Mathematical modeling of typhoid fever disease incorporating unprotected humans 1 in the spread dynamics. 2

3

4 Abstract

In this study we have develop a deterministic mathematical model for spread dynamics of 5 typhoid fever disease incorporating unprotected humans. The model result into a system of 6 7 ordinary differential equations which are used to study the spread dynamics of typhoid fever. The model incorporating Susceptible, unprotected, Infectious and Recovered humans which are 8 analyzed mathematically. The existence of steady states of the mathematical model is 9 determined. More so we show the existence and positivity of a solution and finally computed the 10 basic reproductive number using next generation matrix. 11

Key words: Basic reproduction number, invariant region, positivity of solution, Mathematical 12 13 model, Disease Free Equilibrium, Endemic equilibrium point. And and a second

Introduction 14

Typhoid fever is an endemic disease that is classified as an enteritis disease. The disease is 15 caused by a bacterium called Salmonella Typhi. It is a common infectious disease in human beings 16 and is transmitted through food and water contaminated with faeces and urine of an infected 17 person [3]. The disease is endemic in developing countries where it continuously causes illness 18 and death. This is contributed by unsafe water supply, poor food hygiene and wanting 19 environmental sanitation. According to World Health Organization an estimated 17 million 20 21 illness cases of typhoid fever were reported per year worldwide resulting to 0.6 million deaths annually[4,5]. 22

1. Description and model formulation 23

We formulated a deterministic model for spread dynamics of typhoid fever that considers 24 25 human population at time t. The model is divided into four compartments as follows. Susceptible(S), Unprotected (E), Infective (I) and Recovered(R). The model has the following 26 flow. $S \rightarrow E \rightarrow I \rightarrow R \rightarrow S$. We use the following parameters in our model.(i) μ is the natural 27 death rate (ii) α is the disease induced death rate.(iii) Λ human recruitment rate (birth). (iv) β 28 29 disease interaction rate $(v)\Omega$ unprotected symptoms showing rate $(v)\gamma$ Infective recovery rate and finally(vii) δ this is the rate at which recovered humans loses temporary immunity obtained 30 through treatment and get the disease back again. All the compartments are positive in the 31 feasible region φ where $\{S, E, I, R\} \in \varphi \subset R_+^4$. All the solutions are also bounded in φ such that 32

- $0 \le N \le \frac{\Lambda}{\mu}$. Thus the model is epidemiologically well posed in the region φ . 33
- The following flow chart shows various compartments in the model. 34
- 35
- 36



$$\frac{dS}{dt} = \Lambda + \delta R - \beta SI - \mu S$$

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$$\frac{dE}{dt} = \beta SI - \Omega E - \mu E$$
$$\frac{dI}{dt} = \Omega E - \gamma I - \alpha I - \mu I$$
$$\frac{dR}{dt} = \gamma I - \delta R - \mu R$$

A

(1)

2. Disease free equilibrium point and endemic equilibrium point 46

The disease free equilibrium of the model is obtained by setting 47

$$\frac{dS}{dt} = \frac{dE}{dt} = \frac{dI}{dt} = \frac{dR}{dt} = 0$$

In absence of disease 49

50
$$E = 0, I = 0, R = 0$$
.

Setting the right hand side of equations of system 1 to zero we have 51

52

$$\Lambda + \delta R - \beta SI - \mu S = 0.$$

$$\beta SI - \Omega E - \mu E = 0$$

$$\Omega E - \gamma I - \alpha I - \mu I = 0$$

$$\gamma I - \delta R - \mu R = 0$$
(2)

53 Hence model has a disease free equilibrium given by

54
$$(S^* E^* I^* R^*) = \left(\frac{\Lambda}{\mu}, 0, 0, 0\right)$$
 (3)

- 55 The basic reproductive number (R_0) which is average number of secondary infections caused by
- 56 one infectious individual introduced in a completely susceptible population is obtained using $\beta S\Omega$
- 57 next generation matrix as $R_0 = \frac{\beta S\Omega}{(\Omega + \mu)(\alpha + \mu + \gamma)}$ where at disease free equilibrium

58
$$R_0 = \frac{\beta \Lambda \Omega}{\mu (\Omega + \mu) (\alpha + \mu + \gamma)}.$$

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- 60
- 61 Theorm1
- 62 If $\Omega < \frac{\mu^2 (\alpha + \mu + \gamma)}{\beta \Lambda u (\alpha + \mu + \gamma)}$, there disease free equilibrium will be stable and typhoid disease will not
- 63 have a hand in the population.
- 64 **Proof**

65 When
$$R < 1$$
; this means that $\frac{\beta \Lambda \Omega}{\mu(\Omega + \mu)(\alpha + \mu + \gamma)} < 1$.

66 Making
$$\Omega$$
 the subject, $\Omega < \frac{\mu^2 (\alpha + \mu + \gamma)}{\beta \Lambda - u (\alpha + \mu + \gamma)}$

Disease free equilibrium point therefore is locally asymptotically stable if the basic reproduction number (R_0) less than one $(R_0 < 1)$ and unstable if the basic reproduction number is greater than $(R_0 > 1)$.

- 70 **3. Endemic equilibrium point**
- Endemic equilibrium E_2^* ; disease exists. Evaluating the state variables of equations of the system 2, the endemic equilibrium points are in this form

$$E_2^* = \left\{ S^{**}, E^{**}, I^{**}, R^{**} \right\}$$

Where

$$S^{**} = \frac{(\Omega + \mu)(\gamma + \alpha + \mu)}{\Omega\beta}$$

$$E^{**} = \frac{(\gamma + \alpha + \mu)(\delta + \mu)\{\Lambda\Omega\beta - \mu\{(\Omega + \mu)(\gamma + \alpha + \mu)\}}{\beta\Omega\{(\delta + \mu)(\Omega + \mu)(\gamma + \alpha + \mu)\} - \gamma\Omega\delta}$$

$$I^{**} = \frac{(\delta + \mu)}{\beta} \cdot \{\frac{\Lambda\Omega\beta - \mu\{(\Omega + \mu)(\gamma + \alpha + \mu)\}}{\{(\delta + \mu)(\Omega + \mu)(\gamma + \alpha + \mu)\} - \gamma\Omega\delta}$$

$$R^{**} = \frac{1}{\beta} \left\{ \frac{\{\gamma \Lambda \Omega \beta - \gamma \mu \{ (\Omega + \mu)(\gamma + \alpha + \mu) \}}{\{ (\delta + \mu)(\Omega + \mu)(\gamma + \alpha + \mu) \} - \gamma \Omega \delta} \right\}$$

76 **4. Stability of endemic equilibrium**

Proof: By the use of lyapunov function defined by LaSalle [1976]

78
$$L(S^{**}, E^{**}, I^{**}, R^{**}) = (S - S^{**} - S^{**}\ln(\frac{S^{**}}{S}) + \left(E - E^{**} - E^{**}\ln(\frac{E^{**}}{E}) + I - I^{**} - I^{**}\ln(\frac{I^{**}}{I}) + R - R^{**} - R^{**}\ln(\frac{R^{**}}{R})\right)$$

79 computing the derivative of L along the solutions of the system is directly:

$$80 \qquad \frac{dL}{dt} = \left(\frac{s-s^{**}}{s}\right)\frac{dS}{dt} + \left(\frac{E-E^{**}}{E}\right)\frac{dE}{dt} + \left(\frac{I-I^{**}}{I}\right)\frac{dI}{dt} + \left(\frac{R-R^{**}}{R}\right)\frac{dR}{dt} \tag{5}$$

81 Substituting the equations of system 1 in equation 5, the equation becomes

$$\frac{dL}{dt} = \left[\left(\frac{s - s^{**}}{s} \right) \Lambda + \delta R - (\beta I + \mu) S \right] + \left[\left(\frac{E - E^{**}}{E} \right) \beta S I - (\Omega + \mu) E \right] + \left[\left(\frac{I - I^{**}}{I} \right) \Omega E - (\gamma + \alpha + \mu) I \right] + \left[\left(\frac{R - R^{**}}{R} \right) \gamma I - (\delta + \mu) R \right]$$
(6)

83 Expanding equation 6, it produces

84
$$\frac{dL}{dt} = \Lambda + \delta R - (\beta I + \mu)S - \Lambda \frac{S^{**}}{S} - \delta R \frac{S^{**}}{S} + (\beta I + \mu)S^{**} + \beta SI - (\Omega + \mu)E - \beta SI \frac{E^{**}}{E} + (\Omega + \mu)E^{**} + \Omega E - (\gamma + \alpha + \mu)I - \Omega E \frac{I^{**}}{I} + (\gamma + \alpha + \mu)I^{**} + \gamma I - (\delta + \mu)R - \gamma I \frac{R^{**}}{R} + (\delta + \mu)R^{*}$$

85 Further simplification result to

$$\frac{dL}{dt} = \left[\Lambda + \delta R + (\beta I + \mu)S^{**} + \beta SI + (\Omega + \mu)E^{**} + \Omega E + (\gamma + \alpha + \mu)I^{**} + \gamma I + (\delta + \mu)R^{**}\right]$$

$$+ \left[-(\beta I + \mu)S - \Lambda \frac{S^{**}}{S} - \delta R \frac{S^{**}}{S} - (\Omega + \mu)E - \beta SI \frac{E^{**}}{E} - (\gamma + \alpha + \mu)I - \Omega E \frac{I^{**}}{I} - (\delta + \mu)R - \gamma I \frac{R^{**}}{R}\right]$$
87 Or

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$$\frac{dL}{dt} = \left[\Lambda + \delta R + (\beta I + \mu)S^{**} + \beta SI + (\Omega + \mu)E^{**} + \Omega E + (\gamma + \alpha + \mu)I^{**} + \gamma I + (\delta + \mu)R^{**}\right]$$

$$-\left[(\beta I + \mu)S + \Lambda \frac{S^{**}}{S} + \delta R \frac{S^{**}}{S} + (\Omega + \mu)E + \beta SI \frac{E^{**}}{E} + (\gamma + \alpha + \mu)I + \Omega E \frac{I^{**}}{I} + (\delta + \mu)R + \gamma I \frac{R^{**}}{R}\right]$$

From equation it's clear that ; $\frac{dL}{dt} = A - B$. Where A are the positive terms and B are the 89 90 negative ones, such that;

 $A = \Lambda + \delta R + (\beta I + \mu) S^{**} + \beta SI + (\Omega + \mu) E^{**} + \Omega E + (\gamma + \alpha + \mu) I^{**} + \gamma I + (\delta + \mu) R^{**}$ $B = (\beta I + \mu) S + \Lambda \frac{S^{**}}{S} + \delta R \frac{S^{**}}{S} + (\Omega + \mu) E + \beta SI \frac{E^{**}}{E} + (\gamma + \alpha + \mu) I + \Omega E \frac{I^{**}}{I} + (\delta + \mu) R + \gamma I \frac{R^{**}}{R}$ If A < B then $\frac{dL}{dt} \le 0$ 92

93
$$\frac{dL}{dt} = 0$$
 Only if $S = S^{**}, E = E^{**}, I = I^{**}, R = R^{**}$

The largest invariant set in $\{(S, E, I, R) \in \varphi : \frac{dL}{dt} = 0\}$ is a singleton E_2^* . Where E_2^* is the 94 endemic equilibrium. Therefore, the endemic equilibrium is globally asymptotically stable 95 in the invariant region φ if A < B [1,2]. 96

97 **Conclusion.**

From our finding if $\Omega < \frac{\mu^2 (\alpha + \mu + \gamma)}{\beta \Lambda - u (\alpha + \mu + \gamma)}$, there disease equilibrium will be stable and typhoid 98

disease will not have a hand in the population. However if $\Omega > \frac{\mu^2 (\alpha + \mu + \gamma)}{\beta \Delta - \mu (\alpha + \mu + \gamma)}$, then disease 99

will be dependent on prevailing circumstances. We also performed numerical simulations to 100

101 determine the changes in various compartments with time using MATLAB ode solve software.

There is direct variation relationship between the unprotected and infectious compartments, 102

therefore the unprotected humans contribute significantly to the spread dynamics of typhoid 103 fever disease. 104

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