

Provided for non-commercial research and education use.
Not for reproduction, distribution or commercial use.



This article appeared in a journal published by Elsevier. The attached copy is furnished to the author for internal non-commercial research and education use, including for instruction at the authors institution and sharing with colleagues.

Other uses, including reproduction and distribution, or selling or licensing copies, or posting to personal, institutional or third party websites are prohibited.

In most cases authors are permitted to post their version of the article (e.g. in Word or Tex form) to their personal website or institutional repository. Authors requiring further information regarding Elsevier's archiving and manuscript policies are encouraged to visit:

<http://www.elsevier.com/copyright>



Contents lists available at SciVerse ScienceDirect

Journal of Theoretical Biology

journal homepage: www.elsevier.com/locate/yjtbi



Corrigendum

Corrigendum to “Modeling the transmission dynamics and control of Hepatitis B virus in China” [J. Theor. Biol. 262 (2010) 330–338]

Lan Zou^a, Weinian Zhang^a, Shigui Ruan^{b,*}

^a Yangtze Center of Mathematics and Department of Mathematics, Sichuan University, Chengdu, Sichuan 610064, PR China

^b Department of Mathematics, University of Miami, Coral Gables, FL 33124-4250, USA

In this note, we provide corrections to a theorem and its proof in our paper Zou et al. (2010) and make some additional remarks.

1. Corrections

Theorem 2 in our paper Zou et al. (2010) states that the endemic equilibrium E^* of model (2.1) is a stable node if it exists. Both the statement and the proof were incorrect. We realized the mistake right after the publication of the paper and a correction of the statement and its proof was presented in the Ph.D. Thesis of the first author in March 2010 (Zou, 2010, Theorem 5.1.2, pp. 65–66 and Appendix III, p. 95). The following corrections are adapted from Zou (2010).

In order to discuss the steady states of model (2.1) of Zou et al. (2010), we only need to consider the following system, which is (3.1) in Zou et al. (2010):

$$\begin{cases} \frac{dS}{dt} = \mu\omega(1-\nu C) + \psi V - (\mu_0 + \beta I + \epsilon\beta C + \gamma_3)S, \\ \frac{dL}{dt} = (\beta I + \epsilon\beta C)S - (\mu_0 + \sigma)L, \\ \frac{dI}{dt} = \sigma L - (\mu_0 + \gamma_1)I, \\ \frac{dC}{dt} = \mu\omega\nu C + q\gamma_1 I - (\mu_0 + \mu_1 + \gamma_2)C, \\ \frac{dV}{dt} = \mu(1-\omega) + \gamma_3 S - (\mu_0 + \psi)V. \end{cases}$$

As defined in (2.2) of Zou et al. (2010), the basic reproduction number is given by

$$R_0 = \frac{\mu(\psi + \mu_0\omega)}{\mu_0(\mu_0 + \gamma_3 + \psi)(\mu_0 + \gamma_1)(\mu_0 + \sigma)} \left[1 + \frac{q\gamma_1\epsilon}{\mu_0 + \mu_1 + \gamma_2 - \mu\omega\nu} \right].$$

It is proved in Zou et al. (2010) that the system has an endemic equilibrium $E^* = (S^*, L^*, I^*, C^*, V^*)$ if $R_0 > 1$, where

$$S^* = \frac{(\mu_0 + \mu_1 + \gamma_2 - \mu\omega\nu)(\mu_0 + \sigma)(\gamma_1 + \mu_0)}{(\mu_0 + \mu_1 + \gamma_2 + \epsilon q\gamma_1 - \mu\omega\nu)\beta\sigma} = \frac{S_0}{R_0},$$

$$\begin{aligned} L^* &= \frac{\mu_0(\gamma_3 + \mu_0 + \psi)S^*(\mu_0 + \mu_1 + \gamma_2)(\gamma_1 + \mu_0)(R_0 - 1)}{(\mu_0 + \psi)\sigma[\beta S^*(\mu_0 + \mu_1 + \gamma_2 + \epsilon q\gamma_1 - \mu\omega\nu) + \mu\omega\nu q\gamma_1]}, \\ I^* &= \frac{\mu_0 S^*(\gamma_3 + \mu_0 + \psi)(R_0 - 1)(\mu_0 + \mu_1 + \gamma_2 - \mu\omega\nu)}{(\mu_0 + \psi)[\beta S^*(\mu_0 + \mu_1 + \gamma_2 + \epsilon q\gamma_1 - \mu\omega\nu) + \mu\omega\nu q\gamma_1]}, \\ C^* &= \frac{\mu_0\epsilon\beta q\gamma_1(\mu_0 + \gamma_1)(\gamma_3 + \mu_0 + \psi)(S^*)^2(R_0 - 1)}{(\mu_0 + \psi)[\beta S^*(\mu_0 + \mu_1 + \gamma_2 + \epsilon q\gamma_1 - \mu\omega\nu) + \mu\omega\nu q\gamma_1]}, \\ V^* &= \frac{\mu(1-\omega) + \gamma_3 S^*}{\mu_0 + \psi}. \end{aligned}$$

Theorem 2 in Zou et al. (2010) should be replaced by the following result.

Theorem 2. Assume that $R_0 > 1$. Then the endemic equilibrium E^* is locally asymptotically stable if

- (i) $b_1 b_2 - b_3 > 0$,
- (ii) $b_3(b_1 b_2 - b_3) + b_1(b_5 - b_1 b_4) > 0$,
- (iii) $(b_1 b_2 - b_3)(b_3 b_4 - b_2 b_5) - (b_5 - b_1 b_4)^2 > 0$,

where b_i s are given as follows

$$\begin{aligned} b_1 &= \frac{\sigma L^*}{I^*} + \frac{q\gamma_1 I^*}{L^*} + \frac{(\beta I^* + \epsilon\beta C^*)S^*}{L^*} + \psi + \beta I^* + \epsilon\beta C^* + \gamma_3 + 2\mu_0, \\ b_2 &= \frac{1}{L^* I^* C^*} [I^* S^* C^* (I^* + C^* \epsilon)^2 \beta^2 + (S^* \epsilon C^* I^{*2} q\gamma_1 + 2\mu_0 S^* I^{*2} C^* \\ &\quad + \epsilon C^{*2} L^{*2} \sigma + S^* I^{*3} q\gamma_1 + S^* \epsilon C^{*2} \sigma L^* + \epsilon C^{*2} \mu_0 L^* I^* + I^{*3} L^* q\gamma_1 \\ &\quad + \epsilon C^{*2} \psi L^* I^* + \gamma_3 S^* I^{*2} C^* + \gamma_3 S^* \epsilon C^{*2} I^* + \epsilon C^* L^* I^{*2} q\gamma_1 \\ &\quad + I^{*2} \psi L^* C^* + \psi S^* I^{*2} C^* + \psi S^* \epsilon C^{*2} I^* + 2\mu_0 S^* \epsilon C^{*2} I^* + I^{*2} \mu_0 L^* C^* \\ &\quad + I^* L^{*2} \sigma C^*) \beta + \gamma_3 L^{*2} \sigma C^* + \mu_0^2 L^* I^* C^* + 2\mu_0 L^{*2} \sigma C^* + \psi L^{*2} \sigma C^* \\ &\quad + \gamma_3 \mu_0 L^* I^* C^* + \mu_0 \psi L^* I^* C^* + 2\mu_0 L^* I^{*2} q\gamma_1 \\ &\quad + \gamma_3 L^* I^{*2} q\gamma_1 + L^{*2} \sigma q\gamma_1 I^* + \psi L^* I^{*2} q\gamma_1], \\ b_3 &= \frac{1}{L^* I^* C^*} (\mu_0^2 L^{*2} \sigma C^* + 2\mu_0 L^{*2} \sigma q\gamma_1 I^* + \mu_0^2 \beta S^* I^{*2} C^* \\ &\quad + 2\beta^2 I^{*2} \mu_0 S^* \epsilon C^{*2} + \beta I^* \psi L^{*2} \sigma C^* + \beta I^{*3} \psi L^* q\gamma_1 \\ &\quad + \epsilon\beta C^* \psi L^* I^{*2} q\gamma_1 + \epsilon^2 \beta^2 C^{*3} S^* \sigma L^* + \epsilon^2 \beta^2 C^{*2} S^* I^{*2} q\gamma_1 \\ &\quad + \epsilon\beta C^* L^{*2} \sigma q\gamma_1 I^* + \beta I^{*2} L^{*2} \sigma q\gamma_1 + \mu_0^2 \beta S^* \epsilon C^{*2} I^* + \epsilon^2 \beta^2 C^{*3} \mu_0 S^* I^* \\ &\quad + \epsilon\beta C^{*2} \mu_0 L^{*2} \sigma + \mu_0 \psi \beta S^* I^{*2} C^* + \mu_0 \psi \beta S^* \epsilon C^{*2} I^* + \mu_0 \psi L^{*2} \sigma C^* \\ &\quad + \mu_0 \psi L^* I^{*2} q\gamma_1 + \epsilon\beta C^* \mu_0 L^* I^{*2} q\gamma_1 + \epsilon^2 \beta^2 C^{*3} \psi S^* I^* \end{aligned}$$

DOI of original article: <http://dx.doi.org/10.1016/j.jtbi.2009.09.035>

* Corresponding author.

E-mail address: ruan@math.miami.edu (S. Ruan).

$$\begin{aligned}
 & + \varepsilon\beta C^* \psi L^* \sigma + \gamma_3 \mu_0 \beta S^* I^* C^* + \gamma_3 \mu_0 \beta S^* \varepsilon C^* I^* + \gamma_3 \mu_0 L^* \sigma C^* \\
 & + \gamma_3 \mu_0 L^* I^* q \gamma_1 + \mu_0^2 L^* I^* q \gamma_1 + \beta^2 \sigma L^* S^* C^* + \beta^2 I^* \mu_0 S^* C^* \\
 & + \gamma_3 \beta S^* I^* q \gamma_1 + \gamma_3 \beta S^* \varepsilon C^* \sigma L^* + \gamma_3 \beta S^* \varepsilon C^* I^* q \gamma_1 + \psi \beta S^* I^* q \gamma_1 \\
 & + \gamma_3 L^* \sigma q \gamma_1 I^* + \psi \beta S^* \varepsilon C^* I^* q \gamma_1 + \psi \beta S^* \varepsilon C^* \sigma L^* \\
 & + \psi L^* \sigma q \gamma_1 I^* + \beta I^* \mu_0 L^* \sigma C^* + \beta I^* \mu_0 L^* q \gamma_1 + \beta^2 I^* \psi S^* C^* \\
 & + \beta^2 I^* S^* q \gamma_1 + 2\beta^2 I^* \psi S^* \varepsilon C^* + 2\mu_0 \beta S^* I^* q \gamma_1 + 2\mu_0 \beta S^* \varepsilon C^* \sigma L^* \\
 & + 2\mu_0 \beta S^* \varepsilon C^* I^* q \gamma_1 + 2\beta^2 I^* S^* \varepsilon C^* \sigma L^* + 2\beta^2 I^* S^* \varepsilon C^* q \gamma_1, \\
 b_4 = & \frac{1}{L^* I^* C^*} (2\beta^2 \sigma L^* q \gamma_1 C^* \varepsilon S^* + 2\beta^2 I^* \psi S^* \varepsilon C^* \sigma L^* + 2\beta^2 I^* \psi S^* \varepsilon C^* q \gamma_1 \\
 & + 2\beta^2 I^* \mu_0 S^* \varepsilon C^* \sigma L^* + \beta^2 \sigma L^* I^* S^* q \gamma_1 + \beta^2 I^* \psi S^* q \gamma_1 \\
 & + \beta \sigma L^* I^* q \gamma_1 C^* \mu \omega \nu + \beta^2 \sigma L^* I^* \varepsilon^2 C^* q \gamma_1 S^* + \beta \sigma L^* I^* \varepsilon C^* q \gamma_1 \mu \omega \nu \\
 & + \beta I^* \mu_0 L^* \sigma q \gamma_1 + \beta I^* \psi L^* \sigma q \gamma_1 + \mu_0^2 \beta S^* I^* q \gamma_1 + \beta^2 I^* \mu_0 S^* q \gamma_1 \\
 & + \mu_0^2 \beta S^* \varepsilon C^* \sigma L^* + \mu_0^2 \beta S^* \varepsilon C^* I^* q \gamma_1 + \varepsilon \beta C^* \psi L^* \sigma q \gamma_1 I^* \\
 & + \gamma_3 \mu_0 \beta S^* I^* q \gamma_1 + \mu_0^2 L^* \sigma q \gamma_1 I^* + \varepsilon^2 \beta^2 C^* \mu_0 S^* \sigma L^* \\
 & + \varepsilon^2 \beta^2 C^* \mu_0 S^* I^* q \gamma_1 + \varepsilon \beta C^* \mu_0 L^* \sigma q \gamma_1 I^* + \varepsilon^2 \beta^2 C^* \psi S^* \sigma L^* \\
 & + \varepsilon^2 \beta^2 C^* \psi S^* I^* q \gamma_1 + \mu_0 \psi L^* \sigma q \gamma_1 I^* \\
 & + \gamma_3 \mu_0 \beta S^* \varepsilon C^* \sigma L^* + \gamma_3 \mu_0 \beta S^* \varepsilon C^* I^* q \gamma_1 \\
 & + \gamma_3 \mu_0 L^* \sigma q \gamma_1 I^* + \mu_0 \psi \beta S^* I^* q \gamma_1 + \mu_0 \psi \beta S^* \varepsilon C^* \sigma L^* \\
 & + \mu_0 \psi \beta S^* \varepsilon C^* I^* q \gamma_1 + \beta^2 \sigma L^* I^* \mu_0 S^* C^* \\
 & + \beta^2 \sigma L^* I^* \psi S^* C^* + 2\beta^2 I^* \mu_0 S^* \varepsilon C^* q \gamma_1), \\
 b_5 = & \frac{1}{L^* I^* C^*} (\beta \sigma L^* I^* \varepsilon C^* \psi q \gamma_1 \mu \omega \nu + \beta^2 \sigma L^* I^* \varepsilon^2 C^* \psi q \gamma_1 S^* \\
 & + \beta \sigma L^* I^* \varepsilon C^* \mu_0 q \gamma_1 \mu \omega \nu + \beta^2 \sigma L^* I^* \varepsilon^2 C^* \mu_0 q \gamma_1 S^* \\
 & + \beta^2 \sigma L^* I^* \psi S^* q \gamma_1 + \beta^2 \sigma L^* I^* \mu_0 S^* q \gamma_1 + \beta \sigma L^* I^* \mu_0 q \gamma_1 C^* \mu \omega \nu \\
 & + 2\beta^2 \sigma L^* I^* \mu_0 q \gamma_1 C^* \varepsilon S^* + 2\beta^2 \sigma L^* I^* \psi q \gamma_1 C^* \varepsilon S^* \\
 & + \beta \sigma L^* I^* \psi q \gamma_1 C^* \mu \omega \nu).
 \end{aligned}$$

Proof. The Jacobian matrix of the system at E^* is

$$J(E^*) = \begin{pmatrix} -\mu_0 - \beta I^* - \varepsilon \beta C^* - \gamma_3 & 0 & -\beta S^* & -\mu \omega \nu - \varepsilon \beta S^* & \psi \\ \beta I^* + \varepsilon \beta C^* & -\mu_0 - \sigma & \beta S^* & \varepsilon \beta S^* & 0 \\ 0 & \sigma & -\mu_0 - \gamma_1 & 0 & 0 \\ 0 & 0 & q \gamma_1 & \mu \omega \nu - \mu_0 - \mu_1 - \gamma_2 & 0 \\ \gamma_3 & 0 & 0 & 0 & -\mu_0 - \psi \end{pmatrix}.$$

Its eigenvalues satisfy the following characteristic equation:

$$\Phi^*(\lambda) := \lambda^5 + b_1 \lambda^4 + b_2 \lambda^3 + b_3 \lambda^2 + b_4 \lambda + b_5 = 0.$$

By Routh–Hurwitz criterion, we know that every root of $\Phi^*(\lambda) = 0$ has negative real part if and only if every determinant of the matrix H_i is positive, $i = 1, \dots, 5$, where

$$\begin{aligned}
 H_1 &= (b_1), \quad H_2 = \begin{pmatrix} b_1 & 1 \\ b_3 & b_2 \end{pmatrix}, \quad H_3 = \begin{pmatrix} b_1 & 1 & 0 \\ b_3 & b_2 & b_1 \\ b_5 & b_4 & b_3 \end{pmatrix}, \\
 H_4 &= \begin{pmatrix} b_1 & 1 & 0 & 0 \\ b_3 & b_2 & b_1 & 1 \\ b_5 & b_4 & b_3 & b_2 \\ 0 & 0 & b_5 & b_4 \end{pmatrix}, \quad H_5 = \begin{pmatrix} b_1 & 1 & 0 & 0 & 0 \\ b_3 & b_2 & b_1 & 1 & 0 \\ b_5 & b_4 & b_3 & b_2 & b_1 \\ 0 & 0 & b_5 & b_4 & b_3 \\ 0 & 0 & 0 & 0 & b_5 \end{pmatrix}.
 \end{aligned}$$

Note that all $b_i > 0$ ($i = 1, \dots, 5$) since all parameters are positive. Hence, condition (i) of the theorem implies that the determinant of H_2 is positive; condition (ii) ensures that the determinant of H_3 is positive; and condition (iii) guarantees that the determinant of H_4 is positive. Finally, the determinant of H_5 is positive if and

only if

$$b_5[(b_1 b_2 - b_3)(b_3 b_4 - b_2 b_5) - (b_5 - b_1 b_4)^2] > 0,$$

which follows from the fact that $b_5 > 0$ and condition (iii). The conclusion thus follows. \square

2. Remarks

The conditions (i)–(iii) can also be expressed as

$$\begin{cases} D := (b_5 - b_1 b_4)^2 (b_3^2 - 4b_1 b_5) > 0, \\ \max \left\{ \frac{b_3}{b_1}, \frac{b_3^2 - b_1(b_5 - b_1 b_4)}{b_1 b_3}, \frac{b_3(b_1 b_4 + b_5) - \sqrt{D}}{2b_1 b_5} \right\} \\ < b_2 < \frac{b_3(b_1 b_4 + b_5) + \sqrt{D}}{2b_1 b_5}. \end{cases}$$

Define $N(t) := S(t) + L(t) + I(t) + C(t) + R(t) + V(t)$. We obtain that

$$\frac{dN(t)}{dt} = \mu - \mu_0 N(t) - \mu_1 C(t),$$

which implies that

$$N(t) \leq \frac{\mu}{\mu_0} + e^{-\mu_0 t} \left(N(t_0) - \frac{\mu}{\mu_0} \right) \rightarrow \frac{\mu}{\mu_0}$$

as $t \rightarrow +\infty$. Thus, for any choice of parameter values satisfying our assumptions, $(S(t), L(t), I(t), C(t), R(t), V(t))$ will fall into the set $\{(S, L, I, C, R, V) \in R_+^6 : 0 < S + L + I + C + R + V \leq \mu/\mu_0\}$ in a finite time t .

3. HBV is endemic in China

We would like to point out that in Section 5 in Zou et al. (2010), we numerically simulated the data on acute hepatitis B reported by the Ministry of Chinese Health from 2003 to 2008 using the parameter values in Table 1. Since our simulation matched the reported data well, based on the model and the used parameter values, the basic reproduction number was estimated to be $R_0 = 2.406$. We then stated: ‘‘This indicates that hepatitis B is endemic in mainland China: it stabilizes and is

Table 1
Reported hepatitis B data in China, 2009–2010 (MOHC, 2012).

Year	2009	2010	2011
Cases	1 179 607	1 060 582	1 093 335

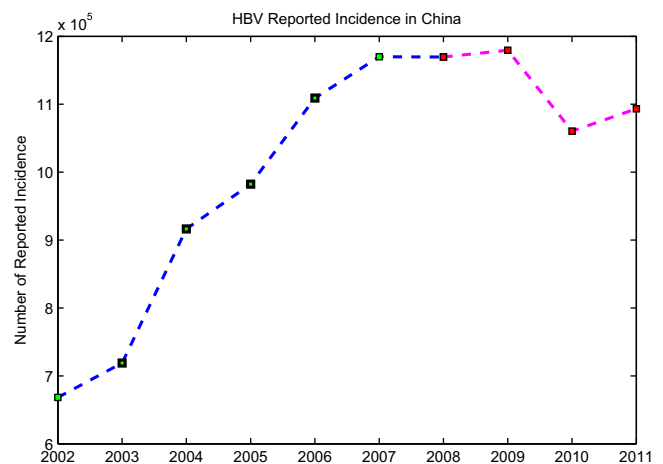


Fig. 1. Hepatitis B data reported by the Ministry of Health of China from 2002 to 2011.

approaching its equilibrium.” This conclusion (and those proposed control strategies) was made based on the evaluation and sensitivity analysis of the basic reproduction number and numerical simulations using reported data while the stability of the positive equilibrium E^* was never used.

The hepatitis B data reported by the Ministry of Chinese Health from 2009 to 2011 are available now (MOHC, 2012), see Table 1.

Fig. 1(a) in Zou et al. (2010) showed the hepatitis B data reported by the Ministry of Health of China from 2003 to 2008. Adding the data of 2002 and new data from 2009 to 2011 in Table 1 into Fig. 1(a) in Zou et al. (2010), we obtain the incidence

numbers of HBV in China since 2002 (Fig. 1), which further confirms our conclusion that hepatitis B is endemic in mainland China: it stabilizes and is approaching its equilibrium.

References

- MOHC, 2012. Ministry of Health of the People's Republic of China. <<http://www.moh.gov.cn/publicfiles/business/htmlfiles/zwgkzt/pyq/list.htm>>.
- Zou, L., 2010. Center and degenerate equilibria of differential equations and applications to biological dynamical systems. Ph.D. Thesis, Sichuan University.
- Zou, L., Zhang, W., Ruan, S., 2010. Modeling the transmission dynamics and control of hepatitis B virus in China. *J. Theor. Biol.* 262, 330–338.