective TB treatment during the pandemic and the increasingly difficult living conditions under COVID, such as challenges with accessing food : *"Mentally, I feel like garbage; all I'm worried about right now is how to get my health back. . I don't feel well right now, you see that by yourself! Due to food problems I barely earn, it's just filling my stomach. Currently, it's not okay, and sometimes I don't have transport to get to the centre or sometimes even if you leave you are told that the centre is out of medication. I have lost a lot of weight at the moment. I lost much hope despite the encouragement of the doctors"*.

Theme 2: Economic aspects

Sub-theme: Financial situation of TB patients during the COVID-19 pandemic.

Respondants highlighted how the cessation of certain activities during the COVID-19 pandemic directly and/or indirectly affected their financial conditions, which created challenges for accessing TB services and maintaining their treatment. As this participant, who earns money to support their family through artistic activities noted: *".... Financially it [the situation during COVID-19] is not good. There are no events, and as I am an artist, this is where I earn money to feed my family and pay for transportation to the [treatment] centre. But as there is currently no event, even finding transport to get to the treatment centre is a problem because transport is expensive, sometimes we do not come on the date that the doctors tell us to come, because we don't have transportation. You see, it's not easy"..*

Sub-theme: Diet of TB patients during the COVID-19 pandemic.

TB patients revealed during their interviews that their health condition had deteriorated significantly during COVID due to the lack of an appropriate diet. They noted how they were no longer able to comply with the diet recommended by their doctors. As this participant noted: *"I cannot eat all of the foods my doctor recommends. I eat with difficulty, and for example, with this difficult situation, I only eat rice. I have diabetes, and my doctor has advised me to limit my intake a bit. But I have to consume it because that's what I earn .. "*.

Theme 3: Perceptions on government actions during the COVID-19 pandemic.

Sub-theme: Restrictive measures

According to our participants, the Guinean population's life depends on their daily economic activities; therefore, the restrictive measures imposed by the Government to limit the spread of COVID were largely regarded as drastic and seen to cause more harm than good: *"These measures of the state are salutary, but which is not without consequences for some, you knew it as well as me sir... these measures make life difficult by blocking the activities of*

people, and you know here what we see, that's what we eat. Guineans eat from day to day, so these measures increase their suffering".

Sub-theme: Mitigation measures

Unanimously, respondents believed that the Government should financially help its people, especially those who were sick, poor, children or elderly. *"I thank the state for what is already done the price of fuel at the pump, this leads to a reduction in the prices of necessities (rice, sugar ...), and, therefore facilitates access to food by the population. But some do not benefit from it and others do not even know about it. But when the price drops, everyone will feel this impact on their living condition."* Likewise, some believed that special attention should be paid to TB patients given the additional burdens created by their illness: *"Yes, the state should help its population, especially patients like us tuberculosis patients who are rejected by our bosses for fear of being infected"*

4. Discussion

The advent of the COVID-19 pandemic has caused a noticeable imbalance in the smooth running of our society. Healthy and sick individuals feel this impact differently. Regardless of any external disturbances, TB patients constitute a vulnerable group. This study showed significant additional discomfort for these TB patients. Even though our study coincided with the onset of relaxation of the restrictive measures, we found a significant proportion of patients reporting pain or discomfort, and anxiety or depression. Our results are consistent with a recent evaluation of the stress response during COVID-19 in which a significant proportion of patients struggled to cope with stress [22,23]. Apart from the situation induced by COVID-19 pandemic, psychological problems are quite recurrent in TB patients and should be the subject of special attention [12,13]. One of the reasons for this psychosocial state of patients stems from the stigma experienced by TB patients during COVID-19 given the similarity between these two diseases [24,25]. Also, in Guinea, the implementation of restrictive measures limited the population's movements and led to the closure of many businesses which further reduced many patients' support sources, worsening financial problems and exacerbating stress. Our findings underscore the need for comprehensive psycho-social support to TB patients during times of public health emergencies to protect them against further vulnerability.

In our study, we also sought to understand factors explaining the different quality of life scales (EQ 5D 3L). Each item (pain or discomfort, usual activities and anxiety or depression) of EQ-5D-3L made it possible to identify the factors associated with them. Socio-demographically, it emerges that age, the number of people in the household, and the distance between patients' homes and the care site explain the deterioration in life quality. Among the factors degrading the quality of life, we have unsurprisingly the elderly, who presented more difficulties in carrying out daily activities than children. This category of patients is potentially subject to various pathologies and dysfunctions. Due to reduced immunity, their fragility exposes them to more significant risks, especially side effects [13,23,26] and malnutrition. Unlike young people, TB in the elderly poses a real diagnostic and management challenge [25–28]. The number of people in households has also emerged as a factor associated with mobility problems, pain or discomfort, anxiety and stress. This situation could be explained by patients' economic situation, who lacks adequate care and as a result, exposed to these various traumatic events. The socioeconomic conditions are also manifested by the fact that the patients who were far from care had a degraded HRQOL. Indeed, not often having the means to travel regularly to health centres increases considerably the risk of having a degraded HRQOL.

We found that patients clinically diagnosed, coinfecting or with extra-pulmonary or drug-resistant TB were dominant factors associated with a deterioration in the HRQOL. Previously treated patients had a lower risk of poor quality of life compared to new ones. Newly diagnosed patients struggle to adapt to the disease's new life, especially in a particular pandemic context. HRQOL strongly depends on clinical forms of tuberculosis [5], which depend on apparent clinical and subclinical manifestations. The approximate duration of the anti-tuberculosis drug procurement remains the primary factor identified in this study. Patients expressed a lot of difficulties related to travel and the financial problems involved. We have observed that the shorter time for drug procurement, the more the patients' HRQOL deteriorated. Thus, the close approximation of supply time increases the cost and the fear and discomfort when travelling. Tuberculosis patients in the African context often face catastrophic health costs related to their living conditions [29]. Therefore, any gait that requires frequent movement is affecting the HRQOL.

Beyond the associated factors revealed by the quantitative method, the other recurring element expressed by patients in the qualitative part is malnourishment. The nutritional support program only benefits multidrug-resistant patients in Guinea, leaving the rest of the TB patients neglected. This is not without consequences for tuberculosis patients, and for this reason, the substantial cessation of activities has impacted the possibility for these tuberculosis patients to meet their nutritional needs.

Our study has the advantage of combining two complementary approaches. But the study design did not allow us to fit all quantitative questions into the qualitative scheme. This made the comparison of the two approaches difficult. It should also be noted that the study's implementation coincided with the start of the relaxation of restrictive measures. This implies that the resulting image would be even worse at the onset of COVID-19 in Guinea. In any case, a comparison of the different periods would have been useful. We were unable to fully explore the real economic impact (associated costs) on patients' lives. The simplification of the study design, the need for information at a brief time, social distancing concerns, and other barrier measure limited our ability to go into depth in the assessment. Although we were unable to survey tuberculosis patients outside the capital, our sample remains representative of most tuberculosis patients in the Guinean context, because the Capital Conakry takes care of 60% of tuberculosis patients.

5. Conclusions

This study confirmed common problems that influence tuberculosis patients' quality of life and how such challenges are exacerbated by the COVID-19 pandemic in Guinea. It is urgent to put a mitigation plan that should consider patients with particular clinical characteristics, nutritional issues, and the strategy to reduce tuberculosis patients' movement for accessing TB care during COVID-19 pandemic. Finally, this plan must necessarily include a psychosocial care component

Supplementary Materials: The following supporting information can be downloaded at: www.mdpi.com/xxx/s1, Table S1: Univariate analysis between dependant variables and the characteristic of the patients.

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Informed Consent Statement: Informed consent was obtained from all subjects involved in the study. For children under 18, consent was obtained from their parents and/or legal guards. All procedures in this study were carried out in compliance with principles and regulations in effect in Guinea.

Data Availability Statement: Requests for original datasets used in this manuscript can be directed to the corresponding author.

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Disclaimer

VV and CSM are staff members of the World Health Organization; the authors alone are responsible for the views expressed in this publication and they do not necessarily represent the decisions, policy or views of the WHO.

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Supplementary material

Table S1. Univariate analysis between dependant variables and the characteristic of the patients.

	Stress p.trend ¹	Anxiety depression p.trend ¹	Pain Discomfort p.trend ¹	Selfcare p.trend ¹	Usual p.trend ¹	activities	Mobility p.trend ¹
age	0.256	0.120	0.017	0.048	<0.001		0.001
Gender	0.575	0.926	0.865	0.556	0.834		0.846
Matrimonial status	0.210	0.627	0.136	0.137	0.686		0.325
residence	0.116	0.099	0.174	0.381	0.687		0.422
Education	0.391	0.027	0.073	0.963	0.008		0.153
Occupation	0.341	0.885	0.333	0.703	0.206		0.638
Household number/size	<0.001	<0.001	0.005	0.063	0.069		0.005
Tuberculosis type	0.924	0.155	0.179	0.183	0.700		0.408
Disease localisation	0.008	0.344	0.517	0.322	0.034		0.045
Treatment history	0.656	0.347	0.449	0.622	0.041		0.148
TB diagnostic	0.005	0.113	0.067	0.227	0.072		0.710
Treatment type	0.852	0.198	0.225	0.324	0.861		0.268
Type multi-drug treatment	0.881	0.267	0.072	0.379	0.761		0.326
Anti-retroviral treatment	0.056	0.001	0.007	0.768	0.063		0.071
Distance site Patient residence	0.001	<0.001	0.182	0.025	0.185		0.313
Length time for drug procurement (in days)	0.004	<0.001	0.129	0.242	0.009		0.004
Treatment initiation	0.383	0.430	0.700	0.991	0.975		0.460
Duration of treatment	0.381	0.202	0.276	0.199	0.019		0.103

¹ p trend computed with the levels (no problem, moderate, serious) for each dependent variable