# On the solvability of a mathematical model for prion proliferation 

Gieri Simonett, Christoph Walker*<br>Vanderbilt University, Department of Mathematics, 1326 Stevenson Center, Nashville, TN 37240, USA

Received 19 August 2005
Available online 18 January 2006
Submitted by M. Iannelli


#### Abstract

We show that a model describing the interaction between normal and infectious prion proteins admits global solutions. More precisely, supposing the involved degradation rates to be bounded, we prove global existence and uniqueness of classical solutions. Based on this existence theory, we provide sufficient conditions for the existence of global weak solutions in the case of unbounded splitting rates. Moreover, we prove global stability of the disease-free steady state.


© 2005 Elsevier Inc. All rights reserved.
Keywords: Global existence; Classical and weak solutions; Asymptotic behavior; Fragmentation; Prion proliferation

## 1. Introduction

The present paper aims at investigating mathematically a recent model that describes the dynamics of prion proliferation. Prions seem to be widely regarded as the infectious agent causing fatal diseases known as bovine spongiform encephalopathy (BSE) for cattle, scrapie for sheep, or Kuru and Creutzfeld-Jacob for humans. In this theory, prions are thought to be a polymeric form of a normal protein monomer $\operatorname{PrP}^{\mathrm{C}}$. The polymeric infectious prions $\mathrm{PrP}^{\mathrm{Sc}}$ have a tendency to attach units of $\mathrm{PrP}^{\mathrm{C}}$ in a stringlike formation, converting the latter to the infectious form. This mechanism makes $\mathrm{PrP}^{\mathrm{Sc}}$ polymers more stable and is called nucleated polymerization. Above some critical size, $\mathrm{PrP}^{\mathrm{Sc}}$ is very stable and polymerizes rapidly to form chains, possibly involv-

[^0]ing several thousands of monomer units. Nevertheless, $\operatorname{PrP}{ }^{\mathrm{Sc}}$ prions also can split, usually into smaller infectious prions. However, if a polymer falls below the critical size, it degrades immediately into $\operatorname{PrP}^{\mathrm{C}}$ monomers.

A model for nucleated polymerization has recently been proposed in [4,5] (see also the references therein) describing the mechanism by which prions are hypothesized to replicate. Denoting the number of $\operatorname{PrP}^{\mathrm{C}}$ monomers at time $t \geqslant 0$ by $v(t) \geqslant 0$ and introducing a population density $u=u(t, y) \geqslant 0$ for the infectious $\operatorname{PrP}^{S c}$ polymers at time $t \geqslant 0$ and size $y$ greater than the minimum length $y_{0}>0$, the interaction of the $\mathrm{PrP}^{\mathrm{C}}$ monomers and the $\mathrm{PrP}^{\mathrm{Sc}}$ polymers can be described by the coupled system consisting of the ordinary differential equation

$$
\begin{equation*}
\dot{v}=\lambda-\gamma v-\tau v \int_{y_{0}}^{\infty} u(t, y) \mathrm{d} y+2 \int_{y_{0}}^{\infty} u(t, y) \beta(y) \int_{0}^{y_{0}} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \tag{1}
\end{equation*}
$$

and the partial differential equation

$$
\begin{equation*}
\partial_{t} u+\tau v(t) \partial_{y} u=-(\mu(y)+\beta(y)) u(y)+2 \int_{y}^{\infty} \beta\left(y^{\prime}\right) \kappa\left(y, y^{\prime}\right) u\left(y^{\prime}\right) \mathrm{d} y^{\prime} \tag{2}
\end{equation*}
$$

for $y \in\left(y_{0}, \infty\right)$ subject to the boundary condition

$$
\begin{equation*}
u\left(t, y_{0}\right)=0, \quad t>0 \tag{3}
\end{equation*}
$$

These equations are supplemented with the initial conditions

$$
\begin{equation*}
v(0)=v^{0}, \quad u(0, y)=u^{0}(y), \quad y \in\left(y_{0}, \infty\right) \tag{4}
\end{equation*}
$$

Equation (1) includes a source term $\lambda \geqslant 0$, while the term $-\gamma v(t)$, with $\gamma \geqslant 0$, takes into account metabolic degradation of monomers. The constant $\tau>0$ denotes the polymerization rate. Moreover, $\beta(y) \geqslant 0$ is the length-dependent fragmentation rate of polymers of size $y>y_{0}$, and $\kappa\left(y^{\prime}, y\right)$ is the probability of a polymer of size $y>y_{0}$ splitting into two pieces $y^{\prime}<y$ and $y-y^{\prime}<y$. The transport term $\tau v(t) \partial_{y} u(t, y)$ in Eq. (2) accounts for the loss of polymers of size $y$ due to lengthening. A loss of polymers according to metabolic degradation is reflected by the term $-\mu(y) u(y)$. Finally, the terms involving $\beta$ on the right-hand side of Eq. (2) represent the loss and gain of $\mathrm{PrP}^{\mathrm{Sc}}$ polymers caused by splitting. For a more detailed explanation of each process we refer to $[4,5]$ and the references therein.

Let us point out that (1), (2) is a coupled system of non-linear, non-local equations. In order to solve this equations, we employ Kato's theory for hyperbolic evolution equations. That is, given a function $v$ with appropriate regularity properties, we construct an evolution system for the partial differential equation (2). We should remark that in the absence of the kernel operator on the right-hand side of (2), an evolution system can readily be obtained by using the method of characteristics.

It should also be pointed out that Eqs. (1), (2) can be handled as an abstract quasilinear hyperbolic system in order to obtain local existence, see for instance [9, §6.4]. However, this approach does not seem to yield optimal results for Eqs. (1), (2).

Before outlining the contents of this paper, we summarize the present-state of knowledge on the above model. It seems that only kernels of the form

$$
\begin{equation*}
\mu \equiv \mathrm{const}, \quad \beta(y)=\beta y, \quad \kappa\left(y^{\prime}, y\right)=\frac{1}{y} \tag{5}
\end{equation*}
$$

have been considered so far. This choice of kernels leads to a closed system of ordinary differential equations for $v$ and

$$
U(t):=\int_{y_{0}}^{\infty} u(t, y) \mathrm{d} y, \quad P(t):=\int_{y_{0}}^{\infty} y u(t, y) \mathrm{d} y .
$$

Indeed, (1) reduces to

$$
\begin{equation*}
\dot{v}=\lambda-\gamma v-\tau v U+\beta y_{0}^{2} U \tag{6}
\end{equation*}
$$

and integrating (2) yields the equations

$$
\begin{align*}
\dot{U} & =\beta P-\mu U-2 \beta y_{0} U  \tag{7}\\
\dot{P} & =\tau v U-\mu P-\beta y_{0}^{2} U \tag{8}
\end{align*}
$$

which, together with (6), are uniquely globally solvable. In addition, it has been shown in [5] that the disease-free steady state $(v, U, P)=(\lambda / \gamma, 0,0)$ for Eqs. (6)-(8) is globally stable provided

$$
\begin{equation*}
\beta y_{0}+\mu>\sqrt{\frac{\beta \lambda \tau}{\gamma}} \tag{9}
\end{equation*}
$$

If one reverses the strict inequality sign in (9) it has also been proved in [5] that there exists a prion disease steady state which is locally asymptotically stable. These results have been improved in [10] in that the disease-free steady state is globally asymptotically stable also for an equality sign in (9) and in that the disease steady state is even globally asymptotically stable for (9) with a reversed strict inequality sign.

Observe that the solvability of (6)-(8) implies that the original equations (1), (2) are no longer coupled since $v$ is completely determined for all $t \geqslant 0$. Hence, as shown in [3], the partial differential equation (2) (with kernels as in (5)) can be solved for $u=u(t, y)$ by using the method of characteristics combined with semigroup theory. Moreover, it has also been shown in [3] that $u$ converges either to 0 or to the disease steady state according to whether or not (9) holds.

Our aim is to consider quite general kernels, merely assuming suitable growth conditions. More precisely, after collecting some auxiliary results in Section 2, we show in Section 3 that (1)-(4) is globally well-posed provided $\mu$ and $\beta$ are bounded, see Theorem 3.1. The basic idea is to solve Eq. (1) for a fixed, suitable function $\bar{u}$ and then to substitute the obtained solution $v_{\bar{u}}$ into Eq. (2). Using Kato's theory for hyperbolic evolution equations, we solve then Eq. (2) in order to obtain a classical solution $u_{\bar{u}}$. A fixed point argument for the map $\bar{u} \mapsto u_{\bar{u}}$ yields then local existence and uniqueness of a solution pair $(v, u)$ for (1)-(4). Suitable a priori estimates guarantee global existence. A weak formulation of (2) allows then to extend in Section 4 the existence results to unbounded kernels by using a weak compactness method, see Theorem 4.3. We also prove finite speed of propagation for the weak (and classical) solutions to (2). Finally, in Section 5 we show that the disease-free steady state is globally asymptotically stable provided some suitable lower and upper bounds for the splitting kernels are available. We refer to Theorem 5.3 for a precise statement.

Clearly, the method described above does not yield uniqueness of weak solutions. This issue will be the topic of future work [8].

## 2. Preliminaries

In the following, we set $Y:=\left(y_{0}, \infty\right)$ and assume that

$$
\begin{equation*}
\mu, \beta \in L_{\infty}^{+}(Y) \tag{10}
\end{equation*}
$$

where $L_{\infty}^{+}(Y)$ stands for the positive cone in $L_{\infty}(Y)$. We also assume that $\kappa \geqslant 0$ is measurable on $K:=\left\{\left(y^{\prime}, y\right) ; y_{0}<y<\infty, 0<y^{\prime}<y\right\}$ and satisfies

$$
\begin{equation*}
\kappa\left(y^{\prime}, y\right)=\kappa\left(y-y^{\prime}, y\right), \quad\left(y^{\prime}, y\right) \in K, \tag{11}
\end{equation*}
$$

which means binary splitting. Moreover, we suppose the number of monomer units to be preserved during splitting, that is,

$$
\begin{equation*}
2 \int_{0}^{y} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}=y, \quad \text { a.e. } y \in Y \tag{12}
\end{equation*}
$$

Furthermore, we let

$$
\begin{equation*}
\tau>0, \quad \lambda, \gamma \geqslant 0 \tag{13}
\end{equation*}
$$

It is easy to check that (11), (12) imply

$$
\begin{equation*}
\int_{0}^{y} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}=1, \quad \text { a.e. } y \in Y . \tag{14}
\end{equation*}
$$

Observe that the natural constraints (11), (12) hold if $\kappa$ is of self-similar form

$$
\begin{equation*}
\kappa\left(y^{\prime}, y\right)=\frac{1}{y} \kappa_{0}\left(\frac{y^{\prime}}{y}\right), \quad y>y_{0}, 0<y^{\prime}<y \tag{15}
\end{equation*}
$$

where $\kappa_{0}$ is a non-negative integrable function defined on $(0,1)$ such that

$$
\begin{equation*}
\kappa_{0}(y)=\kappa_{0}(1-y), \quad y \in(0,1), \quad \int_{0}^{1} \kappa_{0}(y) \mathrm{d} y=1 \tag{16}
\end{equation*}
$$

This allows to capture $\kappa$ in (5) by taking $\kappa_{0} \equiv 1$. Also note that the operator $L$, given by

$$
\begin{equation*}
L[u](y):=-(\mu(y)+\beta(y)) u(y)+2 \int_{y}^{\infty} \beta\left(y^{\prime}\right) \kappa\left(y, y^{\prime}\right) u\left(y^{\prime}\right) \mathrm{d} y^{\prime}, \quad \text { a.e. } y \in Y, \tag{17}
\end{equation*}
$$

defines a linear and bounded operator from $L_{1}(Y, y \mathrm{~d} y)$ into itself according to (10)-(12) and that

$$
\begin{align*}
\int_{y_{0}}^{\infty} \varphi(y) L[u](y) \mathrm{d} y= & -\int_{y_{0}}^{\infty} \varphi(y) \mu(y) u(y) \mathrm{d} y \\
& +\int_{y_{0}}^{\infty} u(y) \beta(y)\left(-\varphi(y)+2 \int_{y_{0}}^{y} \varphi\left(y^{\prime}\right) \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}\right) \mathrm{d} y \tag{18}
\end{align*}
$$

for $u \in L_{1}(Y, y \mathrm{~d} y)$ and a suitable test function $\varphi$. We then put

$$
E_{0}:=L_{1}(Y, y \mathrm{~d} y) \quad \text { and } \quad E_{1}:=\stackrel{\circ}{W}_{1}^{1}(Y, y \mathrm{~d} y):=\mathrm{cl}_{W_{1}^{1}(Y, y \mathrm{~d} y)} \mathcal{D}(Y)
$$

where $\mathcal{D}(Y)$ denotes the space of all test functions on $Y$. By $E_{0}^{+}$we mean the positive cone in $E_{0}$ and $E_{1}^{+}:=E_{1} \cap E_{0}^{+}$. Finally, given any interval $J$ and any function $v: J \rightarrow \mathbb{R}^{+}$, we define

$$
\begin{equation*}
\mathbb{A}_{v}(t) u:=\tau v(t) \partial_{y} u-L[u], \quad u \in E_{1}, t \in J . \tag{19}
\end{equation*}
$$

Lemma 2.1. The operator $-A$, defined as

$$
\begin{equation*}
A \varphi:=\partial_{y} \varphi, \quad \varphi \in E_{1}, \tag{20}
\end{equation*}
$$

generates a strongly continuous semigroup $\left\{e^{-t A} ; t \geqslant 0\right\}$ on $E_{0}$. It is given by

$$
\left[e^{-t A} \varphi\right](y)=\left\{\begin{array}{ll}
\varphi(y-t), & y>y_{0}+t,  \tag{21}\\
0, & y_{0}<y \leqslant y_{0}+t,
\end{array} \quad t \geqslant 0\right.
$$

and satisfies

$$
\begin{equation*}
\left\|e^{-t A}\right\|_{\mathcal{L}\left(E_{0}\right)} \leqslant e^{t / y_{0}}, \quad t \geqslant 0 \tag{22}
\end{equation*}
$$

Proof. Clearly, (21) defines a strongly continuous semigroup on $E_{0}$ satisfying

$$
\left\|e^{-t A} \varphi\right\|_{E_{0}} \leqslant\left(1+\frac{t}{y_{0}}\right)\|\varphi\|_{E_{0}} \leqslant e^{t / y_{0}}\|\varphi\|_{E_{0}}, \quad t \geqslant 0
$$

for $\varphi \in E_{0}$, whence (22). It thus remains to show that its generator $-A$ is indeed given by (20). Note that Lebesgue's theorem guarantees that the test functions are contained in the domain of $A$ and that

$$
\begin{equation*}
A \varphi=\partial_{y} \varphi, \quad \varphi \in \mathcal{D}(Y) \tag{23}
\end{equation*}
$$

Since (21) is a right translation, $\mathcal{D}(Y)$ is invariant under $e^{-t A}$ and therefore is a core for $A$. In particular, $\mathcal{D}(Y)$ is dense in the domain of $A$, which, together with (23), easily yields (20).

In the sequel, we set $J_{T}:=[0, T]$ for $T>0$ and, given $R>1$, we define

$$
\begin{equation*}
\mathcal{V}_{T, R}:=\left\{v \in C^{1}\left(J_{T}\right) ; R^{-1} \leqslant v(t) \leqslant\|v\|_{C^{1}\left(J_{T}\right)} \leqslant R\right\} . \tag{24}
\end{equation*}
$$

Recall then that the operator $\mathbb{A}_{v}(t)$ has been defined in (19).
Proposition 2.2. Fix $R>1, T_{0}>0$ and let $0<T \leqslant T_{0}$. Then $\left(-\mathbb{A}_{v}(t)\right)_{t \in[0, T]}$ generates for each $v \in \mathcal{V}_{T, R}$ a unique evolution system $U_{v}(t, s), 0 \leqslant s \leqslant t \leqslant T$, in $E_{0}$, and there exists a constant $\omega_{0}:=\omega_{0}\left(T_{0}, R\right)>0$ such that

$$
\begin{equation*}
\left\|U_{v}(t, s)\right\|_{\mathcal{L}\left(E_{0}\right)} \leqslant e^{\omega_{0}(t-s)}, \quad 0 \leqslant s \leqslant t \leqslant T, v \in \mathcal{V}_{T, R} \tag{25}
\end{equation*}
$$

and

$$
\begin{equation*}
\left\|U_{v}(t, s)\right\|_{\mathcal{L}\left(E_{1}\right)} \leqslant \omega_{0}, \quad 0 \leqslant s \leqslant t \leqslant T, v \in \mathcal{V}_{T, R} \tag{26}
\end{equation*}
$$

Moreover, for $v, w \in \mathcal{V}_{T, R}$, it holds that

$$
\begin{equation*}
\left\|U_{v}(t, s)-U_{w}(t, s)\right\|_{\mathcal{L}\left(E_{1}, E_{0}\right)} \leqslant \omega_{0}(t-s)\|v-w\|_{C\left(J_{T}\right)}, \quad 0 \leqslant s \leqslant t \leqslant T \tag{27}
\end{equation*}
$$

Proof. Since $L$ is a bounded operator on $E_{0}$, Lemma 2.1 and a well-known perturbation result (see [9, Theorem 3.1.1]) ensure that, for any fixed $v \in \mathcal{V}_{T, R}$ and any $s \in J_{T},-\mathbb{A}_{v}(s)$ generates a strongly continuous semigroup on $E_{0}$ with

$$
\begin{equation*}
\left\|e^{-t \mathbb{A}_{v}(s)}\right\|_{\mathcal{L}\left(E_{0}\right)} \leqslant e^{\bar{\omega} t}, \quad t \geqslant 0 \tag{28}
\end{equation*}
$$

where $\bar{\omega}:=\tau R y_{0}^{-1}+\|L\|_{\mathcal{L}\left(E_{0}\right)}$. Hence, putting $\omega:=\bar{\omega}+1$ it follows that $\left\{\mathbb{A}_{v}(t)\right\}_{t \in J_{T}}$ is stable in the sense of $[9, \S 5.2]$ for each $v \in \mathcal{V}_{T, R}$. Next, given any $t \in J_{T}$, the definition $Q_{v}(t):=\omega+\mathbb{A}_{v}(t)$ yields an isomorphism from $E_{1}$ onto $E_{0}$ satisfying

$$
\begin{equation*}
\left\|Q_{v}(t)\right\|_{\mathcal{L}\left(E_{1}, E_{0}\right)} \leqslant \omega+\tau R+\|L\|_{\mathcal{L}\left(E_{0}\right)}, \quad t \in J_{T}, v \in \mathcal{V}_{T, R} \tag{29}
\end{equation*}
$$

Moreover, for $u \in E_{1}$,

$$
Q_{v}(\cdot) u \in C^{1}\left(J_{T}, E_{0}\right) \quad \text { with } \dot{Q}_{v}(t) u:=\frac{\mathrm{d}}{\mathrm{~d} t} Q_{v}(t) u=\tau \dot{v}(t) \partial_{y} u
$$

Therefore, assumptions $\left(H_{1}\right),\left(H_{2}\right)^{+},\left(H_{3}\right)$ of $[9, \S 5]$ hold, thus implying that there indeed exists a unique evolution system $U_{v}(t, s), 0 \leqslant s \leqslant t \leqslant T$, in $E_{0}$ for each $v \in \mathcal{V}_{T, R}$, which, in addition, satisfies statements $\left(E_{1}\right)-\left(E_{5}\right)$ of $[9, \S 5]$. In particular, (25) holds (with $\omega_{0}$ replaced by $\left.\bar{\omega}\right)$.

We now refer to the proof of [9, Theorem 5.4.6]: The evolution system $U_{v}(t, s)$ can be written as

$$
\begin{equation*}
U_{v}(t, s)=Q_{v}(t)^{-1} W_{v}(t, s) Q_{v}(s), \quad 0 \leqslant s \leqslant t \leqslant T \tag{30}
\end{equation*}
$$

where $W_{v}(t, s) \in \mathcal{L}\left(E_{0}\right)$ satisfies

$$
W_{v}(t, s) u=U_{v}(t, s) u+\int_{s}^{t} W_{v}(t, r) C_{v}(r) U_{v}(r, s) u \mathrm{~d} r
$$

for $0 \leqslant s \leqslant t \leqslant T$ and $u \in E_{0}$ with

$$
C_{v}(t):=\dot{Q}_{v}(t) Q_{v}(t)^{-1} \in \mathcal{L}\left(E_{0}\right), \quad t \in J_{T} .
$$

We then claim that there is a constant $c_{0}(R)>0$ such that

$$
\begin{equation*}
\left\|Q_{v}(t)^{-1}\right\|_{\mathcal{L}\left(E_{0}, E_{1}\right)} \leqslant c_{0}(R), \quad t \in J_{T}, v \in \mathcal{V}_{T, R} \tag{31}
\end{equation*}
$$

Indeed, (28) implies

$$
\left\|Q_{v}(t)^{-1}\right\|_{\mathcal{L}\left(E_{0}\right)} \leqslant 1, \quad t \in J_{T}
$$

and therefore, for $u \in E_{0}$ and $t \in J_{T}$,

$$
\begin{aligned}
\left\|Q_{v}(t)^{-1} u\right\|_{E_{1}} & =\left\|Q_{v}(t)^{-1} u\right\|_{E_{0}}+\left\|\partial_{y} Q_{v}(t)^{-1} u\right\|_{E_{0}} \\
& \leqslant\|u\|_{E_{0}}+\frac{1}{\tau v(t)}\left\|u-(\omega-L) Q_{v}(t)^{-1} u\right\|_{E_{0}} \\
& \leqslant\left(1+R / \tau\left(1+\omega+\|L\|_{\mathcal{L}\left(E_{0}\right)}\right)\right)\|u\|_{E_{0}}
\end{aligned}
$$

whence (31). Consequently, we have

$$
\begin{aligned}
\left\|C_{v}(t)\right\|_{\mathcal{L}\left(E_{0}\right)} & \leqslant\left\|\dot{Q}_{v}(t)\right\|_{\mathcal{L}\left(E_{1}, E_{0}\right)}\left\|Q_{v}(t)^{-1}\right\|_{\mathcal{L}\left(E_{0}, E_{1}\right)} \\
& \leqslant \tau\|\dot{v}\|_{C\left(J_{T}\right)} c_{0}(R) \leqslant c_{0}^{\prime}(R)
\end{aligned}
$$

for $t \in J_{T}$ and $v \in \mathcal{V}_{T, R}$. From the proof of [9, Lemma 5.4.5] (see in particular Eq. (4.11) therein) and from (25) it thus follows that there exists a constant $c\left(T_{0}, R\right)>0$ such that

$$
\begin{equation*}
\left\|W_{v}(t, s)\right\|_{\mathcal{L}\left(E_{0}\right)} \leqslant c\left(T_{0}, R\right), \quad 0 \leqslant s \leqslant t \leqslant T, v \in \mathcal{V}_{T, R} \tag{32}
\end{equation*}
$$

Applying estimates (29), (31), and (32) to (30) we conclude that (26) is true.
Finally, let $v, w \in \mathcal{V}_{T, R}$ and $u \in E_{1}$ be arbitrary. Then, for $0 \leqslant s<t \leqslant T$,

$$
N:=\left[\sigma \mapsto U_{v}(t, \sigma) U_{w}(\sigma, s) u\right] \in C^{1}\left((s, t), E_{0}\right) \cap C\left([s, t], E_{1}\right)
$$

by $\left(E_{2}\right)-\left(E_{5}\right)$ in $[9, \S 5]$ with

$$
\dot{N}(\sigma)=U_{v}(t, \sigma)\left(\mathbb{A}_{v}(\sigma)-\mathbb{A}_{w}(\sigma)\right) U_{w}(\sigma, s) u
$$

Therefore, (25) and (26) yield

$$
\begin{aligned}
& \left\|U_{w}(t, s) u-U_{v}(t, s) u\right\|_{E_{0}} \\
& \quad \leqslant \int_{s}^{t}\left\|U_{v}(t, \sigma)\right\|_{\mathcal{L}\left(E_{0}\right)}\left\|\mathbb{A}_{v}(\sigma)-\mathbb{A}_{w}(\sigma)\right\|_{\mathcal{L}\left(E_{1}, E_{0}\right)}\left\|U_{w}(\sigma, s)\right\|_{\mathcal{L}\left(E_{1}\right)} \mathrm{d} \sigma\|u\|_{E_{1}} \\
& \quad \leqslant c\left(T_{0}, R\right)(t-s)\|v-w\|_{C\left(J_{T}\right)}\|u\|_{E_{1}}
\end{aligned}
$$

for $0 \leqslant s \leqslant t \leqslant T$, hence statement (27).
Remark 2.3. As observed in the previous proof, the evolution system $U_{v}(t, s), 0 \leqslant s \leqslant t \leqslant T$, corresponding to $v \in \mathcal{V}_{T, R}$ satisfies $\left(E_{1}\right)-\left(E_{5}\right)$ in [9, §5]. In particular, we have for $u^{0} \in E_{1}$ that

$$
\left[t \mapsto U_{v}(t, 0) u^{0}\right] \in C^{1}\left(J_{T}, E_{0}\right) \cap C\left(J_{T}, E_{1}\right)
$$

The existence of weak solutions will require the following auxiliary result.
Lemma 2.4. For $v \in C\left(J_{T}\right)$ with $v(t) \geqslant 0$ put $A_{v}(t):=\tau v(t) \partial_{y}, t \in J_{T}$, and let $U_{A_{v}}(t, s)$, $0 \leqslant s \leqslant t \leqslant T$, be the corresponding evolution system in $L_{1}(Y)$. Then, for any $\delta>0$, it holds that

$$
\sup _{|\mathcal{E}| \leqslant \delta} \int_{\mathcal{E}} U_{A_{v}}(t, s) \varphi \mathrm{d} y \leqslant \sup _{|\mathcal{E}| \leqslant \delta} \int_{\mathcal{E}} \varphi \mathrm{d} y, \quad 0 \leqslant s \leqslant t \leqslant T, \varphi \in L_{1}^{+}(Y),
$$

the supremum being taken over all measurable sets $\mathcal{E} \subset Y$.
Proof. Noticing that $-\partial_{y}$ with domain $\stackrel{\circ}{W}_{1}^{1}(Y)$ generates a strongly continuous positive semigroup of contractions on $L_{1}(Y)$ given as in (21), it follows that

$$
\left\|e^{-t A_{v}(s)}\right\|_{\mathcal{L}\left(L_{1}(Y)\right)} \leqslant 1, \quad\left\|e^{-t A_{v}(s)}\right\|_{\mathcal{L}\left(\dot{W}_{1}^{1}(Y)\right)} \leqslant 1, \quad t \geqslant 0, s \in J_{T} .
$$

Hence, the corresponding evolution system $U_{A_{v}}(t, s), 0 \leqslant s \leqslant t \leqslant T$, in $L_{1}(Y)$ is well defined according to [9, Theorems 5.2.2, 5.3.1]. Let then $\mathcal{E} \subset Y$ be any measurable subset of $Y$ with measure $|\mathcal{E}| \leqslant \delta$ and choose $\varphi \in L_{1}^{+}(Y)$. Denoting by $\chi_{S}$ the characteristic function on a set $S$, we have

$$
\int_{\mathcal{E}}\left[e^{-t A_{v}(s)} \varphi\right](y) \mathrm{d} y=\int_{y_{0}}^{\infty} \chi_{\{-t \tau v(s)+\mathcal{E}\}}(y) \varphi(y) \mathrm{d} y \leqslant \sup _{\left|\mathcal{E}^{\prime}\right| \leqslant \delta} \int_{\mathcal{E}^{\prime}} \varphi(y) \mathrm{d} y
$$

for $s \in J_{T}$ and $t \geqslant 0$. From Eqs. (3.5) and (3.15) in [9, §5] we thus deduce

$$
\int_{\mathcal{E}} U_{A_{v}}(t, s) \varphi \mathrm{d} y \leqslant \sup _{\left|\mathcal{E}^{\prime}\right| \leqslant \delta} \int_{\mathcal{E}^{\prime}} \varphi \mathrm{d} y, \quad 0 \leqslant s \leqslant t \leqslant T,
$$

and the assertion follows.

## 3. Classical solutions

In this section we show that problem (1)-(4) is globally well-posed for bounded kernels $\mu$ and $\beta$. In order to do this, let us denote by $|\cdot|_{1}$ the norm in $L_{1}(Y)$ and put

$$
g(u):=2 \int_{y_{0}}^{\infty} u(y) \beta(y) \int_{0}^{y_{0}} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y .
$$

Defining $L$ by (17) and $\mathbb{A}_{v}(t)$ by (19), we may rewrite (1)-(4) as

$$
\begin{equation*}
\dot{v}=\lambda-\gamma v-\tau v|u|_{1}+g(u), \quad t>0, \quad v(0)=v^{0}, \tag{33}
\end{equation*}
$$

provided $u \geqslant 0$, and

$$
\begin{equation*}
\dot{u}+\mathbb{A}_{v}(t) u=0, \quad t>0, \quad u(0)=u^{0} . \tag{34}
\end{equation*}
$$

Theorem 3.1. Suppose (10)-(13) hold. Then, given any $v^{0}>0$ and $u^{0} \in E_{1}^{+}$, problem (33), (34) possesses a unique global classical solution $(v, u)$ such that $v \in C^{1}\left(\mathbb{R}^{+}\right), v(t)>0$ for $t>0$, and $u \in C^{1}\left(\mathbb{R}^{+}, E_{0}\right) \cap C\left(\mathbb{R}^{+}, E_{1}^{+}\right)$.

Proof. (i) We first prove that, for any $S>0$, there exists $T:=T(S) \in(0,1]$ such that (33), (34) possesses a unique solution $(v, u)$ on $J_{T}$ with regularity properties as stated in the theorem, provided that $\left(v^{0}, u^{0}\right) \in \mathbb{R}^{+} \times E_{1}^{+}$satisfies

$$
\begin{equation*}
S^{-1} \leqslant v^{0} \quad \text { and } \quad v^{0}+\left\|u^{0}\right\|_{E_{1}} \leqslant S \tag{35}
\end{equation*}
$$

In the following, we denote by $c(S)>0$ a generic constant depending on $S$ but not on $T \in(0,1]$. Let us then define the complete metric space

$$
X_{T}:=\left\{u \in C\left(J_{T}, E_{0}^{+}\right) ;\|u(t)\|_{E_{0}} \leqslant S+1, t \in J_{T}\right\},
$$

and let us choose $\bar{u} \in X_{T}$ arbitrarily. Then, since $g(\bar{u}),|\bar{u}|_{1} \in C\left(J_{T}\right)$ due to (12), it follows that (33), with $u$ replaced by $\bar{u}$, admits a unique solution $v_{\bar{u}} \in C^{1}\left(J_{T}\right)$. Clearly,

$$
v_{\bar{u}}(t)=e^{-\gamma t-\tau \int_{0}^{t}|\bar{u}(\sigma)|_{1} \mathrm{~d} \sigma} v^{0}+\int_{0}^{t} e^{-\gamma(t-s)-\tau \int_{s}^{t}|\bar{u}(\sigma)|_{1} \mathrm{~d} \sigma}(\lambda+g(\bar{u}(s))) \mathrm{d} s
$$

for $t \in J_{T}$, hence

$$
\begin{equation*}
v_{\bar{u}}(t) \geqslant e^{-\gamma t-\tau / y_{0}(S+1) t} v^{0} \geqslant c(S), \quad 0 \leqslant t \leqslant T \leqslant 1 . \tag{36}
\end{equation*}
$$

Moreover, since $v^{0} \leqslant S$ and $g(\bar{u}(t)) \leqslant\|\beta\|_{\infty}(S+1)$ for $t \in J_{T}$, we deduce

$$
\begin{equation*}
v_{\bar{u}}(t) \leqslant c(S), \quad t \in J_{T}, \tag{37}
\end{equation*}
$$

from which it follows

$$
\begin{equation*}
-c(S) \leqslant-\left(\gamma+\tau|\bar{u}(t)|_{1}\right) v_{\bar{u}}(t) \leqslant \dot{v}_{\bar{u}}(t) \leqslant \lambda+g(\bar{u}(t)) \leqslant c(S), \quad t \in J_{T} . \tag{38}
\end{equation*}
$$

Therefore, (36)-(38) entail the existence of $R:=R(S)>1$, depending on $S>0$ but not on $T \in(0,1]$, such that $v_{\bar{u}} \in \mathcal{V}_{T, R}$ whenever $\bar{u} \in X_{T}$, where $\mathcal{V}_{T, R}$ is given by (24). Furthermore, we readily derive from the explicit representation of $v_{\bar{u}}$ and the linearity of $g$ that

$$
\begin{equation*}
\left|v_{\bar{u}_{1}}(t)-v_{\bar{u}_{2}}(t)\right| \leqslant c(S)\left\|\bar{u}_{1}-\bar{u}_{2}\right\|_{X_{T}}, \quad 0 \leqslant t \leqslant T \leqslant 1, \bar{u}_{1}, \bar{u}_{2} \in X_{T} . \tag{39}
\end{equation*}
$$

Let $U_{v_{\bar{u}}}(t, s), 0 \leqslant s \leqslant t \leqslant T$, denote the unique evolution system in $E_{0}$ corresponding to $\left\{\mathbb{A}_{v_{\bar{u}}}(t)\right\}_{t \in J_{T}}$ and by $\omega_{0}=\omega_{0}(1, R(S))$ the constant occurring in Proposition 2.2. Defining

$$
\Lambda(\bar{u})(t):=U_{v_{\bar{u}}}(t, 0) u^{0}, \quad t \in J_{T}, \bar{u} \in X_{T},
$$

we obtain by Remark 2.3 the unique solution in $C\left(J_{T}, E_{1}\right) \cap C^{1}\left(J_{T}, E_{0}\right)$ to

$$
\dot{u}+\mathbb{A}_{v_{\bar{u}}}(t) u=0, \quad t>0, \quad u(0)=u^{0} .
$$

Next we show that $\Lambda: X_{T} \rightarrow X_{T}$ is a contraction, which, consequently, would imply our first claim. Provided $T:=T(S) \in(0,1]$ is chosen sufficiently small, we deduce from (25) that, for $\bar{u} \in X_{T}$ and $t \in J_{T}$,

$$
\|\Lambda(\bar{u})(t)\|_{E_{0}} \leqslant e^{\omega_{0} T}\left\|u^{0}\right\|_{E_{0}} \leqslant S+1,
$$

and (27) and (39) ensure for $\bar{u}_{1}, \bar{u}_{2} \in X_{T}$ and $t \in J_{T}$,

$$
\left\|\Lambda\left(\bar{u}_{1}\right)(t)-\Lambda\left(\bar{u}_{2}\right)(t)\right\|_{E_{0}} \leqslant \omega_{0} T\left\|v_{\bar{u}_{1}}-v_{\bar{u}_{2}}\right\|_{C\left(J_{T}\right)}\left\|u^{0}\right\|_{E_{1}} \leqslant \frac{1}{2}\left\|\bar{u}_{1}-\bar{u}_{2}\right\|_{X_{T}}
$$

In order to prove that $\Lambda(\bar{u})(t)$ is non-negative observe that $\Lambda(\bar{u})$ also solves

$$
\dot{u}+\left(A_{v_{\bar{u}}}(t)+r\right) u=L[u]+r u=: B(u), \quad t>0, \quad u(0)=u^{0}
$$

with $r:=\|\mu+\beta\|_{\infty}$ and $A_{v_{\bar{u}}}(t):=\tau v_{\bar{u}}(t) \partial_{y}$. Then $B(u) \in E_{0}^{+}$for $u \in E_{0}^{+}$. Since Lemma 2.1 ensures that $-A_{v_{\bar{u}}}(s)$ generates a positive semigroup on $E_{0}$, it readily follows from the proof of [9, Theorem 5.3.1] that the evolution system $\bar{U}(t, s)$ generated by $\left\{A_{v_{\bar{u}}}(t)+r\right\}_{t \in J_{T}}$ is positive. Defining then

$$
F(w)(t):=\bar{U}(t, 0) u^{0}+\int_{0}^{t} \bar{U}(t, s) B(w(s)) \mathrm{d} s,
$$

one shows that $\underset{\tilde{T}}{ }$ is a contraction from a suitable closed ball in $C\left([0, \tilde{T}], E_{0}\right)$, containing $u^{0}$, into itself provided $\tilde{T} \in(0, T]$ is sufficiently small. Hence, putting

$$
u_{0}:=u^{0}, \quad u_{n+1}:=F\left(u_{n}\right), \quad n \in \mathbb{N},
$$

we obtain a sequence in $C\left([0, \tilde{T}], E_{0}^{+}\right)$that converges to $\left.\Lambda(\bar{u})\right|_{[0, \tilde{T}]}$. This shows that

$$
T^{\star}:=\sup \left\{T^{\prime} \in(0, T] ; \Lambda(\bar{u})(t) \in E_{0}^{+}, 0 \leqslant t \leqslant T^{\prime}\right\} \geqslant \tilde{T}
$$

Assuming $T^{\star}<T$, a repetition of the above arguments with $u^{0}$ replaced by $\Lambda(\bar{u})\left(T^{\star}\right) \in E_{1}^{+}$ would lead to a contradiction. Thus $T^{\star}=T$, which entails that $\Lambda: X_{T} \rightarrow X_{T}$ is indeed a contraction.
(ii) It follows from part (i) that (33), (34) admit a unique maximal solution

$$
(v, u) \in C\left(J, \mathbb{R}^{+} \times E_{1}^{+}\right) \cap C^{1}\left(J, \mathbb{R} \times E_{0}\right),
$$

where $J$ is open in $\mathbb{R}^{+}$. We claim that, if $t^{+}:=\sup J<\infty$, then

For, suppose to the contrary that there are $t_{j} \nearrow t^{+}<\infty$ and $S>0$ such that

$$
v\left(t_{j}\right) \geqslant S^{-1} \quad \text { and } \quad v\left(t_{j}\right)+\left\|u\left(t_{j}\right)\right\|_{E_{1}} \leqslant S
$$

Let $T(S)>0$ be the corresponding constant from part (i) and fix $t_{N}>t^{+}-T(S)$. Then we may choose $\left(v\left(t_{N}\right), u\left(t_{N}\right)\right) \in \mathbb{R}^{+} \backslash\{0\} \times E_{1}^{+}$as initial value and deduce that the solution $(v, u)$ can be extended to a solution on $\left[0, t_{N}+T(S)\right]$, contradicting its maximality.
(iii) We now show that (40) does not occur in finite time. Observe that (12) and (18) imply

$$
\begin{equation*}
\dot{v}(t)+\frac{\mathrm{d}}{\mathrm{~d} t} \int_{y_{0}}^{\infty} y u(t, y) \mathrm{d} y=\lambda-\gamma v(t)-\int_{y_{0}}^{\infty} y \mu(y) u(t, y) \mathrm{d} y, \quad t \in J, \tag{41}
\end{equation*}
$$

hence

$$
\begin{equation*}
v(t)+\|u(t)\|_{E_{0}} \leqslant v^{0}+\left\|u^{0}\right\|_{E_{0}}+\lambda t, \quad t \in J . \tag{42}
\end{equation*}
$$

Suppose now that $t^{+}<\infty$. Then (42) entails that

$$
\dot{v}(t) \leqslant \lambda+g(u(t)) \leqslant \lambda+\|\beta\|_{\infty}\|u(t)\|_{E_{0}} \leqslant c\left(t^{+}\right), \quad t \in J,
$$

and

$$
\dot{v}(t) \geqslant-\gamma v(t)-\tau|u(t)|_{1} v(t) \geqslant-c\left(t^{+}\right), \quad t \in J .
$$

Therefore

$$
\begin{equation*}
\|v\|_{C^{1}(J)} \leqslant c\left(t^{+}\right) \tag{43}
\end{equation*}
$$

and

$$
\begin{equation*}
v(t) \geqslant e^{-\left(\gamma+\tau|u(t)|_{1}\right) t} v^{0} \geqslant e^{-\left(\gamma+\tau c\left(t^{+}\right)\right) t^{+}} v^{0}>0, \quad t \in J . \tag{44}
\end{equation*}
$$

Taking (26) into account, we derive from (43), (44) that the evolution system $U_{v}(t, s)$ satisfies

$$
\left\|U_{v}(t, s)\right\|_{\mathcal{L}\left(E_{1}\right)} \leqslant c\left(t^{+}\right), \quad 0 \leqslant s \leqslant t<t^{+}
$$

But then

$$
\begin{equation*}
\|u(t)\|_{E_{1}}=\left\|U_{v}(t, 0) u^{0}\right\|_{E_{1}} \leqslant c\left(t^{+}\right)\left\|u^{0}\right\|_{E_{1}}, \quad t \in J \tag{45}
\end{equation*}
$$

thus (40) cannot be true in view of (43)-(45). This contradiction proves that the solution $(v, u)$ exists for all times, hence the assertion follows.

If $(v, u)$ denotes the solution to (1)-(4) provided by Theorem 3.1, the next proposition shows that $u$ propagates with finite speed. The proof is adapted from the proof of [6, Theorem 2.6].

Proposition 3.2. Suppose (10)-(13) hold. For $v^{0}>0$ and $u^{0} \in E_{1}^{+}$let ( $v, u$ ) denotes the unique global classical solution to (1)-(4). If $\operatorname{supp} u^{0} \subset\left[y_{0}, S_{0}\right]$ for some $S_{0}>y_{0}$, then $\operatorname{supp} u(t) \subset$ [ $\left.y_{0}, S(t)\right], t \geqslant 0$, where

$$
S(t):=S_{0}+\tau \int_{0}^{t} v(s) \mathrm{d} s, \quad t \geqslant 0 .
$$

Proof. Define $P \in C^{1}\left(\mathbb{R}^{+}, L_{1}(Y)\right)$ by

$$
P(t, y):=\int_{y}^{\infty} u\left(t, y^{\prime}\right) \mathrm{d} y^{\prime}, \quad y \in Y, t \geqslant 0 .
$$

Then, since

$$
\frac{\mathrm{d}}{\mathrm{~d} t} P(t, y)=\int_{y}^{\infty} \dot{u}\left(t, y^{\prime}\right) \mathrm{d} y^{\prime}=\tau v(t) u(t, y)+\int_{y}^{\infty} L[u(t)]\left(y^{\prime}\right) \mathrm{d} y^{\prime},
$$

we derive from (2) and (14),

$$
\begin{aligned}
\frac{\mathrm{d}}{\mathrm{~d} t} \int_{S(t)}^{\infty} P(t, y) \mathrm{d} y & =\int_{S(t)}^{\infty} \frac{\mathrm{d}}{\mathrm{~d} t} P(t, y) \mathrm{d} y-S^{\prime}(t) P(t, S(t)) \\
& =\int_{S(t)}^{\infty} \int_{y}^{\infty} L[u(t)]\left(y^{\prime}\right) \mathrm{d} y^{\prime} \mathrm{d} y \\
& \leqslant 2 \int_{S(t)}^{\infty} \int_{y}^{\infty} \int_{y^{\prime}}^{\infty} \beta\left(y^{\prime \prime}\right) \kappa\left(y^{\prime}, y^{\prime \prime}\right) u\left(t, y^{\prime \prime}\right) \mathrm{d} y^{\prime \prime} \mathrm{d} y^{\prime} \mathrm{d} y \\
& =2 \int_{S(t)}^{\infty} \int_{y}^{\infty} \beta\left(y^{\prime \prime}\right) u\left(t, y^{\prime \prime}\right) \int_{y}^{y^{\prime \prime}} \kappa\left(y^{\prime}, y^{\prime \prime}\right) \mathrm{d} y^{\prime} \mathrm{d} y^{\prime \prime} \mathrm{d} y \\
& \leqslant 2\|\beta\|_{\infty} \int_{S(t)}^{\infty} P(t, y) \mathrm{d} y
\end{aligned}
$$

which implies

$$
\int_{S(t)}^{\infty} P(t, y) \mathrm{d} y \leqslant e^{2\|\beta\|_{\infty} t} \int_{S_{0}}^{\infty} \int_{y}^{\infty} u^{0}\left(y^{\prime}\right) \mathrm{d} y^{\prime} \mathrm{d} y=0, \quad t \geqslant 0 .
$$

Hence $u(t, y)=0$ for $y \in(S(t), \infty)$ and $t \geqslant 0$.
Remark 3.3. Note that if $\mu(y) \geqslant \underline{\mu}>0$ for a.e. $y \in Y$ and $\gamma>0$, then (41) entails

$$
\begin{equation*}
v(t)+\int_{y_{0}}^{\infty} y u(t, y) \mathrm{d} y \leqslant \frac{\lambda}{v}+e^{-v t}\left(v^{0}+\left\|u^{0}\right\|_{E_{0}}-\frac{\lambda}{v}\right), \quad t \geqslant 0, \tag{46}
\end{equation*}
$$

where $\nu:=\min \{\underline{\mu}, \gamma\}>0$. In particular,

$$
\begin{equation*}
\int_{0}^{t} v(s) \mathrm{d} s \leqslant \frac{\lambda t}{v}+\frac{1}{v}\left(1-e^{-v t}\right)\left(v^{0}+\left\|u^{0}\right\|_{E_{0}}-\frac{\lambda}{v}\right), \quad t \geqslant 0 . \tag{47}
\end{equation*}
$$

## 4. Weak solutions

The aim of this section is to relax condition (10) and to prove existence of weak solutions for unbounded kernels $\mu$ and $\beta$. More precisely, instead of (10) we assume in the following that

$$
\left\{\begin{array}{l}
\text { there exist } \alpha \geqslant 1 \text { and } \varrho \in L_{\infty}^{+}(Y) \text { such that }  \tag{48}\\
\varrho(y) \rightarrow 0 \text { as } y \rightarrow \infty \text { and } \mu(y)+\beta(y) \leqslant \varrho(y) y^{\alpha}, \text { a.e. } y \in Y .
\end{array}\right.
$$

In addition, we require that

$$
\left\{\begin{array}{l}
\text { for each } \varepsilon>0 \text { there exists } \delta>0 \text { such that }  \tag{49}\\
\sup _{|\mathcal{E}| \leqslant \delta} \frac{\beta(y)}{y^{\alpha}} \int_{y_{0}}^{y} \chi \mathcal{E}\left(y^{\prime}\right) \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \leqslant \varepsilon, \text { a.e. } y \in Y,
\end{array}\right.
$$

the supremum being taken over all measurable subsets $\mathcal{E}$ in $Y$ with measure $|\mathcal{E}| \leqslant \delta$. Observe that if $\kappa$ is subject to the self-similar form (15), (16), then

$$
\lim _{|\mathcal{E}| \rightarrow 0} \operatorname{ess-sup} \int_{y>y_{0}}^{y} \int_{y_{0}}^{y} \mathbf{1}_{\mathcal{E}}\left(y^{\prime}\right) \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}=\lim _{|\mathcal{E}| \rightarrow 0} \operatorname{ess}-\mathrm{sup}_{y>y_{0}} \int_{y_{0} / y}^{1} \mathbf{1}_{\frac{1}{y}} \mathcal{E}\left(y^{\prime}\right) \kappa_{0}\left(y^{\prime}\right) \mathrm{d} y^{\prime}=0
$$

due to $y_{0}>0$ and the integrability of $\kappa_{0}$, so (49) automatically holds by (48).
In the following we denote by $L_{1, \mathrm{w}}(Y)$ the space $L_{1}(Y)$ equipped with its weak topology.
Definition 4.1. Given $v^{0}>0$ and $u^{0} \in L_{1}^{+}(Y, y \mathrm{~d} y)$, we call $(v, u)$ a (global) weak solution to (1)-(4) if
(i) $g(u) \in C\left(\mathbb{R}^{+}\right)$,
(ii) $v \in C^{1}\left(\mathbb{R}^{+}\right)$is a non-negative solution to (1),
(iii) $u \in C\left(\mathbb{R}^{+}, L_{1, \mathrm{w}}(Y)\right) \cap L_{\infty, \text { loc }}\left(\mathbb{R}^{+}, L_{1}^{+}(Y, y \mathrm{~d} y)\right)$,
(iv) for all $t>0$ and $\varphi \in W_{\infty}^{1}(Y)$ it holds that $L[u] \in L_{1}((0, t) \times Y)$ and

$$
\begin{aligned}
& \int_{y_{0}}^{\infty} \varphi(y) u(t, y) \mathrm{d} y-\tau \int_{0}^{t} v(s) \int_{y_{0}}^{\infty} \varphi^{\prime}(y) u(s, y) \mathrm{d} y \mathrm{~d} s \\
& =\int_{y_{0}}^{\infty} \varphi(y) u^{0}(y) \mathrm{d} y+\int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y) L[u(s)](y) \mathrm{d} y \mathrm{~d} s .
\end{aligned}
$$

We first need the following auxiliary result.
Lemma 4.2. Suppose that $h_{n}$ and $h$ are measurable functions on $Y$ such that $h_{n} \rightarrow h$ a.e. and let $u_{n} \rightarrow u$ in $L_{1, \mathrm{w}}^{+}(Y)$.
(i) If $\left\|h_{n}\right\|_{\infty} \leqslant c$, then $h_{n} u_{n} \rightarrow h u$ in $L_{1, \mathrm{w}}(Y)$.
(ii) If $\varrho$ and $\alpha$ are as in (48) and if $\left|h_{n}(y)\right| \leqslant \varrho(y) y^{\alpha}$ for a.e. $y \in Y$ and

$$
\int_{y_{0}}^{\infty} y^{\alpha} u_{n}(y) \mathrm{d} y \leqslant c, \quad n \in \mathbb{N}
$$

then $h_{n} u_{n} \rightarrow h u$ in $L_{1, w}(Y)$.

Proof. In case that $Y$ is a finite interval, a proof of (i) is implicitly contained in [11, Lemma 4.1] (a detailed proof can also be found in [13, Appendix]). The case of unbounded $Y$ is a slight modification thereof. Statement (ii) can be shown along the lines of [7, Appendix A, Corollary 4.1]. Nevertheless, for the reader's convenience, we include here a proof. First note that the assumptions imply $|h(y)| \leqslant \varrho(y) y^{\alpha}$, a.e. $y \in Y$, and

$$
\int_{y_{0}}^{\infty} y^{\alpha} u(y) \mathrm{d} y \leqslant c
$$

Putting $\bar{u}_{n}(y):=\varrho(y) y^{\alpha} u_{n}(y)$ and $\bar{u}(y):=\varrho(y) y^{\alpha} u(y)$, we obtain for $\varphi \in L_{\infty}(Y)$ and $R>y_{0}$,

$$
\begin{aligned}
\left|\int_{y_{0}}^{\infty} \varphi(y)\left(\bar{u}_{n}(y)-\bar{u}(y)\right) \mathrm{d} y\right| \leqslant & \left|\int_{y_{0}}^{R} \varphi(y) \varrho(y) y^{\alpha}\left(u_{n}(y)-u(y)\right) \mathrm{d} y\right| \\
& +2 c\|\varphi\|_{\infty}\|\varrho\|_{L_{\infty}(R, \infty)} .
\end{aligned}
$$

Taking first the limsup as $n \rightarrow \infty$ on both sides and letting then $R \rightarrow \infty$, we conclude from (48) that $\bar{u}_{n} \rightarrow \bar{u}$ in $L_{1, \mathrm{w}}(Y)$. Therefore, it follows from (i) that the right-hand side of the estimate

$$
\begin{aligned}
& \left|\int_{y_{0}}^{\infty} \varphi(y)\left(h_{n}(y) u_{n}(y)-h(y) u(y)\right) \mathrm{d} y\right| \\
& \quad \leqslant\left|\int_{y_{0}}^{\infty} \varphi(y)\left(\varrho(y) y^{\alpha}\right)^{-1}\left(h_{n}(y)-h(y)\right) \bar{u}_{n}(y) \mathrm{d} y\right| \\
& \quad+\left|\int_{y_{0}}^{\infty} \varphi(y)\left(\varrho(y) y^{\alpha}\right)^{-1} h(y)\left(\bar{u}_{n}(y)-\bar{u}(y)\right) \mathrm{d} y\right|
\end{aligned}
$$

converges to 0 , leading to the assertion.
Now we are in a position to relax the boundedness assumptions on $\mu$ and $\beta$ and also the assumption on $u^{0}$ can be weakened.

Theorem 4.3. Suppose that (11)-(13) and (48), (49) hold. Then, given any $v^{0}>0$ and $u^{0} \in$ $L_{1}^{+}\left(Y, y^{\alpha} \mathrm{d} y\right)$, problem (1)-(4) admits at least one global weak solution $(v, u)$. In addition, $u$ belongs to $L_{\infty, \text { loc }}\left(\mathbb{R}^{+}, L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)\right)$.

Proof. (i) Let $u_{n}^{0} \in \mathcal{D}^{+}(Y)$ be such that $u_{n}^{0} \rightarrow u^{0}$ in $L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)$. We define $\mu_{n}:=\min \{\mu, n\}$ and $\beta_{n}:=\min \{\beta, n\}$. Observe that $\mu_{n}, \beta_{n}$ also satisfy (48) and (49). Then Theorem 3.1 guarantees the existence of

$$
\left(v_{n}, u_{n}\right) \in C\left(\mathbb{R}^{+}, \mathbb{R}^{+} \times E_{1}^{+}\right) \cap C^{1}\left(\mathbb{R}^{+}, \mathbb{R} \times E_{0}\right)
$$

satisfying

$$
\begin{equation*}
\dot{v}_{n}=\lambda-\gamma v_{n}-\tau v_{n}\left|u_{n}\right|_{1}+g_{n}\left(u_{n}\right), \quad t>0, \quad v_{n}(0)=v^{0} \tag{50}
\end{equation*}
$$

and

$$
\begin{equation*}
\partial_{t} u_{n}+\tau v_{n}(t) \partial_{y} u_{n}=L_{n}\left[u_{n}\right], \quad t>0, \quad u_{n}(0)=u_{n}^{0} \tag{51}
\end{equation*}
$$

where

$$
g_{n}(u):=2 \int_{y_{0}}^{\infty} u(y) \beta_{n}(y) \int_{0}^{y_{0}} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y
$$

and

$$
L_{n}[u](y):=-\left(\mu_{n}(y)+\beta_{n}(y)\right) u(y)+2 \int_{y}^{\infty} \beta_{n}\left(y^{\prime}\right) \kappa\left(y, y^{\prime}\right) u\left(y^{\prime}\right) \mathrm{d} y^{\prime}
$$

Let $T>0$ be arbitrary. According to (42) there exists $c_{0}(T)>0$ independent of $n \geqslant 1$ such that

$$
\begin{equation*}
v_{n}(t)+\left\|u_{n}(t)\right\|_{E_{0}} \leqslant c_{0}(T), \quad t \in J_{T}, n \geqslant 1 \tag{52}
\end{equation*}
$$

Moreover, we claim that

$$
\begin{equation*}
\left\|u_{n}(t)\right\|_{L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)} \leqslant c_{0}(T), \quad t \in J_{T}, n \geqslant 1 . \tag{53}
\end{equation*}
$$

For, recall that $u_{n}(t)$ has compact support due to Proposition 3.2. Hence, we may test (51) by $y^{\alpha}$ and obtain

$$
\begin{aligned}
& \frac{\mathrm{d}}{\mathrm{~d} t} \int_{y_{0}}^{\infty} y^{\alpha} u_{n}(t, y) \mathrm{d} y= \alpha \tau v_{n}(t) \int_{y_{0}}^{\infty} y^{\alpha-1} u_{n}(t, y) \mathrm{d} y \\
&-\int_{y_{0}}^{\infty} y^{\alpha}\left(\mu_{n}(y)+\beta_{n}(y)\right) u_{n}(t, y) \mathrm{d} y \\
&+2 \int_{y_{0}}^{\infty} u_{n}(t, y) \beta_{n}(y) \int_{y_{0}}^{y}\left(y^{\prime}\right)^{\alpha} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \\
& \leqslant \alpha \tau v_{n}(t) \int_{y_{0}}^{\infty} y^{\alpha-1} u_{n}(t, y) \mathrm{d} y
\end{aligned}
$$

for $t \geqslant 0$, since (12) ensures

$$
2 \int_{y_{0}}^{y}\left(y^{\prime}\right)^{\alpha} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \leqslant y^{\alpha}, \quad \text { a.e. } y>y_{0} .
$$

Therefore, Gronwall's inequality and estimate (52) yield (53). In particular, combining (53), (48) and (14), we deduce

$$
g_{n}\left(u_{n}(t)\right) \leqslant 2 y_{0}\|\varrho\|_{\infty}\left\|u_{n}(t)\right\|_{L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)} \leqslant c(T), \quad t \in J_{T}, n \geqslant 1 .
$$

(ii) It follows from (1) and the estimate on $g_{n}\left(u_{n}(t)\right)$ that

$$
\left|v_{n}(t)-v_{n}(s)\right| \leqslant c(T)|t-s|, \quad t, s \in J_{T}, n \geqslant 1,
$$

where $c(T)>0$ is independent of $n \geqslant 1$. Taking (52) into account, the Arzelà-Ascoli theorem warrants that the sequence $\left(v_{n}\right)$ is relatively compact in $C\left(J_{T}\right)$.
(iii) We show that $\left(u_{n}\right)$ is relatively sequentially compact in $C\left(J_{T}, L_{1, \mathrm{w}}(Y)\right)$. According to a variant of the Arzelà-Ascoli theorem (see [12, Theorem 1.3.2]), we merely have to check that the set $\left\{u_{n}(t) ; n \geqslant 1\right\}$ is relatively compact in $L_{1, \mathrm{w}}(Y)$ for every $t \in J_{T}$ and that the set $\left\{u_{n} ; n \geqslant 1\right\}$ is equicontinuous in $L_{1, \mathrm{w}}(Y)$ at every $t \in J_{T}$. First observe that (52) entails

$$
\begin{equation*}
\lim _{R \rightarrow \infty} \sup _{\substack{n \geqslant 1 \\ t \in J_{T}}} \int_{R}^{\infty} u_{n}(t, y) \mathrm{d} y=0 . \tag{54}
\end{equation*}
$$

Let $U_{v_{n}}(t, s)$ denote the evolution system in $L_{1}(Y)$ corresponding to the operator $A_{v_{n}}(t):=$ $\tau v_{n}(t) \partial_{y}$. Then

$$
u_{n}(t)=U_{v_{n}}(t, 0) u_{n}^{0}+\int_{0}^{t} U_{v_{n}}(t, s) L_{n}\left[u_{n}(s)\right] \mathrm{d} s, \quad t \in J_{T}
$$

Consequently, given $\delta>0$, Lemma 2.4 and the positivity of $u_{n}(t)$ imply that

$$
\begin{aligned}
\sup _{|\mathcal{E}| \leqslant \delta} \int_{\mathcal{E}} u_{n}(t, y) \mathrm{d} y \leqslant & \sup _{|\mathcal{E}| \leqslant \delta} \int_{\mathcal{E}} u_{n}^{0}(y) \mathrm{d} y \\
& +2 \int_{0}^{t} \sup _{|\mathcal{E}| \leqslant \delta} \int_{y_{0}}^{\infty} u_{n}(s, y) \beta_{n}(y) \int_{y_{0}}^{y} \chi \mathcal{E}\left(y^{\prime}\right) \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \mathrm{~d} s .
\end{aligned}
$$

Since $u_{n}^{0} \rightarrow u^{0}$ in $L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)$ and in view of (49) and (53), we conclude that

$$
\begin{equation*}
\lim _{|\mathcal{E}| \rightarrow 0} \sup _{\substack{n \geqslant 1 \\ t \in J_{T}}} \int_{\mathcal{E}} u_{n}(t, y) \mathrm{d} y=0 . \tag{55}
\end{equation*}
$$

From (52), (54), (55) and the Dunford-Pettis theorem (cf. [2, Theorem 4.21.2]) we hence derive that $\left\{u_{n}(t) ; t \in J_{T}, n \geqslant 1\right\}$ is relatively compact in $L_{1, \mathrm{w}}(Y)$.

Now let $\varphi \in \mathcal{D}(Y)$ be arbitrary. Testing (51) by $\varphi$, we infer

$$
\begin{aligned}
& \left|\int_{y_{0}}^{\infty} \varphi(y)\left[u_{n}(t, y)-u_{n}(s, y)\right] \mathrm{d} y\right| \\
& \leqslant \tau \int_{s}^{t} v_{n}(\sigma) \int_{y_{0}}^{\infty}\left|\varphi^{\prime}(y)\right| u_{n}(\sigma, y) \mathrm{d} y \mathrm{~d} \sigma \\
& \quad+\int_{s}^{t} \int_{y_{0}}^{\infty}|\varphi(y)|\left(\mu_{n}(y)+\beta_{n}(y)\right) u_{n}(\sigma, y) \mathrm{d} y \mathrm{~d} \sigma \\
& \quad+2 \int_{s}^{t} \int_{y_{0}}^{\infty} u_{n}(\sigma, y) \beta_{n}(y) \int_{y_{0}}^{y}\left|\varphi\left(y^{\prime}\right)\right| \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \mathrm{~d} \sigma
\end{aligned}
$$

for $0 \leqslant s \leqslant t \leqslant T$, whence, from (14), (48), (52) and (53),

$$
\begin{equation*}
\left|\int_{y_{0}}^{\infty} \varphi(y)\left[u_{n}(t, y)-u_{n}(s, y)\right] \mathrm{d} y\right| \leqslant c(T, \varphi)|t-s|, \quad t, s \in J_{T} . \tag{56}
\end{equation*}
$$

For $\varphi \in L_{\infty}(Y)$ let $\varphi_{j} \in \mathcal{D}(Y)$ be such that $\varphi_{j} \rightarrow \varphi$ a.e. and $\left\|\varphi_{j}\right\|_{\infty} \leqslant\|\varphi\|_{\infty}$ (see [1, p. 131f]). Given $\varepsilon>0$ it follows from (54), from the fact that $\left\{u_{n}(t) ; t \in J_{T}, n \geqslant 1\right\}$ is relatively compact in $L_{1, \mathrm{w}}(Y)$, and from Egorov's theorem that there are $R>y_{0}$, a measurable subset $\mathcal{E}$ of $\left(y_{0}, R\right)$ and $j \in \mathbb{N}$ such that

$$
\int_{R}^{\infty} u_{n}(t, y) \mathrm{d} y+\int_{\mathcal{E}} u_{n}(t, y) \mathrm{d} y \leqslant \frac{\varepsilon}{12\|\varphi\|_{\infty}}, \quad t \in J_{T}, n \geqslant 1,
$$

and

$$
\left\|\varphi-\varphi_{j}\right\|_{L_{\infty}\left(\left(y_{0}, R\right) \backslash \mathcal{E}\right)} \leqslant \frac{\varepsilon}{6 c_{0}(T)}
$$

where $c_{0}(T)>0$ stems from (52). Therefore, (56) yields

$$
\begin{aligned}
\left|\int_{y_{0}}^{\infty} \varphi(y)\left[u_{n}(t, y)-u_{n}(s, y)\right] \mathrm{d} y\right| \leqslant & \left\|\varphi-\varphi_{j}\right\|_{L_{\infty}\left(\left(y_{0}, R\right) \backslash \mathcal{E}\right)}\left(\left|u_{n}(t)\right|_{1}+\left|u_{n}(s)\right|_{1}\right) \\
& +\left(\|\varphi\|_{\infty}+\left\|\varphi_{j}\right\|_{\infty}\right) \int_{\mathcal{E}}\left(u_{n}(t, y)+u_{n}(s, y)\right) \mathrm{d} y \\
& +\left(\|\varphi\|_{\infty}+\left\|\varphi_{j}\right\|_{\infty}\right) \int_{R}^{\infty}\left(u_{n}(t, y)+u_{n}(s, y)\right) \mathrm{d} y \\
& +c\left(T, \varphi_{j}\right)|t-s| \\
\leqslant & \varepsilon+c\left(T, \varphi_{j}\right)|t-s|
\end{aligned}
$$

for $t, s \in J_{T}$ and $n \geqslant 1$. We conclude

$$
\lim _{s \rightarrow t} \sup _{n \geqslant 1}\left|\int_{y_{0}}^{\infty} \varphi(y)\left[u_{n}(t, y)-u_{n}(s, y)\right] \mathrm{d} y\right|=0
$$

hence $\left\{u_{n} ; n \geqslant 1\right\}$ is equicontinuous in $L_{1, \mathrm{w}}(Y)$ at every $t \in J_{T}$.
(iv) Since now $\left(v_{n}, u_{n}\right)$ is relatively compact in $C\left(J_{T}, \mathbb{R} \times L_{1, \mathrm{w}}(Y)\right)$ for each $T>0$, we may choose a subsequence (again denoted by $\left.\left(\left(v_{n}, u_{n}\right)\right)_{n \in \mathbb{N}}\right)$ and a function $(v, u) \in C\left(\mathbb{R}^{+}\right.$, $\left.\mathbb{R} \times L_{1, \mathrm{w}}(Y)\right)$ such that

$$
\begin{equation*}
\left(v_{n}, u_{n}\right) \rightarrow(v, u) \quad \text { in } C\left(J_{T}, \mathbb{R} \times L_{1, \mathrm{w}}(Y)\right) \tag{57}
\end{equation*}
$$

for each $T>0$.
(v) We then claim that $(v, u)$ is a weak solution to (1)-(4). Evidently, it holds that $(v(t), u(t)) \in \mathbb{R}^{+} \times L_{1}^{+}(Y)$ for $t>0$ since $\left(v_{n}(t), u_{n}(t)\right) \in \mathbb{R}^{+} \times L_{1}^{+}(Y)$. We fix again $T>0$. Then (57) and (53) imply

$$
\begin{equation*}
\|u(t)\|_{L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)} \leqslant c_{0}(T), \quad t \in J_{T} \tag{58}
\end{equation*}
$$

in particular, we have $u \in L_{\infty, \operatorname{loc}}\left(\mathbb{R}^{+} L_{1}\left(Y, y^{\alpha} \mathrm{d} y\right)\right)$. Let $\varphi \in W_{\infty}^{1}(Y)$ be arbitrary. Clearly, (57) yields

$$
\begin{equation*}
\lim _{n \rightarrow \infty} \int_{y_{0}}^{\infty} \varphi(y) u_{n}(t, y) \mathrm{d} y=\int_{y_{0}}^{\infty} \varphi(y) u(t, y) \mathrm{d} y, \quad t \in J_{T} . \tag{59}
\end{equation*}
$$

Moreover, writing

$$
\begin{aligned}
& \left|\int_{0}^{t} v(s) \int_{y_{0}}^{\infty} \varphi^{\prime}(y) u(s, y) \mathrm{d} y \mathrm{~d} s-\int_{0}^{t} v_{n}(s) \int_{y_{0}}^{\infty} \varphi^{\prime}(y) u_{n}(s, y) \mathrm{d} y \mathrm{~d} s\right| \\
& \leqslant \int_{0}^{t}\left|v(s)-v_{n}(s)\right| \int_{y_{0}}^{\infty}\left|\varphi^{\prime}(y)\right| u(s, y) \mathrm{d} y \mathrm{~d} s \\
& \quad+\int_{0}^{t} v_{n}(s)\left|\int_{y_{0}}^{\infty} \varphi^{\prime}(y)\left[u(s, y)-u_{n}(s, y)\right] \mathrm{d} y\right| \mathrm{d} s
\end{aligned}
$$

for $t \in J_{T}$, we infer from (57), (52) and Lebesgue's theorem that, for $t \in J_{T}$,

$$
\begin{equation*}
\lim _{n \rightarrow \infty} \int_{0}^{t} v_{n}(s) \int_{y_{0}}^{\infty} \varphi^{\prime}(y) u_{n}(s, y) \mathrm{d} y \mathrm{~d} s=\int_{0}^{t} v(s) \int_{y_{0}}^{\infty} \varphi^{\prime}(y) u(s, y) \mathrm{d} y \mathrm{~d} s \tag{60}
\end{equation*}
$$

In addition, since $\mu_{n}(y)+\beta_{n}(y) \leqslant \varrho(y) y^{\alpha}$ for a.e. $y \in Y$, we conclude from Lemma 4.2(ii), (53), (57) and Lebesgue's theorem that

$$
\begin{aligned}
& \lim _{n \rightarrow \infty} \int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y)\left(\mu_{n}(y)+\beta_{n}(y)\right) u_{n}(s, y) \mathrm{d} y \mathrm{~d} s \\
& =\int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y)(\mu(y)+\beta(y)) u(s, y) \mathrm{d} y \mathrm{~d} s
\end{aligned}
$$

as well as

$$
\begin{aligned}
& \lim _{n \rightarrow \infty} \int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y) \int_{y}^{\infty} u_{n}\left(s, y^{\prime}\right) \beta_{n}\left(y^{\prime}\right) \kappa\left(y, y^{\prime}\right) \mathrm{d} y^{\prime} \mathrm{d} y \mathrm{~d} s \\
& =\int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y) \int_{y}^{\infty} u\left(s, y^{\prime}\right) \beta\left(y^{\prime}\right) \kappa\left(y, y^{\prime}\right) \mathrm{d} y^{\prime} \mathrm{d} y \mathrm{~d} s
\end{aligned}
$$

where we use Fubini's theorem for the second limit. Therefore,

$$
\begin{equation*}
\lim _{n \rightarrow \infty} \int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y) L_{n}\left[u_{n}(s)\right] \mathrm{d} y \mathrm{~d} s=\int_{0}^{t} \int_{y_{0}}^{\infty} \varphi(y) L[u(s)] \mathrm{d} y \mathrm{~d} s \tag{61}
\end{equation*}
$$

Now, since $\left(v_{n}, u_{n}\right)$ is a weak solution to (1)-(4), we derive from (59)-(61) that $u$ indeed satisfies part (iv) of Definition 4.1. Next, it follows from Lemma 4.2(ii), similarly as above, that

$$
\lim _{n \rightarrow \infty} g_{n}\left(u_{n}(t)\right)=g(u(t)), \quad t \in J_{T}
$$

and also

$$
\lim _{n \rightarrow \infty} \int_{0}^{t}\left|u_{n}(s)\right|_{1} \mathrm{~d} s=\int_{0}^{t}|u(s)|_{1} \mathrm{~d} s, \quad t \in J_{T}
$$

Consequently, (50) yields

$$
v(t)=e^{-\gamma t-\tau \int_{0}^{t}|u(\sigma)|_{1} \mathrm{~d} \sigma} v^{0}+\int_{0}^{t} e^{-\gamma(t-s)-\tau \int_{s}^{t}|u(\sigma)|_{1} \mathrm{~d} \sigma}(\lambda+g(u(s))) \mathrm{d} s
$$

for $t \in J_{T}$. But since $u \in C\left(\mathbb{R}^{+}, L_{1, \mathrm{w}}(Y)\right)$, Lemma 4.2(ii) and (58) warrant that $g(u) \in C\left(J_{T}\right)$. In addition, $|u|_{1} \in C\left(J_{T}\right)$, so we deduce that $v \in C^{1}\left(J_{T}\right)$ solves (1). This proves the theorem.

Also the weak solution propagates with finite speed as shown in the next corollary.
Corollary 4.4. Suppose (11)-(13), (48), (49). If $v^{0}>0$ and if $u^{0} \in L_{1}^{+}\left(Y, y^{\alpha} \mathrm{d} y\right)$ is such that $\operatorname{supp} u^{0} \subset\left[y_{0}, S_{0}\right]$, then the weak solution $(v, u)$ provided by Theorem 4.3 satisfies $\operatorname{supp} u(t) \subset$ [ $\left.y_{0}, S(t)\right]$ for $t \geqslant 0$, where

$$
S(t):=S_{0}+\tau \int_{0}^{t} v(s) \mathrm{d} s, \quad t \geqslant 0 .
$$

Proof. We may choose the sequence $\left(u_{n}^{0}\right) \subset \mathcal{D}^{+}(Y)$ in the proof of Theorem 4.3 such that $\operatorname{supp} u_{n}^{0} \subset\left(y_{0}, S_{0}\right)$. Then Proposition 3.2 ensures that the approximating sequence $\left(\left(v_{n}, u_{n}\right)\right)_{n \in \mathbb{N}}$ given in (50), (51) satisfies $\operatorname{supp} u_{n}(t) \subset\left[y_{0}, S_{n}(t)\right]$ for $t \geqslant 0$, where

$$
S_{n}(t):=S_{0}+\tau \int_{0}^{t} v_{n}(s) \mathrm{d} s, \quad t \geqslant 0, n \geqslant 1
$$

Evidently, $\lim _{n \rightarrow \infty} S_{n}(t)=S(t)$ and

$$
\int_{S(t)}^{\infty} u(t, y) \mathrm{d} y=\lim _{n \rightarrow \infty} \int_{S_{n}(t)}^{\infty} u_{n}(t, y) \mathrm{d} y=0
$$

by (57) and Lemma 4.2(i), thus $\operatorname{supp} u(t) \subset\left[y_{0}, S(t)\right]$ for $t \geqslant 0$.

Remark 4.5. In addition to (11)-(13), (48), (49) suppose that $\mu(y) \geqslant \mu>0$ for a.e. $y \in Y$ and that $\gamma>0$. Then the weak solution $(v, u)$ also satisfies the estimates (46) and (47). Indeed, (46) follows immediately from the corresponding estimate for $\left(v_{n}, u_{n}\right)$ and (57).

## 5. Stability of the disease-free steady state

This section is devoted to the investigation of stability properties of the disease-free steady state $(v, u)=(\lambda / \gamma, 0)$ of (1), (2).

In the sequel, we always assume that (11)-(13) are satisfied with $\gamma>0$ and that either

$$
\left\{\begin{array}{l}
(10) \text { holds },  \tag{62}\\
v^{0}>0, \quad u^{0} \in E_{1}^{+},
\end{array}\right.
$$

or

$$
\left\{\begin{array}{l}
\text { (48), (49) hold, }  \tag{63}\\
v^{0}>0, \quad u^{0} \in L_{1}^{+}\left(Y, y^{\alpha} \mathrm{d} y\right) .
\end{array}\right.
$$

Then we denote by ( $v, u$ ) either the classical solution provided by Theorem 3.1 if (62) holds, or the weak solution provided by Theorem 4.3 if (63) holds.

We assume that

$$
d_{0}:=\underset{y \in Y}{\operatorname{ess}-\sup } \frac{\beta(y)}{y \mu(y)} \in(0, \infty)
$$

and introduce $\varepsilon_{k}, \delta_{k}$ such that

$$
0 \leqslant \delta_{k} \leqslant \beta(y) \int_{0}^{y_{0}}\left(y^{\prime}\right)^{k} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \leqslant \varepsilon_{k}, \quad \text { a.e. } y \in Y
$$

for $k=0,1$, assuming at least $\varepsilon_{1}$ to be finite. In the following we suppose that

$$
\begin{equation*}
\underline{\mu}:=\underset{y \in Y}{\operatorname{ess-inf}} \mu(y)>0 \tag{64}
\end{equation*}
$$

and

$$
\begin{equation*}
\frac{1}{2 d_{0}}\left(\underline{\mu}+2 \delta_{0}\right)>\frac{\tau \lambda}{2 \gamma}+\varepsilon_{1}-2 \delta_{1}+\frac{2 d_{0} \delta_{1}\left(\varepsilon_{1}-\delta_{1}\right)}{\underline{\mu}+2 \delta_{0}} \tag{65}
\end{equation*}
$$

Given the assumptions above, we can construct a Lyapunov function as follows.

Lemma 5.1. Suppose (62) or (63) and that (64) and (65) are satisfied. Then there are constants $a, b, p, q>0$ such that for

$$
F(v, u):=\left(v-\frac{\lambda}{\gamma}\right)^{2}+a \int_{y_{0}}^{\infty} y u(y) \mathrm{d} y+b \int_{y_{0}}^{\infty} u(y) \mathrm{d} y
$$

there holds

$$
F(v, u)(t)+p \int_{0}^{t} \int_{y_{0}}^{\infty} u(s, y) \mathrm{d} y \mathrm{~d} s+q \int_{0}^{t} \int_{y_{0}}^{\infty} y u(s, y) \mathrm{d} y \mathrm{~d} s \leqslant F\left(v^{0}, u^{0}\right)
$$

for each $t \geqslant 0$, where $(v, u)$ is either the classical solution or the weak solution constructed in Theorems 3.1 or 4.3, respectively.

Proof. Defining

$$
A:=\frac{\tau}{2}\left(\underline{\mu}+2 \delta_{0}\right)>0, \quad B:=2 \delta_{1}-\varepsilon_{1}-\frac{\tau \lambda}{2 \gamma}, \quad C:=4 \delta_{1}\left(\varepsilon_{1}-\delta_{1}\right) \geqslant 0
$$

and $d:=\tau d_{0} / 4$, (65) can be recasted as

$$
\frac{A}{4 d}>-B+\frac{C d}{A} .
$$

Therefore, with

$$
b:=\frac{A}{4 d^{2}}+\frac{C}{A}>\frac{C}{A} \geqslant 0
$$

we have $b d<B+\sqrt{A b-C}$, hence

$$
\begin{equation*}
0<\frac{4}{\tau} b d<a<\frac{4}{\tau}(B+\sqrt{A b-C}) \quad \text { and } \quad \frac{4}{\tau}(B-\sqrt{A b-C})<a \tag{66}
\end{equation*}
$$

for $a:=2 / \tau(\max \{b d, B-\sqrt{A b-C}\}+B+\sqrt{A b-C})$. We set

$$
R:=b\left(\underline{\mu}+2 \delta_{0}\right)+\frac{4 \lambda \delta_{1}}{\gamma}-\frac{\tau \lambda^{2}}{2 \gamma^{2}}-\frac{2 \varepsilon_{1}^{2}}{\tau}-\frac{2 \lambda \varepsilon_{1}}{\gamma}
$$

and notice that $0<A b-C=B^{2}+\tau R / 2$, hence $p:=-\tau a^{2} / 8+B a+R>0$ by (66). Since (66) also warrants that $d_{0}<a / b$, we infer from (64) the existence of $q>0$ such that

$$
\begin{equation*}
\underset{y \in Y}{\operatorname{ess}-s u p} \frac{\beta(y)}{y \mu(y)}+\frac{q}{b} \underset{y \in Y}{\operatorname{ess}-s u p} \frac{1}{\mu(y)}<\frac{a}{b} . \tag{67}
\end{equation*}
$$

Now, in the case of the classical solution one can show directly that

$$
\frac{\mathrm{d}}{\mathrm{~d} t} F(v, u)(t) \leqslant-p|u(t)|_{1}-q \int_{y_{0}}^{\infty} u(t, y) y \mathrm{~d} y, \quad t \geqslant 0
$$

using estimates very close to the subsequent ones. We hence focus on the case of weak solutions. Let $\left(v_{n}, u_{n}\right)$ be the approximations of $(v, u)$ corresponding to the data $\left(v^{0}, u_{n}^{0}, \beta_{n}, \mu_{n}\right)$ as in the proof of Theorem 4.3. Then it follows from (12), (14) and (18) that

$$
\begin{aligned}
\frac{\mathrm{d}}{\mathrm{~d} t} F\left(v_{n}, u_{n}\right)= & -2 \gamma\left(v_{n}-\frac{\lambda}{\gamma}\right)^{2}-2 \tau v_{n}^{2}\left|u_{n}\right|_{1}+\frac{2 \tau \lambda}{\gamma} v_{n}\left|u_{n}\right|_{1} \\
& +4\left(v_{n}-\frac{\lambda}{\gamma}\right) \int_{y_{0}}^{\infty} u_{n}(y) \beta_{n}(y) \int_{0}^{y_{0}} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \\
& +a \tau v_{n}\left|u_{n}\right|_{1}-a \int_{y_{0}}^{\infty} y \mu_{n}(y) u_{n}(y) \mathrm{d} y \\
& -2 a \int_{y_{0}}^{\infty} u_{n}(y) \beta_{n}(y) \int_{0}^{y_{0}} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y
\end{aligned}
$$

$$
\begin{aligned}
& -b \int_{y_{0}}^{\infty} \mu_{n}(y) u_{n}(y) \mathrm{d} y \\
& +b \int_{y_{0}}^{\infty} u_{n}(y) \beta_{n}(y)\left(1-2 \int_{0}^{y_{0}} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}\right) \mathrm{d} y
\end{aligned}
$$

Recalling that $\underline{\mu}>0$ and $\varepsilon_{1}<\infty$, integration of the above equality yields (for $n>\underline{\mu}$ )

$$
\begin{align*}
& F\left(v_{n}, u_{n}\right)(t)+\int_{0}^{t}\left|u_{n}(s)\right|_{1}\left(2 \tau v_{n}(s)^{2}+b \underline{\mu}\right) \mathrm{d} s \\
& \quad+\int_{0}^{t} \int_{y_{0}}^{\infty} u_{n}(s, y) \beta_{n}(y)\left[\left(\frac{4 \lambda}{\gamma}+2 a\right) \int_{0}^{y_{0}} y^{\prime} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}+2 b \int_{0}^{y_{0}} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime}\right] \mathrm{d} y \mathrm{~d} s \\
& \quad+a \int_{0}^{t} \int_{y_{0}}^{\infty} y \mu_{n}(y) u_{n}(s, y) \mathrm{d} y \mathrm{~d} s \\
& \leqslant F\left(v^{0}, u_{n}^{0}\right)+b \int_{0}^{t} \int_{y_{0}}^{\infty} u_{n}(s, y) \beta_{n}(y) \mathrm{d} y \mathrm{~d} s \\
& \quad+\int_{0}^{t}\left|u_{n}(s)\right|_{1} v_{n}(s) \mathrm{d} s\left(\frac{2 \tau \lambda}{\gamma}+a \tau+4 \varepsilon_{1}\right) \tag{68}
\end{align*}
$$

Observe then that (57) ensures

$$
\begin{equation*}
F(v, u)(t) \leqslant \varlimsup_{n \rightarrow \infty} F\left(v_{n}, u_{n}\right)(t), \quad t \geqslant 0 . \tag{69}
\end{equation*}
$$

Next, (57) and Lebesgue's theorem imply

$$
\begin{equation*}
\lim _{n \rightarrow \infty} \int_{0}^{t}\left|u_{n}(s)\right|_{1} v_{n}(s) \mathrm{d} s=\int_{0}^{t}|u(s)|_{1} v(s) \mathrm{d} s, \quad t \geqslant 0 . \tag{70}
\end{equation*}
$$

As in (61) one shows that

$$
\begin{align*}
& \lim _{n \rightarrow \infty} \int_{0}^{t} \int_{y_{0}}^{\infty} u_{n}(s, y) \beta_{n}(y) \int_{0}^{y_{0}}\left(y^{\prime}\right)^{k} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \mathrm{~d} s \\
& =\int_{0}^{t} \int_{y_{0}}^{\infty} u(s, y) \beta(y) \int_{0}^{y_{0}}\left(y^{\prime}\right)^{k} \kappa\left(y^{\prime}, y\right) \mathrm{d} y^{\prime} \mathrm{d} y \mathrm{~d} s \tag{71}
\end{align*}
$$

for $k=0,1$. Owing to Lemma 4.2, (48), (52) and (57), we may apply Lebesgue's theorem to conclude

$$
\lim _{n \rightarrow \infty} \int_{0}^{t} \int_{y_{0}}^{\infty} \chi_{\left(y_{0}, R\right)}(y) u_{n}(s, y) \mu_{n}(y) y \mathrm{~d} y \mathrm{~d} s=\int_{0}^{t} \int_{y_{0}}^{\infty} \chi_{\left(y_{0}, R\right)}(y) u(s, y) \mu(y) y \mathrm{~d} y \mathrm{~d} s
$$

for each $R>y_{0}$, hence

$$
\begin{equation*}
\int_{0}^{t} \int_{y_{0}}^{\infty} u(s, y) \mu(y) y \mathrm{~d} y \mathrm{~d} s \leqslant \varlimsup_{n \rightarrow \infty} \int_{0}^{t} \int_{y_{0}}^{\infty} u_{n}(s, y) \mu_{n}(y) y \mathrm{~d} y \mathrm{~d} s \tag{72}
\end{equation*}
$$

Thus, in view of (69)-(72) we may pass to the limit in (68) to deduce that this inequality is still true if we replace $\left(v_{n}, u_{n}\right)$ by $(v, u)$ and $\left(\beta_{n}, \mu_{n}\right)$ by $(\beta, \mu)$, respectively. Rearranging the terms and using the definition of $\delta_{k}$, we derive

$$
\begin{aligned}
& F(v, u)(t)+\int_{0}^{t}|u(s)|_{1}\left\{2 \tau v(s)^{2}-\left(\frac{2 \tau \lambda}{\gamma}+4 \varepsilon_{1}+a \tau\right) v(s)\right. \\
& \left.\quad+b\left(\underline{\mu}+2 \delta_{0}\right)+\left(\frac{4 \lambda}{\gamma}+2 a\right) \delta_{1}\right\} \mathrm{d} s \\
& \quad+\int_{0}^{t} \int_{y_{0}}^{\infty}(a y \mu(y)-b \beta(y)) u(s, y) \mathrm{d} y \mathrm{~d} s \\
& \leqslant F\left(v^{0}, u^{0}\right)
\end{aligned}
$$

for each $t \geqslant 0$. Minimizing the quadratic function in the curly brackets and observing then that $p>0$ is a lower bound, the assertion follows from (67).

Remark 5.2. In the case of rates subject to (5) it has already been observed in [4] that the function $F$ defined in Lemma 5.1 is a Lyapunov function.

The next theorem shows that the disease-free steady state is asymptotically stable.
Theorem 5.3. Suppose (62) or (63) is satisfied and that (64), (65) hold. Then, given $\varepsilon>0$ there exists $\delta>0$ such that

$$
|v(t)-\lambda / \gamma|+\|u(t)\|_{E_{0}} \leqslant \varepsilon, \quad t \geqslant 0,
$$

whenever

$$
\left|v^{0}-\lambda / \gamma\right|+\left\|u^{0}\right\|_{E_{0}} \leqslant \delta
$$

where $(v, u)$ is either the classical solution or the weak solution constructed in Theorems 3.1 or 4.3, respectively.

Moreover, if $\beta(y) \leqslant B y$ for a.e. $y \in Y$ and some $B>0$, then

$$
(v(t), u(t)) \rightarrow(\lambda / \gamma, 0) \quad \text { in } \mathbb{R} \times L_{1}\left(Y, y^{\sigma} \mathrm{d} y\right) \text { as } t \rightarrow \infty
$$

for each $\sigma<1$ and any initial value $\left(v^{0}, u^{0}\right)$ subject to (62) or (63).
Proof. Defining $F$ as in Lemma 5.1, the first statement readily follows from the fact that $F(v, u)(t) \leqslant F\left(v^{0}, u^{0}\right)$ for $t \geqslant 0$. Next, Lemma 5.1 also ensures that

$$
\begin{equation*}
\|u(t)\|_{L_{1}(Y, y \mathrm{~d} y)} \leqslant \frac{1}{a} F\left(v^{0}, u^{0}\right), \quad t \geqslant 0 . \tag{73}
\end{equation*}
$$

Furthermore, by definition of a weak solution we have

$$
|u(t)|_{1}=\left|u^{0}\right|_{1}+\int_{0}^{t} \int_{y_{0}}^{\infty} L[u(s)](y) \mathrm{d} y \mathrm{~d} s, \quad t \geqslant 0
$$

from which we infer that

$$
\begin{aligned}
\frac{1}{h}\left(|u(t+h)|_{1}-|u(t)|_{1}\right) & =\frac{1}{h} \int_{t}^{t+h} \int_{y_{0}}^{\infty} L[u(s)](y) \mathrm{d} y \mathrm{~d} s \\
& \leqslant \frac{1}{h} \int_{t}^{t+h} \int_{y_{0}}^{\infty} u(s, y) \beta(y) \mathrm{d} y \mathrm{~d} s \\
& \leqslant B \sup _{s \geqslant 0}\|u(s)\|_{L_{1}(Y, y \mathrm{~d} y)}
\end{aligned}
$$

for $t \geqslant 0$ and $h>0$. Thus, (73) yields

$$
\begin{equation*}
|u(t+h)|_{1}-|u(t)|_{1} \leqslant c h, \quad t, h>0 . \tag{74}
\end{equation*}
$$

Lemma 5.1 also ensures that

$$
\begin{equation*}
\int_{0}^{\infty}|u(s)|_{1} \mathrm{~d} s \leqslant \frac{1}{p} F\left(v^{0}, u^{0}\right) . \tag{75}
\end{equation*}
$$

Combining (74) and (75), we conclude that $\lim _{t \rightarrow \infty}|u(t)|_{1}=0$, which, together with (73), warrants that for each $\sigma<1$,

$$
\begin{equation*}
u(t) \rightarrow 0 \quad \text { in } L_{1}\left(Y, y^{\sigma} \mathrm{d} y\right) \text { as } t \rightarrow \infty . \tag{76}
\end{equation*}
$$

Finally, since $\varepsilon_{1}<\infty$ both $g(u(t))$ and $|u(t)|_{1}$ tend to 0 as $t \rightarrow \infty$ due to (76). Since $v \in C^{1}\left(\mathbb{R}^{+}\right)$ solves (1), it is easy to check that $v(t)$ converges to $\lambda / \gamma$.

The result above can be improved in the case of classical solutions as follows.
Corollary 5.4. Suppose (62), (64), and (65) hold. Then the classical solution ( $v, u$ ) corresponding to $v^{0}>0$ and $u^{0} \in E_{1}^{+}$satisfies

$$
(v, u) \rightarrow(\lambda / \gamma, 0) \quad \text { in } \mathbb{R} \times L_{1}(Y, y \mathrm{~d} y) \text { as } t \rightarrow \infty .
$$

Proof. Set

$$
Q(t):=\int_{y_{0}}^{\infty} y u(t, y) \mathrm{d} y \geqslant 0, \quad t \geqslant 0 .
$$

Then $Q \in C^{1}\left(\mathbb{R}^{+}\right)$according to Theorem 3.1. From Lemma 5.1 it follows that

$$
\begin{equation*}
Q(t)+\int_{0}^{\infty} Q(s) \mathrm{d} s \leqslant c, \quad t \geqslant 0 . \tag{77}
\end{equation*}
$$

In addition, $v(t) \leqslant c$ for each $t \geqslant 0$, whence

$$
\begin{equation*}
\dot{Q}(t) \leqslant \tau v(t)|u(t)|_{1} \leqslant c, \quad t \geqslant 0 . \tag{78}
\end{equation*}
$$

Consequently, we deduce $\lim _{t \rightarrow \infty} Q(t)=0$ from (77) and (78).

## Remarks 5.5.

(a) As was pointed out in the introduction, Eqs. (1), (2) are no longer coupled in case the rates are subject to (5), since $v$ is then completely determined for all $t \geqslant 0$. In this case the results in [3] yield a semiflow in the natural phase space $\mathbb{R}^{+} \times L_{1}^{+}(Y, y \mathrm{~d} y)$, whereas Theorem 4.3 guarantees existence of weak solutions only for initial values $\left(v^{0}, u^{0}\right) \in \mathbb{R}^{+} \times L_{1}^{+}\left(Y, y^{\alpha} \mathrm{d} y\right)$ with $\alpha>1$.
However, in this particular case it can be easily verified that the function ( $v, u$ ) in (57) satisfies Definition 4.1 for any initial value $\left(v^{0}, u^{0}\right) \in \mathbb{R}^{+} \times L_{1}^{+}(Y, y \mathrm{~d} y)$, provided one takes test functions $\varphi \in W_{\infty}^{1}(Y)$ with compact support. For this one should note that $\lim _{y \rightarrow \infty} \varrho(y)=0$ is merely required for step (v) in the proof of Theorem 4.3.
(b) If the kernels are of the form (5), then we may take $d_{0}=\beta / \mu, \delta_{0}:=\beta y_{0}$ and $\varepsilon_{1}:=$ $\delta_{1}:=\beta y_{0}^{2} / 2$, so (65) is equivalent to (9). We should like to point out that in this case the authors in [3] prove that the disease-free steady state is globally exponentially stable in $\mathbb{R}^{+} \times L_{1}^{+}(Y, y \mathrm{~d} y)$, and asymptotically stable if $\beta y_{0}+\mu=\sqrt{\beta \lambda \tau / \gamma}$.
(c) If the rates are subject to (5) it has already been observed in [4] that system (1)-(2) admits also a non-trivial (disease) steady state, provided the inequality in (9) is reversed. It is shown in [3] that this steady state is again globally asymptotically stable in $\mathbb{R}^{+} \times L_{1}^{+}(Y, y \mathrm{~d} y)$. For general rates as in the present publication, existence of other equilibria besides $(\lambda / \gamma, 0)$ is an open problem.

## Acknowledgments

We thank Glenn Webb and Laurent Pujo-Menjouet for introducing us to the subject. Our thanks also go to Jan Prüss for helpful discussions and to the anonymous referee for carefully reading the manuscript and pointing out some improvements.

## References

[1] H. Amann, Linear and Quasilinear Parabolic Problems, Volume I: Abstract Linear Theory, Birkhäuser, Basel, 1995.
[2] R.E. Edwards, Functional Analysis. Theory and Applications, Dover, New York, 1995.
[3] H. Engler, J. Prüss, G. Webb, Analysis of a model for the dynamics of prions II, J. Math. Anal. Appl., in press.
[4] M. Greer, A population model of prion dynamics, PhD thesis, Vanderbilt University, Nashville, 2002.
[5] M. Greer, L. Pujo-Menjouet, G. Webb, A mathematical analysis of the dynamics of prion proliferation, preprint.
[6] M. Lachowicz, P. Laurençot, D. Wrzosek, On the Oort-Hulst-Safronov coagulation equation and its relation to the Smoluchowski equation, SIAM J. Math. Anal. 34 (6) (2003) 1399-1421.
[7] P. Laurençot, S. Mischler, The continuous coagulation-fragmentation equations with diffusion, Arch. Ration. Mech. Anal. 162 (1) (2002) 45-99.
[8] P. Laurençot, C. Walker, in preparation.
[9] A. Pazy, Semigroups of Linear Operators and Applications to Partial Differential Equations, Springer, Berlin, 1983.
[10] J. Prüss, L. Pujo-Menjouet, G. Webb, R. Zacher, Analysis of a model for the dynamics of prions, Discrete Contin. Dyn. Syst. Ser. B 6 (1) (2006) 225-235 (electronic).
[11] I.W. Stewart, A global existence theorem for the general coagulation-fragmentation equation with unbounded kernels, Math. Methods Appl. Sci. 11 (1989) 627-648.
[12] I.I. Vrabie, Compactness Methods for Nonlinear Evolutions, second ed., Longman, London, 1995.
[13] C. Walker, On diffusive and non-diffusive coalescence and breakage processes, Doctoral Thesis, Universität Zürich, 2003.


[^0]:    * Corresponding author.

    E-mail addresses: simonett@math.vanderbilt.edu (G. Simonett), cwalker@math.vanderbilt.edu (C. Walker).

