Passive Constant Flow Regulators

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Passive flow control regulators, also known as passive flow control or autoregulated or pressure-compensated valves, deliver, without external control and energy consumption, a constant flow rate regardless of pressure variations.

flow	passive	constant flow regulator	flow control valve			microfluic	MEMS	
Drug del	livery	Hydrocephalus shunt	Microvalves		Lab	-on-a-chip		
Control a	and syste	ms engineering						

1. Introduction

Passive constant flow regulators are widely used in industry for water treatment, process water control (limiting peak flow rate), water authorities (flow limiting, boost mains pressure), centrifugal pump protection, water-saving (domestic showers, drinking fountains), irrigation, etc. These valves are compact, reliable, maintenance-free, and require no energy. Manufacturers provide passive valves dedicated to high flow rate (from a few hundred ml/min to 10,000L/min and more) and large pressure differential (from 1 bar to several hundreds of bar). More recently, microvalves have been developed for microfluidic applications, including drug delivery, flow chemistry, point-of-care tests, hydrocephalus treatment, microdialysis, etc. These passive constant flow regulators fall into two main categories: the mechanical regulators having moving parts (a membrane in silicon or elastomer, a piston, a collapsible tubing, or a flap), and non-mechanical regulators without moving parts (fixed geometry), wherein the rheological properties of the fluid are exploited to achieve a constant flow rate. Most of the regulator designs belong to the first category. The automatic adjustment of the flow rate with varying pressure conditions is therefore obtained by an automatic change of the fluidic pathway dimensions^[1].

2. Working Principle

The passive constant flow regulators considered in this entry are related to microfluidics, which refers to the manipulation of fluids in channels or structures with dimensions of tens of micrometers^[2]. By contrast to macroscopic fluids, viscosity is more important than inertia, and the Reynolds number (Re) that characterizes the ratio of inertial to viscous forces on the fluid is usually lower than 2000 in most microfluidic systems. The flow in such microstructures is in the laminar regime with highly predictable fluid dynamics. The estimation of the hydraulic resistances along the fluid pathway is done using the governing equations of laminar viscous fluid flow (see, e.g.,

References ^{[3][4]}). A constant passive flow regulator is intended to deliver a constant flow rate, Q_0 , in a predefined pressure gradient range comprised between $\Delta P_{reg\,min}$ and $\Delta P_{reg\,max}$ as illustrated in Figure 1. Further increasing the pressure gradient between the inlet and the outlet can, depending on the regulator design and the target application, either stops the flow rate (passive shut-off feature) or generates a flow rate larger than Q_0 , the nominal flow rate.

At a given temperature, the flow rate ${\it Q}$ in the flow regulator is:

$$Q = \frac{\Delta p}{R_h} \tag{1}$$

where $\Delta p = P_{out} - P_{in}$ is the pressure drop between the inlet and the outlet of the device and R_h is the hydraulic resistance of the device. According to Figure 1, the hydraulic resistance is constant and equal to $R_{h0} = \frac{\Delta P_{reg\,min}}{Q_0}$ if $\Delta p \leq \Delta P_{reg\,min}$. A further increase δp of the pressure gradient, with $\delta p \leq \Delta P_{reg\,min}$, induces an increase of the hydraulic resistance, δR_h . Using (1), we obtain:

$$\Delta P_{
m reg\,min} + \delta p = (R_{h0} + \delta R_h) Q_0$$
 (2)

and thus:

$$\frac{\delta R_h}{\delta p} = \frac{1}{Q_0} = Constant \tag{3}$$

To maintain the flow rate constant in the gradient pressure range $\triangle P_{reg\ min} \leq \triangle p \leq \triangle P_{reg\ max}$, the hydraulic resistance of the passive regulator shall linearly increase with the pressure gradient up to $\triangle P_{reg\ max}$,

where
$$R_h(riangle P_{reg\ max}) = rac{ riangle P_{reg\ max}}{Q_0}.$$

The change of slope in the $Q(\Delta p)$ curve is representative of a nonlinear hydraulic resistor. At higher pressures, the regulator could further increase its hydraulic resistance until the flow is stopped or simply keep the hydraulic resistance constant (see Figure 1). A free-flow at high pressure can be obtained by fully by-passing the nonlinear

resistor of the device (possible option for the piston spring valve^[5]) or by adding a crack valve in parallel to the flow regulator (option implemented in the CRx Diamond Valve manufactured by Phoenix Biomedical Corp.^[6]).

The profile shown in Figure 1 is, therefore, representative of a three-stage valve:

- Stage I—low-pressure stage, with low hydraulic resistance for $riangle p \leq riangle P_{reg\ min}$
- Stage II—flow regulation stage, for $riangle P_{reg\,min} \leq riangle p \leq riangle P_{reg\,max}$
- Stage III—high-pressure stage, with a low or high hydraulic resistance for $riangle p > riangle P_{reg\ max}$



Figure 1. Ideal flow rate versus pressure gradient characteristic of a three-stage passive constant flow regulator. At low pressure (stage I), the regulator has a constant and low hydraulic resistance. In stage II, the flow is regulated at Q_0 for a pressure gradient in the range $[\triangle P_{reg\ min}; \triangle P_{reg\ max}]$. At high pressure (stage III), the device could, according to its specific design, either stop the flow at $\triangle P_{off}$ (solid line) or conversely, deliver a large flow rate (dashed line).

The passive constant flow regulator is intended to be placed into a fluidic circuit. To keep the functioning point of the device onto the plateau of the $Q(\triangle p)$ characteristic, it is recommended that the device exhibits the largest hydraulic resistance of the circuit.

An alternative method to maintain the flow rate at Q_0 consists of the control of the pressure at the inlet of a flow restrictor. This method is exploited in constant flow rate implantable pumps for pain management (e.g., Infusaid-400, Medtronic Isomed, Codman 3000, Tricumed IP2000), wherein the reservoir pressure is generated by the vapor pressure of propellant (typically **2.5** *bar*), or in elastomeric pumps dedicated to ambulatory infusion of medication^[7]. In both cases, a fluidic restriction (capillary) is used to set the flow rate in production, but the strategy to maintain the flow rate during the reservoir emptying is different. The propellant vaporizes progressively as the

drug is infused toward the patient, while in the second type of device, the elastomeric balloon is designed to keep the pressure almost constant during the treatment. A discussion of the flow rate accuracy of constant implantable pumps is provided by Dumont-fillon et al.^[8]. For disposable and other portable infusion pumps, including devices powered by a spring, an elastomeric balloon, a chemical reaction, or a vacuum, overall flow rate accuracy of $\pm 40\%$ was reported^{[9][10][11]}. Also, these devices are sensitive to outlet pressure variations and temperature-induced change of the viscosity.

The combination of a pressurized reservoir and a passive constant flow regulator is recommended to avoid startup overflow and the constant decrease of the infusion rate over time as well. The infusion duration is better controlled and a flow rate error less than $\pm 10\%$ can be achieved^[8].

3. Working Fluid

Except for the device developed by Groisman et al.^[12], almost all passive constant flow regulators dedicated to microfluidic applications were designed to work with water, cerebrospinal fluid (CSF), or other Newtonian fluids like viscous mixes of water and glycerol. In principle, the constant passive flow regulators are also compatible with gas. The modeling of a gas flow regulator can still be based on the standard continuum approach, which is still valid if no-slip boundary conditions apply, typically for Knudsen number $Kn < 10^{-2}$ (viscous flow), where Kn is defined as the ratio of the fluid mean free path and the macroscopic length scale of the physical system^[13].

In the laminar regime, a change of temperature induces a change of fluid viscosity which, in turn, leads to flow rate variability as for standard flow restrictors. Since the hydraulic resistance is proportional to the fluid dynamic viscosity η , the relative flow rate error is thus proportional to the opposite of the relative viscosity change with temperature, considering that the reference temperature is $20^{\circ}C$:

$$Q(T) \propto \frac{1}{\eta(T)} \rightarrow \frac{Q(T) - Q(20^{\circ}\text{C})}{Q(20^{\circ}\text{C})} = -\frac{\eta_T - \eta_{20^{\circ}\text{C}}}{\eta_T} = \frac{\eta_{20^{\circ}\text{C}}}{\eta_T} - 1.$$
(4)

Table 1 shows, at selected temperatures, the relative flow rate error due to dynamic viscosity change. The values are derived from the empirical formula proposed by Kestin et al. for the dynamic viscosity of water in the range -8 to $+150^{\circ}C^{[14]}$. Relative flow rate errors vary from -34% to +53% for operating conditions in the range $[+5; +40]^{\circ}C$. It is noteworthy that additional error sources shall be considered to determine the passive flow regulator accuracy (e.g., fluctuations in the flow regulation stage, hysteresis).

Table 1. Effect of dynamic viscosity change with temperature on relative flow rate accuracy of passive constant flow regulators. The reference temperature is $20^{\circ}C$.

Temperature (°C)	5	10	15	20	25	30	35	40
Water dynamic viscosity (mPa.s)	1.520	1.308	1.139	1.002	0.890	0.797	0.719	0.653
Relative water flow rate error ($\%$)	-34.1%	-23.4%	-12.0%	0.0%	12.6%	25.7%	39.3%	53.5%

4. Materials

Silicon, glass, and poly(dimethylsiloxane) (PDMS) are the most common materials in contact with the working fluid in the passive constant flow regulators reported here. Titanium^{[5][15]} was also considered for implantable devices. Early microfluidic and Micro Total Analysis Systems (µTAS) were made in glass and silicon due to the availability of process equipment and etching recipes. Since the Whitesides' group manufactured, in 1998, microfluidic devices in PDMS by soft lithography, PDMS became the preferred material for rapid prototyping of microchannel systems for use with biological samples [16][17][18]. Indeed, PDMS is an inexpensive, optically transparent, gas and vapor permeable, hydrophobic, relatively inert, and biocompatible polymer that, moreover, cures at low temperatures. PDMS is soft and facilitates removal from Si molds for micron-scale feature replication. The surface energy of PDMS can be tuned to increase its wettability. Also, PDMS can be sealed reversibly to itself or other materials. Last but not least, the elasticity of PDMS has been widely exploited to create membrane pumps and check valves in active microfluidic devices for cell sorting and biochemical assays^[19]. PDMS was also considered to make passive flow regulators because this material enables the rapid prototyping of complex three-dimensional (3D) structures. On the other hand, PDMS presents, compared to Si and glass, some disadvantages that can preterite its use as raw material for specific microfluidic applications that require long-term stability, reproducibility, well-controlled surface chemistry, high accuracy, stiffness, high temperature, compatibility with organic solvents, or low gas/vapor permeability. Si/glass biomedical electromechanical systems (BioMEMS) are more expensive to produce at low volume and require interconnections to the other parts of the fluidic systems. Silicon is a rigid material with a welldefined Young's modulus by contrast to the hyperelastic PDMS, whose mechanical properties are highly dependent on its degree of cross-linking and stretching. Si membranes exhibit a highly reproducible displacement and no fatigue, and the electronic properties of silicon enable the smart integration of reliable sensors^[20]. Also, the negative charge surface of glass and silicon support electroosmotic flow (EOF).

5. Modeling

The modeling of the passive constant flow regulator can be based on analytical models^{[5][21][22]}, but in a general way, 3D simulations using finite element methods are required to model the fluid–structure interactions (FSI) and

the nonlinear behavior of membranes or flaps under large deformations. Numerical simulations are also useful to investigate the impact of machining tolerances on flow rate accuracy^[8]. Specific tools using a genetic algorithm have been made to increase the yield in production by design optimization. This method was used to design a regulator with a flexible Si membrane that deflects against pillars. In addition to standard machining tolerances, other non-idealities have been introduced into the model, like wafer misalignment or valve tilt (due to particle contamination for instance) to increase the design robustness^{[21][23]}.

A passive flow regulator is usually part of a complex microfluidic system. In addition to these quasi-static analyses of the fluid flow in fixed pressure conditions, further simulations shall be carried out to calculate the fluid dynamics in a system that can include microchannels, check valves, micropumps, pressure sources, etc. As a passive flow regulator induces nonlinear effects, the equations governing the fluidic behavior of such systems generally have no analytical solutions. A classical approach consists of building an electrical equivalent network based on a subdivision of the system into lumped elements and the analogy between fluidic, mechanical, and electrical parameters^{[24][25]}. The impedance of each element is estimated beforehand. For a passive flow regulator with an integrated check-valve, its electrical analog is a diode in series with a variable resistor. The analysis of the dynamics of complex systems can then be carried out very quickly with an electrical simulation tool. This method was chosen to simulate the dynamics of a hydrocephalus shunt comprising a passive flow regulator to divert CSF from the brain ventricles to the peritoneal cavity^[26]. The ventricles are equivalent to a constant current source that produces about 20 mL of CSF every hour. The numerical model allowed the determination of the time evolution of intracranial pressure during patient postural changes, changes in the CSF formation rate, oscillations due to the vasogenic system, etc., together with a direct comparison with other shunt designs. Dynamic simulations based on electrical equivalent networks are particularly useful during the design phase to determine the behavior of the passive flow regulator. For a complex system such as the hydrocephalus flow control valve, the model can be further refined by the introduction of experimental data[27].

6. Recent trends and future prospects

Active fluid control poses numerous problems of integration and miniaturization, and the development of microvalves and micropumps, which are two important building blocks of a microfluidic platform, was stimulated by recent advances in lab-on-a-chip for diagnostics applications, drug delivery, flow chemistry, point-of-care testing, etc. Compared to active microvalves^[28], passive constant flow control valves provide a simpler way to regulate fluid flows without external control and energy consumption. A comprehensive design review of the passive constant flow regulators developed over the last three decades can be found in Reference^[1]. The designs of typical passive constant flow regulators made using MEMS technology and PDMS are shown in Figures 2 and 3 respectively.



Figure 2. Cross-section of a passive flow regulator with a top Silicon-On-Insulator (SOI) wafer (membrane) and a bottom wafer (substrate) made of Pyrex or silicon (Si). The fluid flows through the inlet holes of the membrane, the radial diffuser (annular restriction between the membrane and the pillar), and the outlet hole of the substrate. As the pressure increases, the deflection of the membrane induces an increase in the hydraulic resistance of the radial diffuser to keep the flow rate constant. The flow direction is indicated by red arrows^[21].

Figure 3. Schematic drawing of a passive flow regulator using autonomous deflection of parallel membrane valves in PDMS. The increase δp of both the inlet and sub-channel pressures induces a deflection of two membranes. The narrowing of the fluid path leads to an increase δR_h of the main channel hydraulic resistance (adapted from Reference^[29]).

The modeling, the fabrication methods, and the resulting performance of these passive microvalves have been constantly improved. Table 2 summarizes the principal pros and cons of each category of passive flow regulators.

Table 2: Comparative table of the different categories of passive constant flow regulators

	Pressure Range	Flow Rate Range	Accuracy/Reproducibility	Connectivity	Fabrication Cost	Prototyping Complexity	Aqueous Solution
MECHANICAL							
Si membrane(s)	Very large	Very Iarge	High/Very High	Medium	Low*/High	High	Yes
Elastomeric membrane(s)	Large	Very Iarge	High/Medium**	High	Low/Medium	Low/Medium	Yes
Tube with lateral aperture	Small	Small	Low/Low	Medium	Low	Low	Yes
Piston	Large	Large	High/Medium**	Medium	Medium	Medium	Yes
Flap	Large	Large	High/Medium**	High	Low/Medium	Low/Medium	Yes
NON- MECHANICAL							
Rheological	Small	Small	Low/Medium**	High	Low	Low	No

*At large volumes only; ** Estimation.

Hysteresis was not included in this comparative table because this effect should be better characterized. Besides, considerations relative to the manufacturing reproducibility of the passive flow regulators made in PDMS are still speculative and have yet to be demonstrated. Recent advances in the development of these devices are promising but Table 2 also suggests that there is still room for further developing the technology and designing passive constant flow regulators that ideally combine the reliability and performance of MEMS-based valves with the ease of production and the large integration capability of their PDMS counterparts.

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