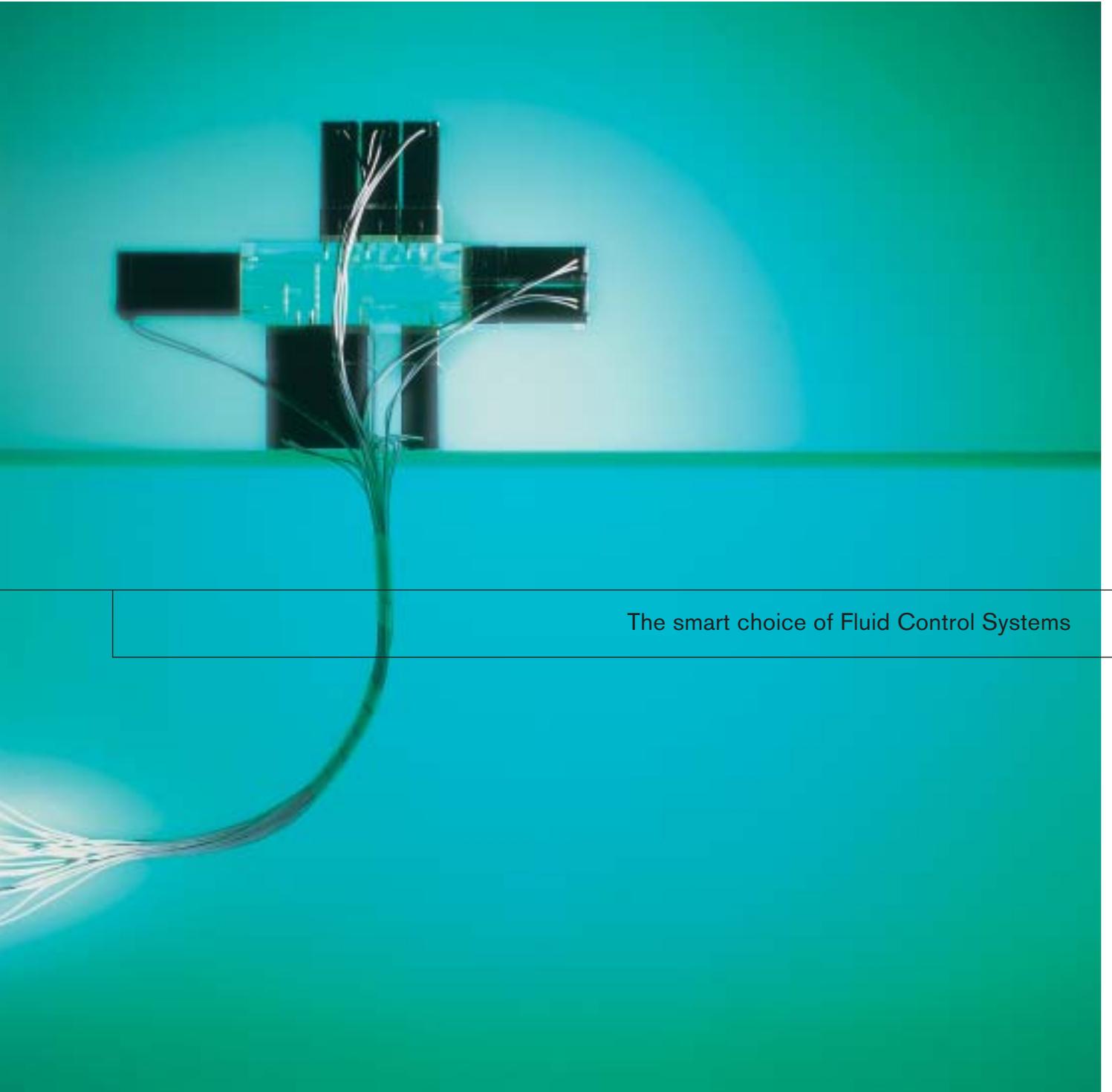


# System Catalog 5

Solenoid valves | Process and control valves | Pneumatics  
Sensors | **MicroFluidics** | MFC and proportional valves



The smart choice of Fluid Control Systems



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# The smart choice of Fluid Control Systems

Your advantage with Bürkert technology

There are only a few white specks on the fluidics world map where a Bürkert solution has not yet been used. And these specks are becoming fewer and fewer with our range of MicroFluidics product and services.

In the same way as the range of applications of fluid control technology is constantly expanding, Bürkert is also expanding its core competence and using the synergy effects of a wide variety of company divisions to meet the new, diversified tasks.

The approaches to solving problems are "fluidic", since in addition to the international experience and know-how of a technology leader, tailored, customer-orientated services are also an essential part of our company.

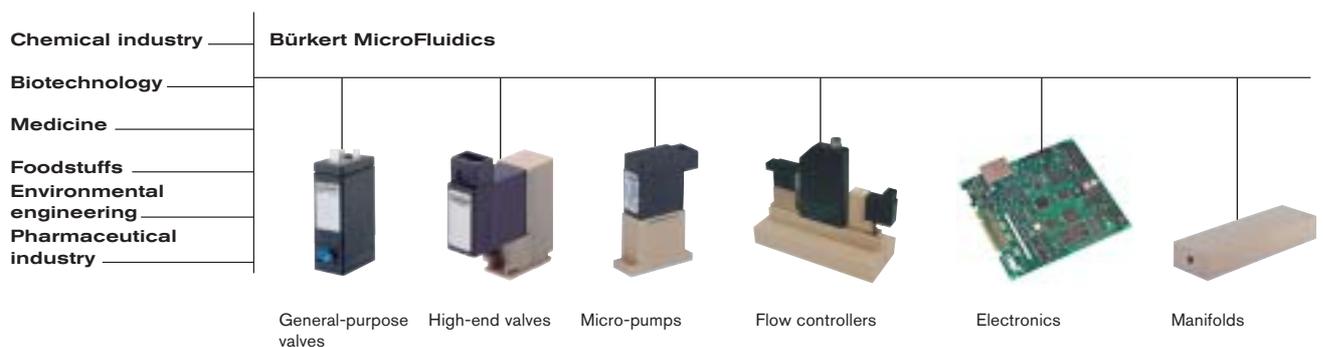
Depending on the specifics of the problem, it is possible to obtain application consulting for serial production and engineering, in addition to development and production of individual systems. Whether they require single components or system solutions, Bürkert customers can always benefit from inclusive innovation, an approach geared both technically and economically to the future.

## New strengths – Bürkert MicroFluidics for medical technology, analysis technology and biotechnology

Along with automation of process sequences, miniaturization of devices is defining the current "trend" in pharmaceuticals, the chemical industry, biotechnology and medical instruments. Controlled and automated handling of ultra-small fluid quantities is in demand. With its MicroFluidic components, Bürkert meets the demand for ultra-

small valves, pumps and flow controllers which enable new processes, e.g. processes in genetic research and pharmaceuticals.

Likewise, successes in biotechnology, genetic engineering and medicine are linked to the use of miniaturized components in workstations and automatic analyzers. Here as well, Bürkert technology makes the small but crucial difference in achieving a growth market with a great future. "What is hot and what is not" – customers throughout the world benefit from the knowledge and experience of a technology leader.



## MicroFluidic components



**Bürkert  
Ingelfingen headquarters  
(Germany)**

- Application consulting
- System development
- Standard component development

**Main focal points of competence: medical technology, analysis technology and biotechnology**



**Bürkert  
Service Center Dresden  
(Germany)**

- Electronics development
- System development
- Job-lot and prototype production
- System production



**Bürkert  
Gerabronn plant  
(Germany)**

- Large-scale serial production and standard device production
- System production
- Electronics development
- System development

### **Dynamics**

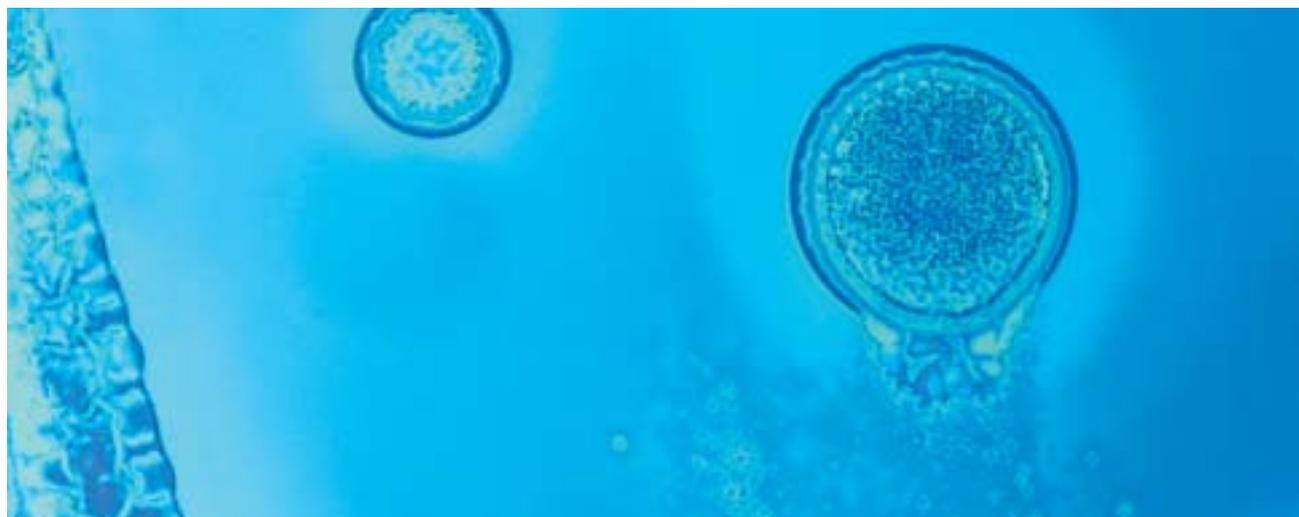
#### **thanks to distributed competence**

A booming market demands fast and flexible solution approaches. That is why Bürkert has set up regionally distributed competence centers for its customers in the MicroFluidics sector. Customers can make use of the entire range of specific services, locally and in their own vicinity, from application consulting through development to production.

Internal competence and transfer of know-how ensure that the best possible solution is selected in each individual case.

# 1. Small, strong and flexible

Fluid control in medical technology, analysis technology and biotechnology



Fluid control ranks as one of the basic technologies in medical technology, analysis technology, biotechnology and related sectors, due to the fact that the substances are primarily present in liquid form, whereby automation and, in particular, miniaturization, are the currently prevailing trends. Existing devices and processes thus become more efficient (faster sequences, higher throughput, lower substance quantities, less solvents and lower costs), and new processes, for example in genetic research (genomics and proteomics) or the pharmaceutical industry, are enabled.

## **1.1.** **Liquid handling**

Liquid handling is the control and automated handling of small and ultra-small fluid quantities in a wide variety of devices and processes. At the same time, it also refers to a sub-function in extensive, modular-structure equipment systems for conducting analyses of all types (screening), for which terms such as workstation or automatic analyzer have also become common. Such devices have become an absolute prerequisite for success in biotechnology, genetic engineering, medicine and other related areas.

## **1.2.** **Bürkert MicroFluidics**

Bürkert has developed an extensive range of miniaturized components (MicroFluidic components), such as

valves, pumps and flow controllers, and the connection manifolds (manifolds) required for setting up the functional units that specifically implement the above trend.

Both the overall dimensions (width per station down to 10 mm) and the performance data (dosing rates down to a few nanoliters) of these components meet the requirements demanded for miniaturization. Components of this size and performance are called MicroFluidic components.

### **Three essentials for MicroFluidic components**

#### **1. Cost-conscious miniaturization**

In the fields of application above, a frequent task involves handling fluids that are available only in very small quantities and are very expensive and which, in addition, often cause an aggressive chemical reaction.

Accordingly, the logical requirement is for:

- Small internal volumes and overall dimensions (miniaturization)
- Low dead volume and good flushability (gap-free shaping of the interior)
- Media separation
- Use of inert and resistant materials.

### 2. Reliable continuous operation

Many applications necessitate operation of devices round the clock to ensure the required high sample throughput.

Accordingly, the logical requirement is for:

- Freedom from wear and tear
- Up to  $10^8$  valve cycles
- Short valve response times
- Low power consumption.

### 3. Modular flexibility

The devices constructed using Micro-Fluidic components must perform greatly differing tasks with frequently similar sub-functions, e.g. dosing or mixing, in their overall function.

Accordingly, the logical requirement is for:

- Various body variants
- Interchangeability of components
- Modular system with standard center spacings
- Application-specific additional developments.

Diverse applications of MicroFluidic components in devices and processes	
Anesthetics systems	Gas mixers
Analysis platforms	GC/MS coupling systems
Analysis robots	Gel electrophoresis systems
Analytical multi-test units	HPLC units
Auto-analyzers	Incubators
Automatic filling machines	Industrial inkjet printing
Automatic high-throughput units	Liquid chromatographs
Automatic pipetting units	Liquid handling systems
Automatic screening units	Liquid transfer systems
Auto-samplers	Mass spectrometers
Biochip readers	Micro-arrays
Biology workstations	Micro-reactors
Biomarkers	Multi-reaction systems
Blood-purifying units	PCR systems
Cell counters	Proteomics analyzers
Cell separators	RNA purification systems
Chromatographs	Robot platforms
CIP systems	Robotic systems
Continuous-flow analysis systems	Sample processors
Kidney dialysis machines	Sample purification/upgrading syst.
Diluter	Sample switches
Dispensers	Samplers
DNA analysis robots	Tissue engineering
Downstream systems (cell harvest)	Titration
Flow cytometers	TOF mass spectrometers
Flow systems	Washing devices
Fraction collators	

### From A(nalysis platform) to W(ashing devices)

The table above lists the diverse applications, which are discussed in the following by way of example.

.



### Specific MicroFluidic tasks and requirements

- Sampling from a container or a sample stream, if applicable with sample switching
- Sample preparation and conditioning (mixing and dilution...)
- Sample transport to the analyzer
- Sample application into the measuring system
- Sample disposal after measurement

Modern analyzers feature a very high level of automation and frequently combine the above processes into one single unit. It is not without reason that the term “workstation” has become the accepted term for such automatic units, primarily in genetic engineering.

## 2.2. Liquid chromatography, HPLC

“Chromatography” is an umbrella term covering physical methods in which substances are separated via distribution between a stationary and mobile phase. HPLC (High-Performance Liquid Chromatography) is a separating process in which a liquid sample is transported through a stationary phase under high pressure. Depending on the nature of the interaction between the phases and the sample, we distinguish between adsorption chromatography, partition chromatography, ion-exchange chromatography, size-exclusion chromatography and affinity chromatography. The processes of adsorption chromatography and partition chromatography are the chief processes used in HPLC. HPLC uses partition particles with grain sizes of a

few  $\mu\text{m}$ ; it thus achieves high numbers of plates or trays, but at the same time, requires a high counter-pressure to be overcome during transport of the mobile phase through the thin separation column (of a few millimeters diameter). All parts must be interconnected without any dead volume wherever possible and must be pressure-stable (up to 300 bar).

An HPLC unit consists of supply unit (mobile solvent reservoir and pump), injection system, separation column, detector, fractionating unit and evaluation system. The sample is initially injected free of pressure into a sample loop. It is then transported into the separation column by appropriate switchover of the eluent stream. The composition of the mobile phase and, thus, its solvent strength parameter is frequently varied during analysis in the case of separation of complex mixtures and in order to shorten analysis time.

### Specific MicroFluidic tasks and requirements

- Sample preparation
- Sample transport
- Mixing the eluent
- Disposal of the sample fluid
- Collection of the fractions (in the case of preparative use)

## 2.2. Mass spectrometry

Mass spectrometry (MS) is a firmly established part of biotechnology, specifically, for example, for analysis of proteomes. It allows unique identification of the proteins after they have been separated or when they are in protein mixtures.

There are a large number of variants, such as:

- Time-of-flight mass spectrometers (TOF), combined with chromatography for removal of non-bound substances on the protein-chip array
- Surface-enhanced laser desorption/ionization (SELDI process)
- Matrix-assisted laser desorption/ionization mass spectrometry (MALDI process).

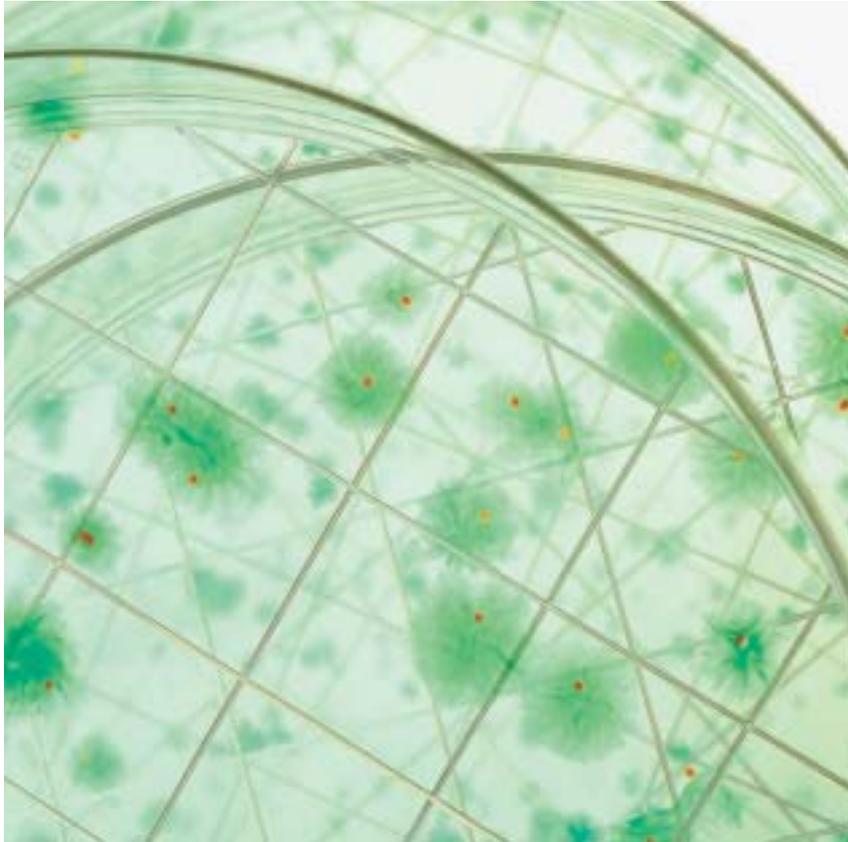
Progress in MS enables protein and genetic analyses with a high throughput. The speed of these analyses necessitates automation of the entire sample handling process.

## 2.3. Electrophoresis

This term refers to the transport of charged particles in an electrolyte solution under the influence of an electric field. It is used for analytical and preparative separation of mixtures of dispersed particles, since individual substances feature different migration rates.

# 3. All about fungi, lightning flashes and cell harvest

MicroFluidics in biotechnology



## 3.1. Gas mixing for fermenters

Biotechnology works with microorganisms such as bacteria, yeast fungi or other fungi, thus allowing the process to run under normal pressure and temperature conditions and in aqueous environments. This is a crucial difference with respect to chemistry where the production processes run subject to high pressures and temperatures and are thus very energy-intensive and frequently include the use of acids or solvents.

However, since the metabolic rates of microorganisms are relatively low, it is necessary to create an environment for them that allows the metabolic process to run optimally. These conditions, some of which are very specific, are created in bioreactors (fermenters), which can be operated intermittently or continuously.

Fermenters are available in an extremely wide variety of sizes, from the bench unit up to building-size reactors for industrial production processes. Parameters for an optimum bioreaction include a defined gas atmosphere in addition to the nutrient content and

pH value (which are adjusted using correction fluids). Oxygen, nitrogen and carbon dioxide in certain mixing ratios, specific to each cell culture, are crucially significant to successful fermentation.

This task is performed by compact gas mixing units, primarily in the case of small fermenters, whose function is ensured by MicroFluidic components.

## 3.2. Downstream systems

After fermentation (i.e. "downstream"), there is generally a process stage of "cell harvesting" via continuously operating separators. Further purification/upgrading of the products is then performed via diverse filtration and chromatography steps.

This downstream area is also an important application field of MicroFluidic components.

### Specific MicroFluidic tasks and requirements

- Simultaneous transport of several gases
- Individual gas mixing for simultaneously connected culture flasks
- Measurement and control of the mass flow rate in the range 0.02 to approx. 50 NI/h
- Pressure measurement (input pressure of the gases)
- Short reaction times, low dead volumes

### 3.3. Microreactors

There is a clearly recognizable trend in chemistry towards smaller reactors, specifically for research and development, but also for production of fine chemicals with production volumes of a few metric tons per annum.

In principle, microreactors are nothing other than reactors as used in the chemical and pharmaceutical process engineering sectors, but which range in the  $\mu\text{m}$  to  $\text{cm}$  range instead of  $\text{dm}$  to  $\text{m}$  range.

Due to their size, they offer certain advantages: for example, they can be operated isothermally despite strong exothermal reaction. This avoids hot spots and means that technical processes can be run at higher temperatures than on conventional systems. Reducing travel, extremely short dwell times, improved heat supply and dissipation and faster response to changed reaction conditions are other resultant phenomena. It is thus possible to achieve enhanced process safety (e.g. in explosion-hazard areas), enhanced process control and enhanced production efficiency.

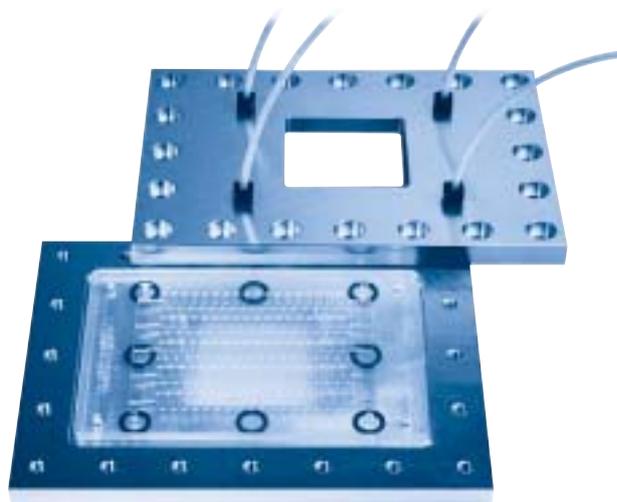
Despite the compact size, it is certainly realistic to produce several metric tons of fine chemicals in reaction container with a few milliliters volume.

Example applications of microreactor technology are as follows:

- Partial oxidation, dehydration (strong heat tonality, short reaction time, thermodynamically unstable, reactive intermediate as target product)
- Distributed production of hazardous substances (e.g. HCN and HF)
- Implementation in mobile reactors (e.g. methanol reforming)
- Mass screening of catalysts and active substances (small substance quantities)

#### Specific MicroFluidic tasks and requirements

- Mixing different media in the case of freely adjustable volume flows
- Transport by pumps or pressurization
- Measuring the flow volumes
- Measuring pressure and temperature
- Switching various infeed streams and discharge streams
- Low dead volumes
- Monitoring process parameters



**Heat exchanger**  
5 channels connected in parallel  
Fluid 1:  $700\ \mu\text{m} \times 1000\ \mu\text{m}$   
Fluid 2:  $400\ \mu\text{m} \times 1000\ \mu\text{m}$   
Heat-transfer layer:  $200\ \mu\text{m}$

Source: MGT mikroglass technik AG

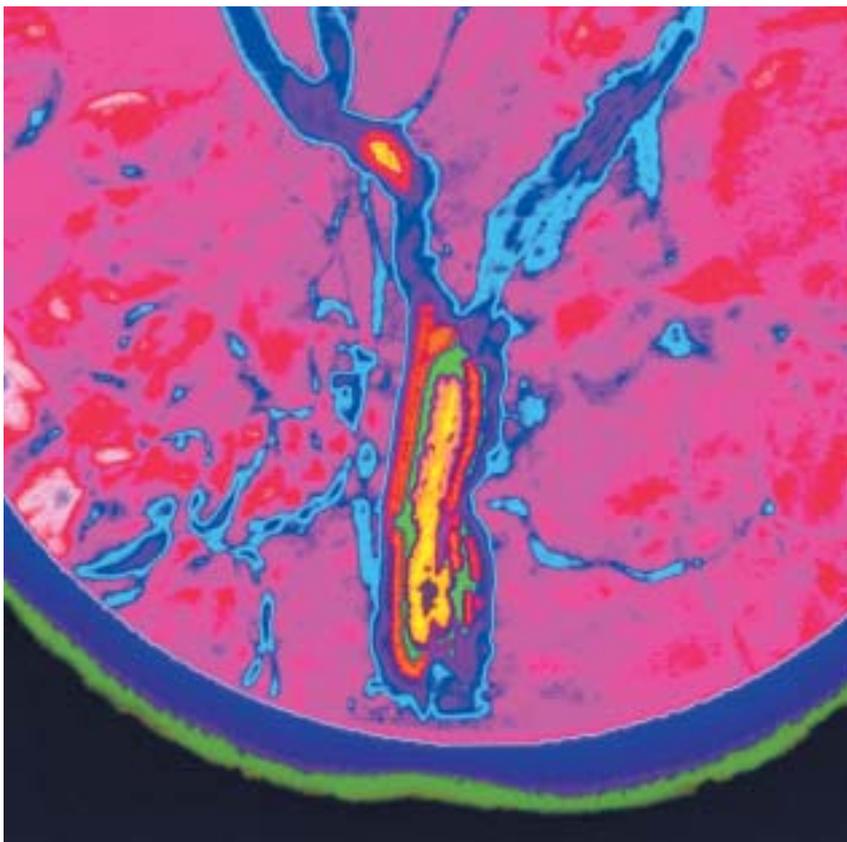
### 3.4.

## Multi-reaction systems

These systems cover devices for simultaneously conducting reactions under individually adjustable conditions such as temperature, pressure and composition. This involves the need for a defined admixing of gases, gas mixtures and/or fluids.

#### Specific MicroFluidic tasks and requirements

- Mixing different media in the case of freely adjustable volume flows
- Transport by pumps or pressurization
- Measuring the flow volumes
- Measuring pressure and temperature
- Switching various infeed streams and discharge streams
- Short response times, low dead volumes



### 3.5.

## Flow cytometry, FACS (Fluorescence Activated Cell Sorting)

Flow cytometry is a method for counting, sorting and isolating particles (e.g. living cells) which, after being specifically marked with a fluorescent dye, are individually moved past an optical detector system in laminar flow and then counted. By deriving parameters from this counting operation, it is possible to use this method to analyze many characteristics of the particles, whereby the property to be determined can be selected by the specific marking.

One promising, current application is dye marking and isolation of antigen-specific cells within the framework of developing treatment methods for e.g. viral infections or cancer.

#### The individual steps of FACS:

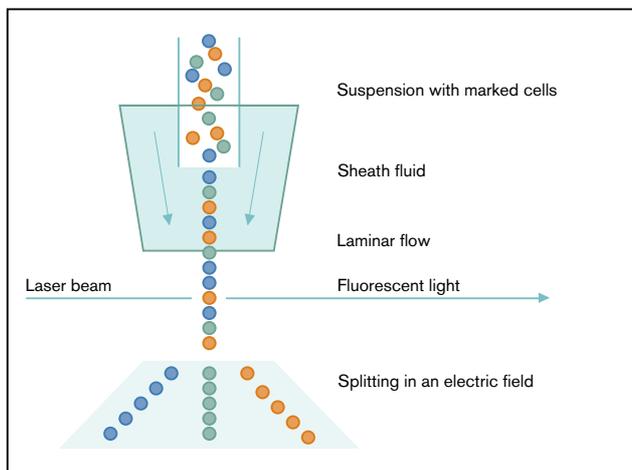
Using various mechanisms, the cell (to be more precise, a certain subset of it) is marked with a specific fluorescent dye. The individual, marked cells, then present in suspension, are transported (by compressed air) into a capillary and, from there, into the measuring chamber. In this case, an additionally applied sheath fluid comprising degassed water produces a laminar flow which allows the particles to individually pass by the actual measuring point at high velocity.

This is irradiated with a light source (laser), whereby the marked particles passing by generate fluorescent light flashes. By counting these signals, it is possible to determine the flow rate of the cells in question and represent it in histograms.

A piezoelectric transducer allows the cell stream to be split into individual droplets at the end of the measuring chamber. These droplets are electrically charged if necessary, triggered by the corresponding light signal and deflected in a downstream electrical field, thus segregating them off from the stream.

#### Specific MicroFluidic tasks and requirements

- Supply of the sample fluid
- Supply of the marking fluid
- Supply of the sheath water
- Transport and collection of the sorted cells
- Transport and storage of "waste"



Basic mode of operation, FACS

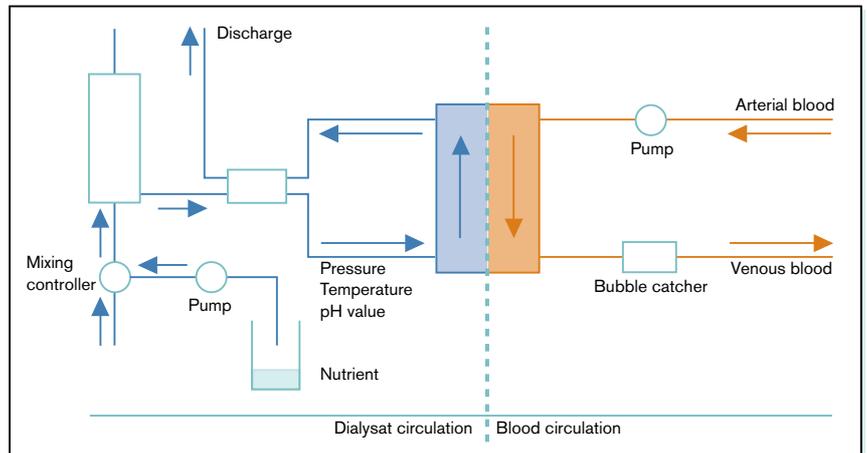
# 4. Technology for infusion, hemodialysis and anesthesia

MicroFluidics in medicine

## 4.1. Kidney dialysis machines

In the generic sense, dialysis is a process for separating low-molecular-weight substances from a fluid by diffusion of these substances through a semipermeable membrane. Use of dialysis in medicine is more than familiar as the principle of the “artificial kidney” for hemodialysis. Similarly, however, dialysis is also used in biotechnology as a “membrane bioreactor” for conducting fermentation processes under specific boundary conditions.

Dialysis in medicine largely replaces the function of the kidney by filtering out water, metabolic end products (e.g. urea) and other toxins from the blood. Blood purification is performed in the dialyser by means of a semipermeable membrane which separates the circuit of the blood to be purified from that of the purification fluid (dialysate). The toxic substances and water from the blood pass through the membrane into the dialysate and are thus removed. The dialysate (deionised water) is enriched with nutrients and its nutrient content, temperature and other parameters are constantly monitored.



Dialysis process

MicroFluidics, including an automatic control and monitoring system, forms the technical basis of a kidney dialysis machine, whose functions are transport, dosing, mixing, distribution and measurement.

### Specific MicroFluidic tasks and requirements

- Circulating the blood
- Transporting the purification fluid
- Dosing the nutrient solution

## 4.2. Adsorbers

In the case of the adsorber process, pathogenic substances, e.g. cholesterol, are removed from the blood outside of the body. The blood flows out of the patient’s vein into the adsorber in which suitable materials selectively remove the substances from the blood. A special machine ensures that the blood circulates through the adsorber.

## 4.3. Cell separators

The cell separator is a device for obtaining blood components. Fields of application include removal of diseased cells or blood plasma from a patient’s blood or collecting cells from a patient’s own blood for subsequent back-transfusion.

#### 4.4.

### Infusion units

Infusions of fluids in medicine are used for various purposes:

- Compensating for water losses
- Producing/maintaining the normal intracellular and extracellular electrolyte concentration
- Normalizing the acid-base balance
- Covering the energy and protein requirement
- Supplying medications, hormones and trace elements.

In addition to the frequently used gravity infusion method, active infusion units are used as well. Stationary infusion units are used e.g. in operating rooms for "total intravenous anesthesia" (TIVA pump), on intensive-care wards and normal wards (syringe pumps) and in accident and emergency medicine (mainly pneumatically operated pressure infusion units). However, portable systems are also increasing in importance. They are required, for instance, for chemotherapy, analgesic therapy (PCA pump: patient-controlled analgesia), parenteral feeding and for treating type-1 diabetics with insulin pump therapy.



The range of application of MicroFluidics extends from simple systems to highly integrated, closed-loop-controlled systems which operate with flow rates in the ml and  $\mu$ l range and, in some cases, also in the nl range.

#### 4.5.

### Anesthetics equipment

Anesthetics equipment contains all components required for performing inhalation anesthesia, such as gas mixer and gas dosing systems, vaporizers and gas extraction. On modern units aim at achieving a largely closed circuit with a low fresh gas supply that, besides achieving a lower gas consumption, also affords the advantage of lower environmental pollution. The fresh gas supply lies in the range 0.5 to 1 l/min.

# 5. On better-tolerated vaccines and decoding the human genome

MicroFluidics in genetic engineering



## 5.1. What does genetic engineering do?

The idioplasm or hereditary substance of virtually all living beings or organisms, from bacterium to humans, consists of double-strand DNAs and is subdivided into sections (genes).

Genes cause the development of a characteristic, in most cases production of a protein. This conversion of a gene into the protein produced by it is performed in two steps:

- transcription (transfer) of the DNA to a "messenger RNA" (mRNA) and
- translation of the messenger RNA to the corresponding protein.

All genetic engineering procedures aim at targeted modification of the hereditary material and subsequent admission of the modified (recombined) genetic material in living cells (host cells) in order to trigger them to produce specific proteins that they would otherwise not produce.

This process can be subdivided as follows:

### Step 1

Modification of the hereditary substance is performed with the aid of specific enzymes (restriction enzymes), which are able to dissect the backbone of a DNA at precisely defined positions.

### Step 2

In order to recombine the (dissected) DNA, the fragments are in turn bonded together again (with a changed frequency) by special enzymes (ligases).

### Step 3

In order to multiply the recombined DNAs, they are transferred to microorganisms, such as bacteria, yeast fungi or cell cultures (transformation). One method of doing this is electroporation or direct injection (if using animal cells).

### Step 4

Not all host cells accept transformed DNA. Consequently, a selection medium must be used to separate the transformed cells from the non-transformed cells. This is generally done using specific antibiotics which cause the non-transformed cells to die off.

## 5.2.

### Polymerase Chain Reaction, PCR

A specific DNA quantity is required for DNA analysis. Previously, preparing this quantity from small quantities of biological parent material was a difficult and very time-consuming process. The PCR method, which was invented in 1983, has eliminated this problem since this method allows DNA fragments to be multiplied exponentially, similar to a nuclear chain reaction, in a simple way by passing them through specific temperature cycles. The steps of the cycle are as follows:

- Denaturation of the double-strand DNA by heating to 94 °C. This separates it into individual strands
- Addition of the primers (starters) and cooling to approx. 50 °C. This

causes hybridization of the primers with the ends of the individual DNA strands and prevents the combination of the single strands

- Heating to 72 °C and addition of a DNA polymerase (group of enzymes), among other substances. This supplements the two single strands to form two double strands.

By repeating this cycle in modern automatic machines, it is possible to produce 1 million “copies” from only one starter molecule within approx. 3 hours!

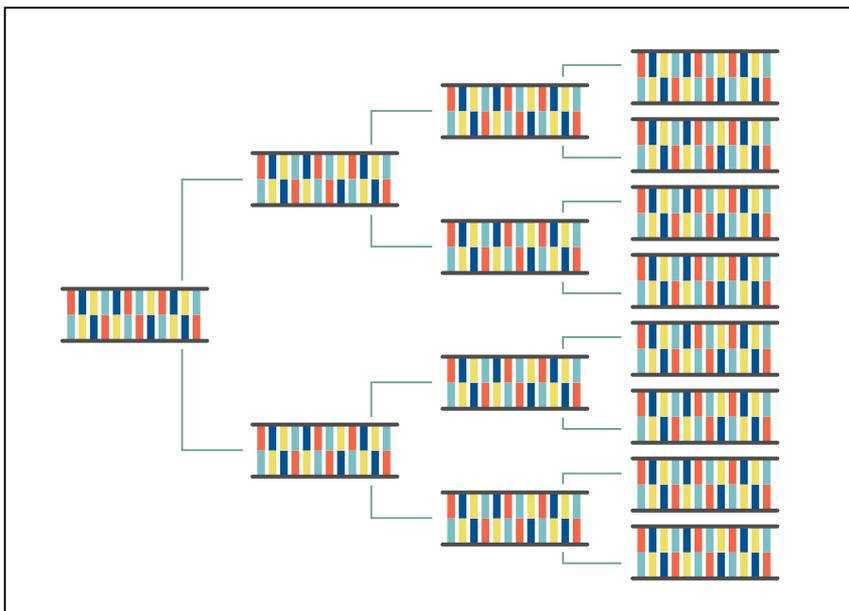
It is only with the aid of this low-cost and effective preparation method that DNA analysis and, thus modern genetic research, has been able to achieve the extraordinary advances it has made.

## 5.3.

### Genomics and proteomics

The term genomics refers to the biotechnology discipline which deals with decoding the human genome. Admittedly, the genes are characteristic and necessary for what occurs in a cell, but proteins are the actual carriers of the cell functions defined by the genes.

Consequently, in order to understand the cellular processes, it is necessary to record and analyze the totality of the existing proteins with their numerous interactions (proteome research, proteomics). The most important instruments of proteomics include the analysis of protein expression, i.e. the number, size and chemical composition of the proteins in a cell. Processes such as gel electrophoresis, mass spectrometry, X-ray diffraction and nuclear magnetic resonance play an important role in this case. MicroFluidics also makes a substantial contribution to technological further development in this sector.



PCR sequence

## 5.4.

### Biochip analysis

This process is a highly innovative method of molecular-biological analysis. Biochips are microscopic slides (generally made of glass or a polymer) on which biological material (e.g. oligonucleotides or cDNA) is fixed in large numbers and with a high density, as a single strand in the form of microarrays, i.e. in a defined arrangement. Such chips perform the function of diagnostic tools and are offered by many manufacturers with various attributes.

In the case of biochip analysis, the substance under investigation, which is also single-strand (e.g. a patient sample or food sample), is brought into contact with the molecules fixed on the chip. If molecules of the sample substance encounter molecules of the chip that are complementary to them, an interaction occurs on the basis of the key-and-lock principle ("precise-fitting parts"), with formation of a double strand (complementary hybridization). This process can be visualized by appropriate pretreatment of



the sample materials with fluorescent dye. The fluorescent pattern thus produced contains the information as to what substances were present in the sample.

Biochip technology has recently also been applied to detection of protein patterns (protein chips) in the cell. In this case, the chips utilize the specific bonding of proteins to interaction partners positioned on the chip. However, this technology is still in its infancy.

#### Specific MicroFluidic tasks and requirements

- Highly precise dosing on the chip
- Minimal channel dimensioning, freedom from dead volumes and perfect flushability
- Effective hybridization with minimized consumption of costly reagents
- Transportation of the reagents by pressurization through inert gas (to replace pumps)
- Pressure control specific to the various pathways in one unit
- Flow monitoring upstream of the chip

## 5.5.

### Genetic engineering in medicine

Genetic engineering serves as a tool in the production of new medicines and in simple and fast diagnosis of hereditary and infectious diseases. One current example relates to the progress being made in research into the causes of diabetes. A gene has been identified which plays a demonstrable role in the occurrence of type 2 diabetes. On the basis of this, a method has been developed to predict the genetic risk of type 2 diabetes.

One other example relates to production of the substance erythropoietin (EPO), required for formation of red corpuscles and which is produced by the kidney. EPO is present only in extremely small traces in the human body, and that is precisely why genetic engineering methods offer the only possibility of producing this substance in larger quantities and making it available for therapeutic purposes.

A person's predisposition to developing cancer may be of genetic origin and thus hereditary. In this case, genetic engineering can be used to identify genes whose detection would anticipate a probability to develop breast cancer. Here as well, initial approaches have been achieved for corresponding therapeutic options.

### **5.6.** **Production of proteins**

A wide range of pharmaceutically active proteins are produced today with the aid of genetically modified microorganisms on the basis of the process steps described above (in refined or extended form). One known example of this is the production of human insulin and the faster-acting analog insulins.

### **5.7.** **Production of DNA vaccines**

DNA vaccines offer clear advantages over conventional vaccines (on the basis of inactivated viruses or bacteria). They are tolerated better, cost less to produce and are easier to store. This involves injecting DNA molecules bearing the hereditary information for specific antigens of the pathogen. Today, DNA vaccines are produced on a 10-100 gram scale.

**Type 7604**  
**Possible application**  
**for continuous**  
**circulation of the**  
**sample medium in**  
**biochip hybridization**



## 6. Our daily bread

MicroFluidics in foodstuff monitoring and inspection



### **6.1. Transborder analysis**

With the increasing practice of intensive farming and innovations in the field of food technology, i.e. genetically modified products and designer foodstuffs, our shopping basket is filled with products that are being more and more critically viewed by the consumer. “Designer” foods, which are intended to enhance well-being and prevent illnesses due to their specific constituents (minerals, vitamins and antioxidants) and other “innovations” increase the distrust of purchasers, who demand that their food contains absolutely no constituents posing a health hazard. Consequently, effective food-

stuff monitoring and inspection are indispensable, not only for consumer protection, but also for manufacturers who endeavor to gain the trust of consumers. In 1999, the European Commission, in a white book, proposed a monitoring mechanism covering the entire chain of production, enabling full-coverage quality analysis from fodder or raw material up to the end product.

This requires powerful examination methods that also have to be harmonized and validated in view of transborder production chains.

Since foodstuffs generally comprise heterogeneous substance mixtures, the individual substances must be analyzed qualitatively and quantitatively, thus placing stringent demands on sample preparation and analysis.

One example is systematic traceability of the origin of meat products. In this case, a digital identity pattern of each animal is created with the aid of genetic analysis; this pattern allows subsequent and clear traceability of the origin of the meat processed. Besides sophisticated sample logistics, this also necessitates highly automated analysis robots with a high sample throughput, which are then integrated into a powerful database concept.

The aim of the checks and inspections is a precise knowledge of the composition of the foodstuff, thus requiring highly sensitive methods of food chemistry and food analysis with respect to the maximum values of contaminants, of which there are generally only traces. The diversity of foodstuffs corresponds to the variability of the methods used, ranging up to mass spectrometry and, recently, also DNA analysis. One special task results from the matrix dependence that frequently occurs in the case of food analysis. Special sample conditioning methods can help in this case, e.g. immunochemical sample conditioning methods.

#### Specific MicroFluidic tasks and requirements

- Foodstuff inspection: conveying, mixing, distributing and dosing
- Equipment engineering: flushability, operational reliability and purity of the materials used etc.
- Detection of flow rates allows documentation, traceability and quality assurance



**Type 6606**  
Diaphragm valve with  
excellent flushability  
for use in food science

# 7. Bürkert MicroFluidics

The products

## General product characteristics

The entire MicroFluidic product series meets the special requirements of the fields of application for which it was designed in line with demand. This means:

- Small and standardized installation dimensions for setting up very compact systems
- Small nominal diameters of the fluid channels with a view to the frequently costly media only available in small quantities
- Use of corrosion-resistant materials
- Minimal dead volumes and easy flushability
- Separation between medium and actuator
- Suitability for battery operation

## Product range

The MicroFluidic product range covers the following segments:

- Valves of diverse designs
- Gas mass flow controllers
- Liquid and gas mass flow meters
- Micro-pumps
- Manifolds
- Sensors for pressure and temperature
- Electronics.

In addition to the information provided on the following pages, information can also be found in the Bürkert Product Catalog, which we will be glad to send to you upon request.

**Note:** The sensors for pressure and temperature are procured from relevant manufacturers in accordance with the specified conditions of the application.

## Product systematics

The Bürkert MicroFluidic product range is subdivided as follows:

### Components

Serial produced valves, pumps, sensors for pressure and temperature, mass flow meters and mass flow controllers



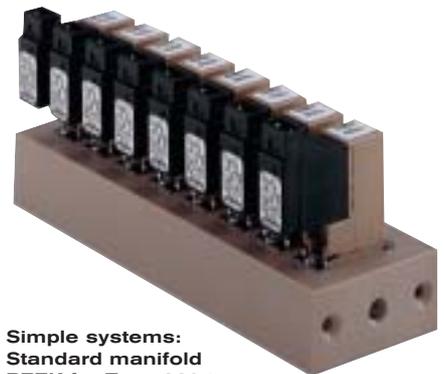
**Individual components:**  
Type 6604

## Manifolds and systems

In various designs (standardized and application-specific), for implementing frequently recurring functions, such as dosing and mixing or distribution.

### Highly integrated systems

Extensive systems with high functionality bandwidth due to the individual combination of specific manifolds, components, electronics and sensor systems.



**Simple systems:**  
Standard manifold  
PEEK for Type 6604



**Highly integrated systems:**  
Gas flow control unit

## 8. Valves

### What valve meets your needs?

Over 20 different valve types are available. They cover a very broad range of application and thus offer the best possible solution for any specific application task. The valves differ in regards to the following features (see table):

- Circuit function: 2/2-way, 3/2-way and 2/2-way proportional
- With or without media separation
- For neutral, slightly aggressive or aggressive media
- For differing orifices (from 0.4 to 8 mm)
- For different pressure ranges (from vacuum to 10 bar).

### Special characteristics

- Very low internal volumes down to 5  $\mu\text{l}$
- Low dead volumes, frequently absolutely no dead volumes
- Easily flushable inner contours
- High back-pressure tightness
- Suitable for vacuum operation
- Virtually no heat transfer between medium and actuator
- Medium temperature up to 50 or 70 °C
- Frequently very low power requirement and, thus, particular suitability for battery operation.

### Flipper valves

On flipper valves, a metal tongue (flipper) vulcanized with elastomer opens and closes the valve seat. Flipper valves allow complete separation of medium and actuator. Compared to diaphragm valves, they produce no pressure pulse when switching and are thus particularly suitable for precise dosing. Less force is required to move the flipper than with the diaphragm valve. Consequently, the response times are particularly short and reproducible.

Flipper valves are insensitive to pressure surges and also withstand higher burst pressures. Since the medium does not act on large areas, the valves remain tight even if a high pressure (back pressure) is applied against the flow direction. Flipper valves have a very long service life owing to the lack of diaphragm flexing.

#### A flipper valve

- operates virtually free of pulsation
- has particularly short and reproducible response times
- requires very little power
- has a very long service life
- is insensitive to pressure peaks
- is also suitable for highly aggressive media.

### Rocker valves

Rocker valves have a diaphragm coupled by a switching rocker that opens or closes two valve seats. They are thus pressure-compensated, in the same way as flipper valves, and also reliably seal off pressures that are applied against the flow direction. The two valve seats and the connecting fluid channel lie in one plane. This means that rocker valves have an extremely small internal volume and virtually no dead volume. The geometry of the fluid channels is designed so that gaps are not produced and to guarantee very good flushability. During switchover, a cross flow is produced between the two valve seats, thus preventing deposits in the valve chamber. If extremely stringent requirements apply to internal volume and dead volume, it is also possible to mount rocker valves directly on user-specific manifolds. The valve seats are then integrated in the manifolds.

#### A rocker valve

- can be flushed particularly well
- can be flange-mounted directly onto a manifold
- has a minimum internal volume and extremely low dead volumes
- remains tight even in the case of pressure opposite to the flow direction (back pressure)
- is also suitable for highly aggressive media.

**Bürkert MicroFluidics valves**

	Circuit function			Med. sep.	Medium				Pressure range (bar)				Orifice (mm)			Miscellaneous	
	2/2-way	3/2-way	Proportional	Yes	Neutral	Slightly aggressive	Aggressive	Suitable for vacuum	up to 2 bar	up to 3 bar	up to 10 bar	> 10 bar	< 0.6	0.8 bis 2	2 bis 6	Bistable version	kv value min./max. [m <sup>3</sup> /h]
Type 0124	▪	▪		▪	▪	▪	▪								up to 5	▪	0.25/0.4
Type 0127	▪	▪		▪	▪	▪	▪							▪			0.025/0.06
Type 6604	▪	▪		▪	▪	▪	▪						▪				0.0074
Type 6606	▪	▪		▪	▪	▪	▪							▪			0.025/0.06
Type 6608	▪	▪		▪	▪	▪	▪								up to 3		0.1/0.18
Type 0330	▪	▪		▪	▪	▪			up to 16						▪	▪	0.11/0.29
Type 0331		▪		▪	▪	▪			up to 16						▪		0.08/0.28
Type 6124	▪	▪		▪	▪	▪							▪				0.0074
Type 6126	▪	▪		▪	▪	▪								▪			0.01
Type 6128	▪	▪		▪	▪	▪									▪		0.11/0.18
Type 6104		▪			▪				up to 7				▪				Q <sub>Nn</sub> max 8.5 l/min
Type 6106		▪			▪									▪		▪	Q <sub>Nn</sub> max 40 l/min
Type 0121	▪	▪		▪	▪	▪	▪		up to 4						up to 8	▪	0.3/1.0
Type 0330 A	▪	▪		▪	▪	▪									▪	▪	0.11/0.29
Type 6011 A	▪				▪	▪			up to 21					▪			0.045/0.13
Type 6013 A	▪				▪	▪			up to 25						▪		0.12/0.55
Type 2822			▪		▪								0.3-1				0.002/0.02
Type 6021			▪		▪				up to 6						▪		0.05
Type 6022			▪		▪				up to 8						▪		0.1/0.33
Type 6023			▪		▪				up to 4					▪			0.4/0.7



#### **Type 6604**

**Direct-acting flipper solenoid valve** with media separation as 2/2-way or 3/2-way valve with monostable or bistable (pulse) switching function. Fast-switching. Very low power consumption and thus very suitable for battery operation. Minimum dead volume and low-gap inner contour. Width per station: 11 mm. Used primarily for very small quantities of aggressive media.



#### **Type 6606**

**Direct-acting rocker solenoid valve** with isolating diaphragm as 2/2-way or 3/2-way valve. With minimum dead volume and low-gap and thus easy-to-flush inner contour. High-quality materials guarantee extreme chemical resistance. The medium only comes into contact with the body and FFKM seal. Coil can be changed easily without having to open the body.



#### **Type 6608**

**Direct-acting rocker solenoid valve with medium-separated PEEK body** with dead volume-optimized and easy-to-flush inner contour. Central screw fitting of the coil allows the coil to be exchanged with the medium applied. Modular body design allows the use of various fluidic connections. For pressures up to 10 bar and orifice DN 3 mm.



**Bürkert MicroFluidics valves**

	Connectors								Body material										Sealing material						
	Flange	G thread	NPT thread	UNF 1/4-28 thread	M5 thread	Tube spigot	Cable plug	Rectangular plug	Leads	SS	Brass	PPS	PA	PVC	PP	PTFE	ETFE	PEEK	PCTFE	PVDF	FFKM	FPM	EPDM	PFTE	NBR
Type 0124		▪	▪				▪								▪						▪	▪	▪		▪
Type 0127	▪	▪	▪	▪		▪	▪	▪									▪	▪	▪	▪	▪	▪	▪		
Type 6604	▪			▪		▪	▪	▪										▪	▪	▪	▪	▪	▪		
Type 6606	▪	▪	▪	▪		▪	▪	▪									▪	▪	▪	▪	▪	▪	▪		
Type 6608	▪	▪	▪				▪	▪										▪	▪	▪	▪	▪	▪		
Type 0330		▪	▪				▪		▪	▪												▪	▪		▪
Type 0331	▪	▪					▪		▪	▪												▪	▪		▪
Type 6124	▪			▪		▪	▪	▪										▪				▪	▪		
Type 6126	▪				▪	▪	▪	▪				▪										▪	▪		
Type 6128	▪	▪	▪				▪	▪	▪			▪										▪	▪		
Type 6104	▪						▪	▪				▪										▪	▪		
Type 6106	▪						▪	▪	▪			▪										▪	▪		
Type 0121		▪	▪				▪		▪				▪	▪	▪					▪	▪	▪	▪		
Type 0330 A		▪	▪				▪		▪													▪	▪		
Type 6011 A	▪	▪					▪		▪	▪												▪	▪		
Type 6013 A	▪	▪					▪		▪	▪												▪	▪		
Type 2822	▪	▪	▪				▪	▪	▪	▪												▪	▪		
Type 6021		▪	▪				▪		▪	▪												▪	▪		
Type 6022		▪	▪				▪		▪	▪												▪	▪		
Type 6023		▪	▪				▪		▪	▪												▪	▪		

**Type 121**

**Direct-acting 2/2-way or 3/2-way pivoted-armature solenoid valve**

The system for separating solenoid system and media chamber consists of an intermediate-vented isolating diaphragm system. Diverse range of applications due to the material selection, including applications involving very aggressive media.





#### **Type 124**

**Direct-acting 2/2-way or 3/2-way pivoted-armature solenoid valve** with isolating diaphragm between solenoid system and media chamber. Available in various materials and with diverse modes of action for opening, shutting off, dosing, aeration or distribution. Particularly used for aggressive, abrasive or lightly contaminated media.



#### **Type 127**

**Direct-acting rocker solenoid valve** with media separation. Developed as minimum-volume, low-dead-volume and easy-to-flush valve (CIP-enabled). An extensive range of bodies allows numerous connection options. All fluidic connections are located in one plane.



#### **Types 330/331**

**Direct-acting 2/2-way or 2/3-way pivoted-armature solenoid valves**  
Solenoid system and media chamber are separated by an isolating diaphragm system. The valves are fast-switching and feature a very long service life, even in the case of dry operation. They are particularly suitable for abrasive and lightly contaminated media. Valve 331 is designed specifically for flange mounting.



**Types 6104/6106**

**Direct-acting 3/2-way rocker solenoid valves**

without media separation, low power consumption, monostable and bistable drive, body made of polyamide. Suitable only for gases.



**Type 6011A and 6013A**

**Direct-acting plunger-type valves**

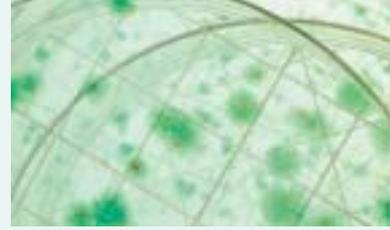
for high cycling rates (up to 1,000 operations per minutes), production under clean room conditions, including thorough cleaning of all wetted parts to remove inorganic and organic substances. Production and inspection are designed specifically for seat tightness and minimal leakage to the outside (permitted leakage rate  $10^{-4}$  mbarl/sec).



**Types 6021/22/23**

**Direct-acting proportional valves**

for continuous closed-loop flow-rate control. Low hysteresis, high reproducibility. Also suitable for technical vacuum in the outlet channel.



#### **Type 6124**

##### **Direct-acting flipper solenoid valve, 2/2-way or 3/2-way**

with media separation. With monostable or bistable (pulse) switching function. Pulse switching with only 20 ms pulse length and extremely low energy demand, consequently particularly suitable for battery operation. Minimum dead volume and easy-to-flush inner contour. Materials used: FPM, EPDM, PEEK. Use for very small quantities of neutral or mildly aggressive gases and liquids.



#### **Type 6126**

##### **Direct-acting rocker solenoid valve, 2/2-way or 3/2-way**

A diaphragm separates the medium from the actuator. In addition, the coil and actuator are separated by means of a stainless steel plate. Universal use for applications involving switching small quantities of compressed air or lightly contaminated fluids.

#### **Type 6128**

##### **Rocker solenoid actuator with medium-separated PPS body**

with dead volume optimized and easy-to-flush inner contour. Central screw fixture of the coil allows the coil to be exchanged even with the medium applied. Modular body design allows the use of various fluidic connections. Type 6128 can be used universally for applications on which compressed air, vacuum or lightly contaminated or slightly aggressive gases and liquids are to be switched.



## 9. Pumps, flow meters and manifolds



### Type 7604 Micro-pump

This micro-pump operates based on the principle of a self-priming diaphragm pump. It was specifically developed for continuous pumping of small quantities of aggressive, inorganic or biological media. Highly precise dosing is possible in combination with an additional flow sensor.

The wetted parts are made of inert and resistant material: PEEK (body) and FFKM (seals). Owing to its resistance, the material combination can also be used for very aggressive media. The temperature of the medium can range from +10 to +60 °C.

The width per station of 11 mm allows an extremely compact set-up, allowing any choice of pump installation position. Owing to the low power consumption of only 1.5 Watt, the pump is also suitable for use in battery-operated equipment.

The delivery rate (max. rate 5 ml/min) can be adapted to the task by varying the control frequency. Virtually pulsation-free dosing is achieved owing to the high maximum permitted frequency (40 Hz).

### Pumps

Micro-pump	Type 7604	Variable control frequency (max. 40 Hz) Virtually pulsation-free dosing Power consumption only 1.5 Watt
Micro-dosing unit	Type 7616	Dosing unit comprising 3 valves and a manifold



### **Type 7616**

#### **Micro-dosing unit**

The self-priming, low-dead-volume micro-dosing unit consists of:

- two valves, Type 6604
- one valve 6606
- one manifold (minimized with a view to the internal volume)
- a control circuitry unit (option).

The delivery rate can be adjusted via the number of cycles (max. 650 cycles/min.) and the optionally adjustable stroke volume (0.5  $\mu$ l ... 5  $\mu$ l). Thanks to the high reproducibility, the unit is suitable for the precise dosing of ultra-small fluid quantities. PEEK and FFKM as the sole wetted materials virtually predestine the unit for use in aggressive media.

### **Type 8005**

#### **Liquid mass flow meter**

The mass flow meter 8005 allows fast and precise flow-rate measurements for fluids down to the nanoliter range. Highly sensitive, intelligent CMOSens microchips are capable of detecting the mass flow rate bi-directionally and with media separation through a thin PEEK wall. The high dynamics of this measurement principle allows a measuring range of 5 to 1,500  $\mu$ l/min. The accuracy is better than 1.5 or 3 %, respectively, depending on the measuring range, and the detection limit is approx. 150 nanoliters/min.

The device is free of dead volume and its response time, at 20 ms (lower limit), is extraordinarily short. The mounting dimension is 14 mm and the type of protection is IP 65.

The mass flow meter 8005 can be interconnected with other components to form functional modules such as:

- with valve 6604 and the micro-pump 7604 to produce a dosing unit and
- with valve 6604 and the proportional valve 2822 to form a mass flow controller.



### Type 8700

#### Mass flow meter

Low-cost version. Measuring range 0.05 ... 30 NI/min. (N<sub>2</sub>). Digital circuitry with analog or digital interface (RS232/RS485). Sensor arranged in bypass, medium-separated, particularly suitable for toxic or aggressive gases. Accuracy  $\pm 1.5\%$  of measured value,  $\pm 0.5\%$  of full scale value, option for calibration with N<sub>2</sub> and conversion factor. Fluidic body made of SS, cover made of chromium-plated sheet steel.



### Type 8702

#### Mass flow meter

Measuring range 0.02 ... 50 NI/min. (N<sub>2</sub>); digital circuitry with analog or digital interface (RS232/RS485) and optional field bus interface. Sensor (semiconductor element) arranged in bypass in medium. Accuracy  $\pm 1\%$  of measured value and  $\pm 0.5\%$  of full-scale value. Fluidic body made of SS, cover made of PPS (IP65). Galvanic isolation of inputs and outputs.

#### Mass flow meters

Liquid mass flow meter	Type 8005	For ultra-small flow rates, max. 1.5 ml/min. Medium-separated, digital interface
Gas flow meter	Type 8700	Measuring range (air) 0.05 ... 30 NI/min. Medium-separated sensor
Gas flow meter	Type 8702	Measuring range (air) 0.02 ... 50 NI/min. Sensor in medium, optional field bus interface

#### Mass flow controllers

Gas mass flow controller	Type 8710	Measuring range (air) 0.05 ... 30 NI/min. Medium-separated sensor
Gas mass flow controller	Type 8712	Measuring range (air) 0.02 ... 50 NI/min. Sensor in medium, settling time < 300 ms



### Manifolds

In addition to customer-specific manifolds, Bürkert offers a selection of standard solutions for Types 6604, 6124, 6126 and 6606. Mixer or distributor functions (2-way) can be implemented via a common collecting channel in each case. The various solutions jointly feature PEEK as the base material and UNF 1/4-28 as fluidic connections, whereby the individual outlet and inlet ports are each in one plane.

Manifolds		
For Types 6604 and 6124 11 mm mounting dimension	2 – 10-fold Standard	All ports in one plane
For Types 6606 and 6126 16 mm mounting dimension	2 – 8-fold Standard	All ports in one plane
For Types 6606 and 6126 16 mm mounting dimension	2 – 8-fold High-end	With extremely low dead volume, direct flange mounting of the valve with seat contour in the manifold, particularly suitable for vacuum, common port at side, individual ports pointing downwards

# MANIFOLD

# 10. Always a unique product

The application-specific solution from Bürkert

## Every problem is different

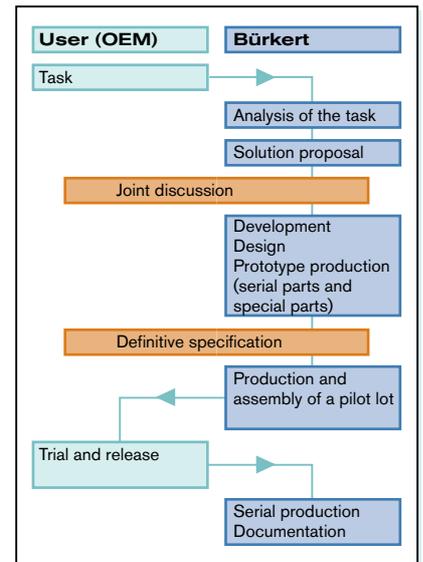
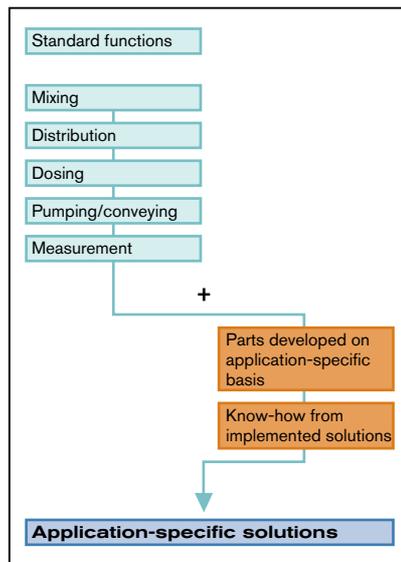
The variety of available components, systems and manifolds from Bürkert is founded on the many decades of experience with products required by the market. This forms the basis for meeting many of the tasks required. In some individual cases, they may not suffice to configure the optimum solution for a specific task.

Individual, application-specific solutions form our concept for technically implementing special requirements with a coordinated performance and service mix. Such solutions frequently comprise systems for standard functions, such as mixing or dosing, combined with specially developed and manufactured components. In most cases, these are individually designed and manufactured connection elements.

resources required for your task in close cooperation with you, the customer.

## System partnership

You know what you want. We know what we can do. Close cooperation and continual exchange of information is the course we pursue. Regardless of whether you are an OEM partner who manufactures their own equipment or a "normal" user, in Bürkert, you have



Complete functional unit from a single source: water supply for dentists' chairs

## Every Bürkert solution is systematic

When developing new, customer-specific solutions, it goes without saying that Bürkert draws on its comprehensive development know-how, modern production technology, application knowledge and experience from already implemented solutions. But this also involves another factor: the incentive and commitment to utilize all the

chosen the right place for achieving a fast and efficient solution. Generally, it takes only a few weeks to solve the problem.

## The project – step by step

The leitmotif of this cooperation: The combination of specific user knowledge with the know-how of an experienced equipment manufacturer.



MicroFluidics production facility in Gerabronn, Germany

## Enough of theory, let's talk about practice!

### One plant – one promise

Bürkert has set up a production facility specifically for MicroFluidic products where development and production of application-specific systems and serial production of standard components are organized under one roof.

A specially trained, experienced staff, equipped with the know-how acquired over many years, fully concentrates on offering a solution geared towards the needs of the future.

### Highest priority: quality

MicroFluidic components would be inconceivable without an optimum standard for production quality. Clean room conditions are just as much a part of this as strict compliance with permitted tolerances. And settings accurate to the  $\mu\text{m}$  are an entirely normal step in the production process.

### State-of-the-art development workstations

One example of many: a special software package allows the user's only sketchily documented task to be immediately converted, and visualized, to a solution proposal with a 3-D representation. After review, revision and approval by the customer, the same development tool is used to produce a fully dimensioned production drawing.

### Tested and approved part for part

Assembly and inspection are performed individually for each part and in directly consecutive operations.

Inspection programs control the sequence and record the values.

In special cases, it is possible to permanently assign the test results to a part and document them. This is one important aspect, e.g., for validation processes.



Use of 3-D CAD systems



Maximum precision in assembly



Assembly and inspection workstation

# 11. Bürkert MicroFluidics: beyond all expectations

Example applications

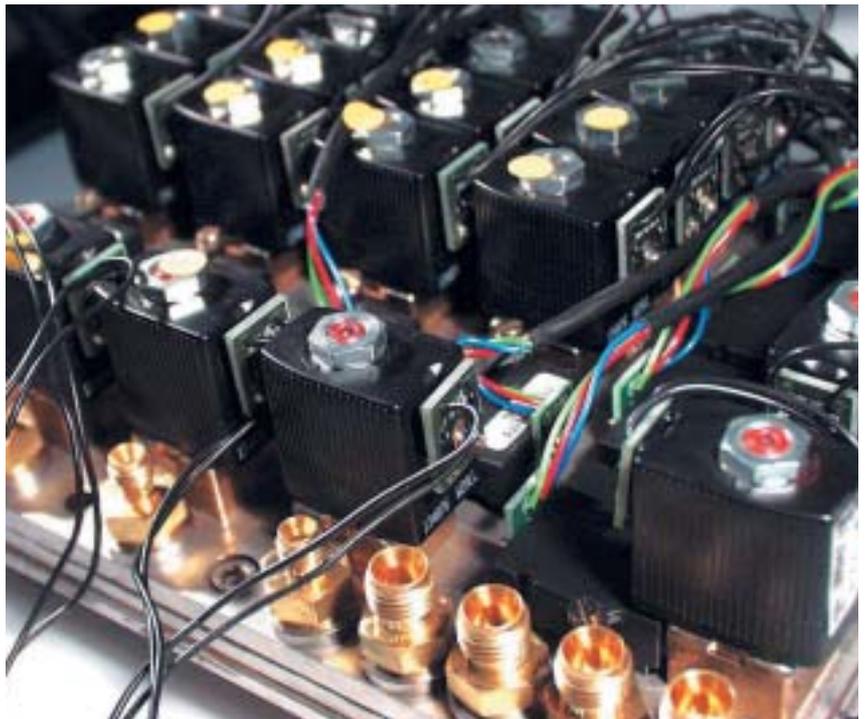
Provided there are no laws of nature against it, you can expect Bürkert to come up with virtually any possible solution to your specific application problem. That also means that if we bring up what appears to be an unrealizable idea, do not hesitate to ask us about it. There is always a way, especially with the competence of a technology leader!

## 11.1.

### **Application: “gas mixing for small reactors”**

#### **Field of application**

The system serves to simultaneously supply several items of consumption equipment with individually composed gas mixtures from several clean gases or parent gases. It unites all the functions of gas ducting and pressure and flow-rate detection into one compact block. It is used e.g. for calibration units or supplying small reactors of all types.



Proportional valves, Type 6021, in the system

**Mechanical construction**

The system consists of a manifold which is designed on an application-specific basis and incorporates multi-layer technology with 3 layers and various standard components mounted on this. The illustrations below show an example for 3 gases and 3 consumption units.

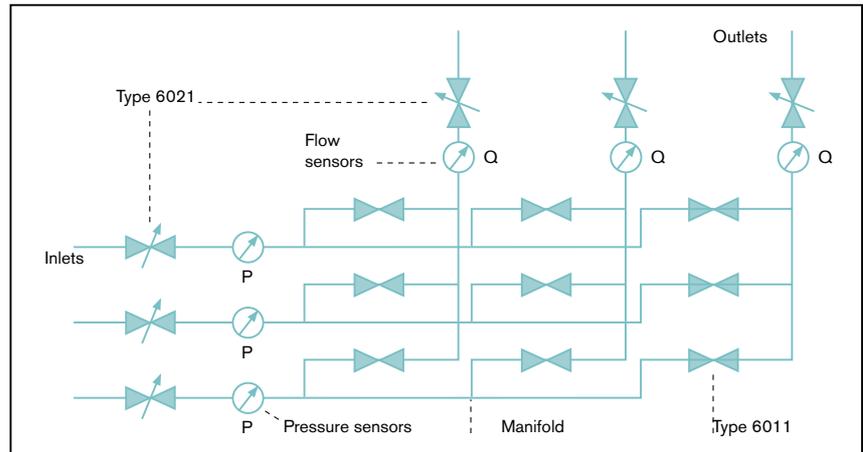
**Characteristics**

- Several gas inlets for unmixed gases, e.g. N<sub>2</sub>, O<sub>2</sub> or air
- Several gas outlets for mixed gases, each with individual mixing ratios, for simultaneous supply to several consumption units
- Admixing by abrasion-free on/off valves (standard valves 6011)
- Proportional valves 6021 at inlet and outlet
- Mass flow rate controllable in the range 0.5 to approx. 50 NI/h

- Detection of the input pressure of the gases
- Detection of the mass flow rate of gas inlets and gas outlets
- Low dead volumes, compact design and low-wear technology
- Short response times for changing the mixing ratio by entrainment-free

- ducting; mixing is already performed on the manifold by a reservoir between the layers
- Less line-ducting, hose-connection and connection-system complexity due to multilayer manifold.

**Gas mixing for small reactors**



Description/Function	Quantity	Typical characteristic parameters
Manifolds with gas ducting	1	Multilayer technology (3 layers)
Proport. valve, Type 6021, for gas inlet and outlet	6	Low hysteresis for continuous closed-loop flow-rate control
Stand. valve, Type 6011, for internal gas admixing	9	for extremely high cycling rate particularly low leakage rate
Pressure sensors	3	Application-specific
Flow sensors	3	Application-specific



### 11.2.

#### Application: “low-dead volume mixer/distributor”

##### Field of application

In many applications in medicine or biotechnology, it is necessary to feed reaction vessels or measuring chambers with differing media or defined media mixtures following a stipulated time sequence. Switchover between the individual feeding stages should occur without any entrainment effects and flushing operations should be possible.

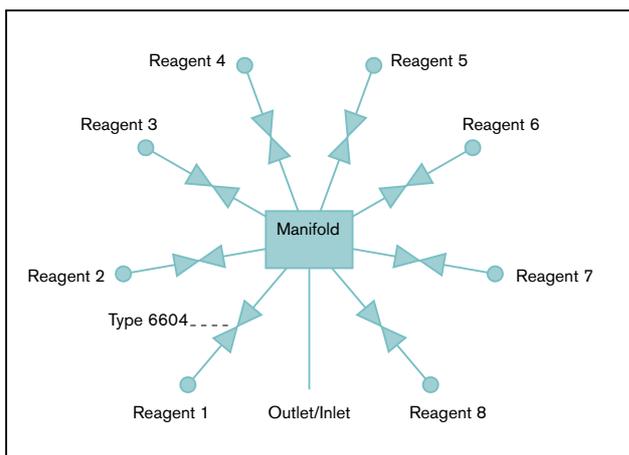
##### Mechanical construction

8 valves for mixing and distribution are arranged in a radial pattern, with uni-

formly small connecting channels to the outlet around a central manifold.

##### Characteristics

- Suitable for neutral and aggressive gases and fluids
- Pressure range from vacuum to 3 bar
- Operating temperature 0 to +50 °C
- Operating voltage 24 V DC
- Minimum channel length the same at all points
- Internal volume (per channel) 5 µl
- UNF 10-32 connections.



**Fluidic schematic  
Low-dead volume  
mixer/distributor**

Description/Function	Quantity	Typical characteristic parameters
Manifold	1	Material acrylic or PEEK, 5µl internal volume per channel
Valve, Type 6604	8	Fast switching flipper valve without pump effect when opening or closing

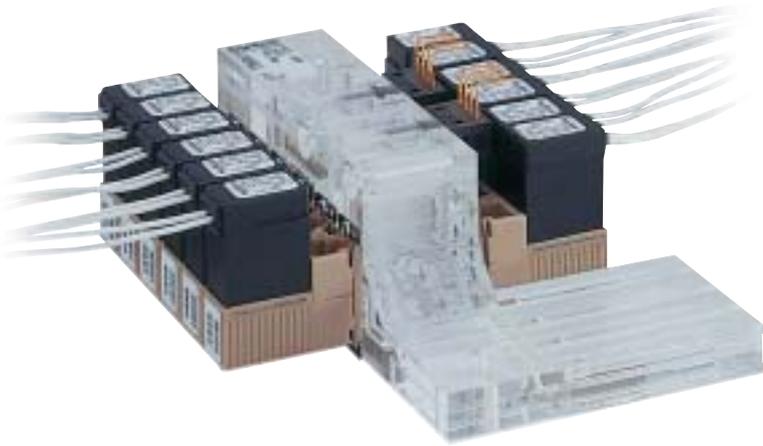
### 11.3.

#### Application: “pressure supply, AutoPatch unit”

##### Field of application

The system serves to provide a defined vacuum for an AutoPatch unit. This allows a significant increase in efficiency in comparison to conventional patch clamp technology in terms of receptor research and medicine development. The electrical signals that are converted by ion channels featuring a receptor function from chemical signals (ligands) are used for this since numerous diseases are attributable to disturbances of ion channels.

The AutoPatch unit isolates a single cell from the cell suspension under examination via an applied vacuum through a channel the size of a few micrometers, passes this over an electrode and thus establishes an electrical connection to the inside of the cell.



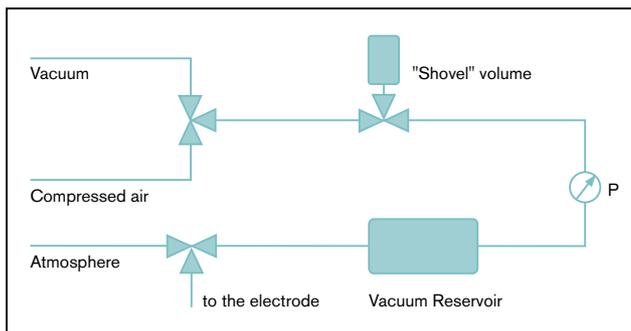
### Mechanical construction

The three-layer, diffusion-bonded, multilayer base made of acrylic material guarantees an extremely compact construction. A precisely dosed vacuum or compressed air can be applied to 4 channels per unit. All 4 channels are supplied centrally via one vacuum, with compressed air and atmosphere connection in each case. Vacuum is

“shoveled”, as it were, into an integrated chamber via an interconnected 3/2-way valve, whereby this operation is controlled via a pressure sensor in the chamber. If the required vacuum has been built up, the outlet channel is connected, thus exerting a controlled suction pressure via the electrode onto the medium under investigation.

### Characteristics

- Highly integrated functionality in a very compact space thanks to three-layer, diffusion-bonded manifolds
- 36 mm overall width allows butt mounting with micro-titer base center spacing (4 x 9 mm)
- Build-up of precisely defined vacuum is possible with integrated pressure sensor
- Production process (diffusion bonding) ensures dead volume-optimized channel ducting, no entrainment and no swarf or chips in the fluidic chambers
- Transparency allows visual inspection of channels for impurities
- No additional sealing materials required on/in the manifold since sealing is performed by the material connection (no parting lines).



**Function schematic for a channel for pressure supply, AutoPatch unit**

Description/Function	Quantity	Typical characteristic parameters
Manifold with gas ducting	1	three-layer PMMA multilayer base (diffusion bonded), 4 separate channel ducts
Flipper valve, Type 6124	12	fast-switching 3/2-way valve wetted materials: PEEK and FPM
Pressure sensor	4	uncalibrated, non-amplified $\pm 1$ bar, repeat accuracy $\pm 0.15$ % Span

## 11.4.

### **Application: “dosing head for liquids”**

#### **Field of application**

The “dosing head” system is used e.g. for filling stations or dispensers. On account of its compact design without traditional piping, the manner in which it performs dosing without switchover operations for filling and its optional quantity determination function using a flow sensor (mass flow rate which can be documented), it forms an interesting alternative to traditional syringe pumps. Diverse utilization of the principle in automated devices of all types, e.g. for filling micro-titer plates.

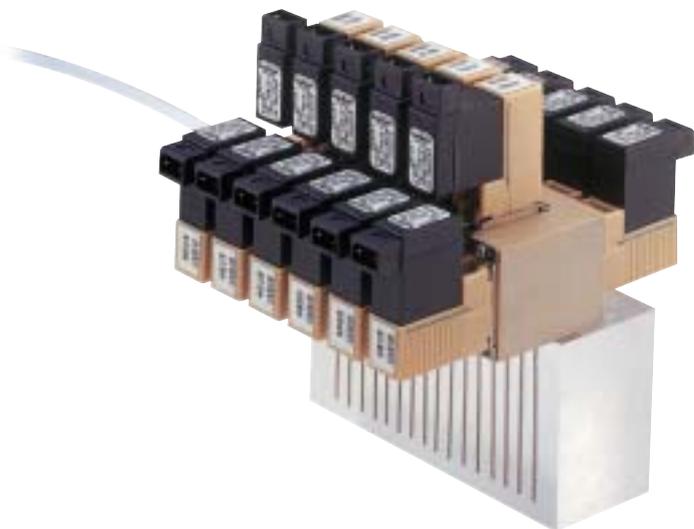
#### **Mechanical construction**

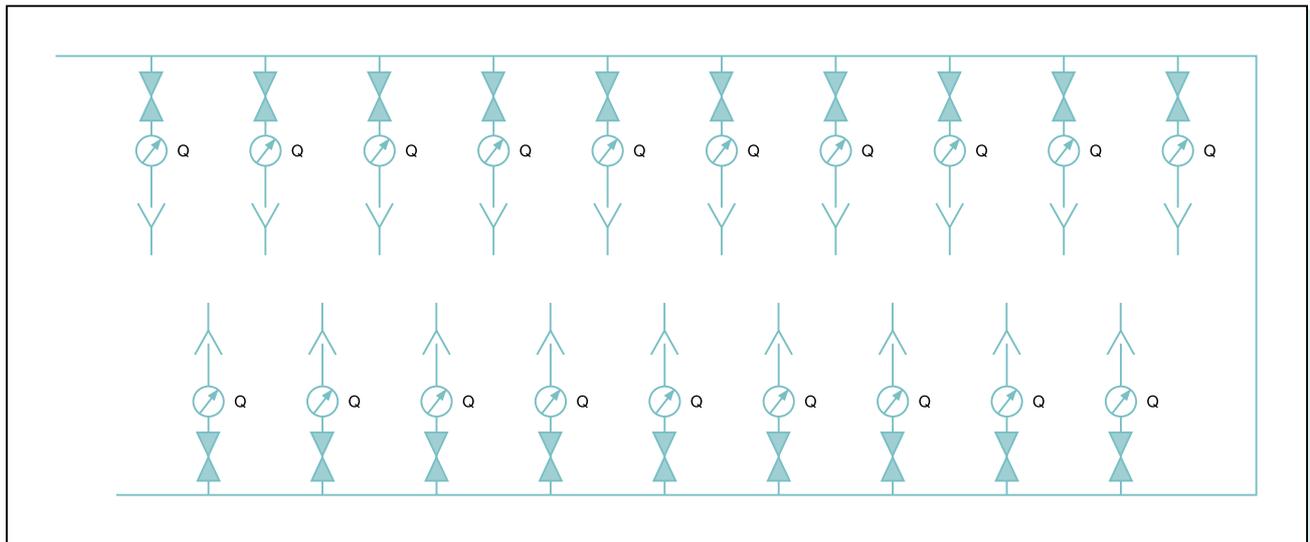
The system consists of a manifold designed on an application-specific basis and Type 6604 valves mounted on it, optionally with Type 8005 flow sen-

sor as well. Alternatively, it is possible to work with micro-pumps, Type 7604, instead of valves, thus eliminating an external pressure supply due to self-priming. Since the design of the nozzle is crucial to precise dosing and for preventing droplet formation at the nozzle, Bürkert offers a wide variety of possible alternatives, ranging from stainless steel cannulas to ceramic nozzles to photolithographically manufactured micro-nozzles with a customer-specific layout.

#### **Characteristics**

- Arrangement (4.5 mm spacing) allows micro-titer plate dosing
- Constant dosing readiness from a pressurized supply vessel
- Without pressure supply vessel if using a micro-pump
- No switchover operation required for filling
- No piping
- Flow rates in the nl/min to ml/min range
- Flushing in-between can be implemented simply
- Optional control loop with flow sensor
- Sensor with media separation
- Dosed quantity can be documented.



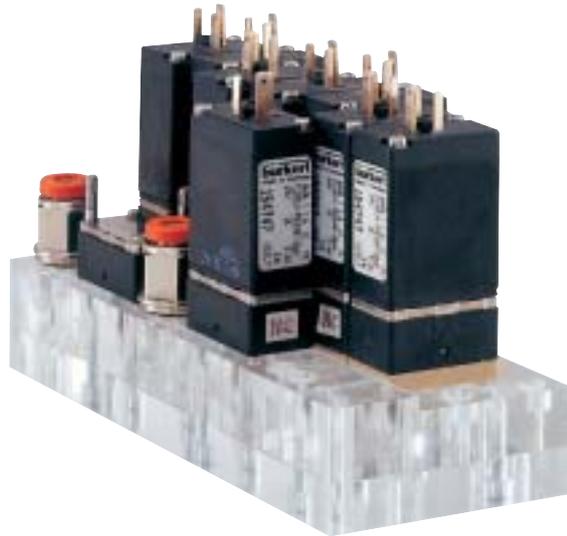


Flowchart for dosing head for 16 channels with flow-rate metering

Description/Function	Quantity	Typical characteristic parameters
Distributor, manifold	1	1 central inlet, 16 outlets lined up with 4.5 mm spacing, material: PEEK
Flipper solenoid valve Type 6604	16	fast-switching, wetted materials: PEEK and FFKM (Simriz), option for PC board contacting
Dosing needles	16	press-in stainless steel cannulas ID 0.3 mm, optionally other nozzle types available
Optional Liquid mass flow meter Type 8005	16	fully media-separated measuring method (only PEEK as wetted material), thermal measuring principle, measuring range down to 1.5 ml/min with dynamics 1:200, smallest detectable quantity 20 nl

## 11.5.

### Application: “determining oil viscosity”



#### Field of application

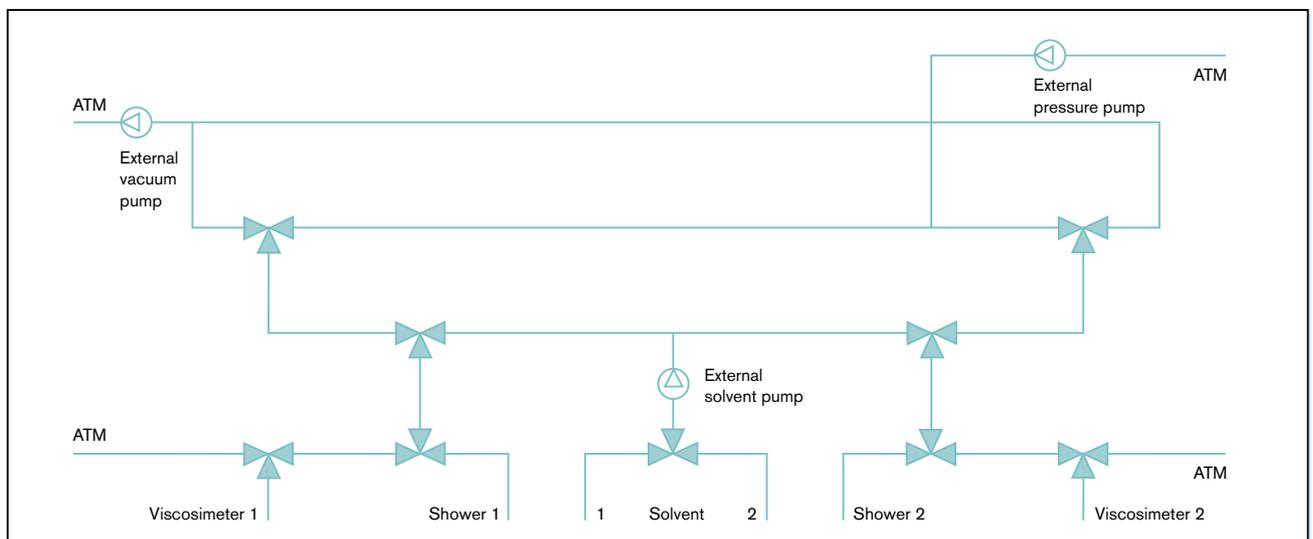
The set-up performs the task of the entire fluid handling for automated viscosity determination of oils. The measuring instrument contains two multi-range viscosimeters. Over those, the Bürkert system applies a certain vacuum to suck in the oil to be analyzed. The sample remains in the viscosimeter until the tempering period has elapsed and then drains off freely. The discharge time is measured thermally by NTC sensors and the viscosity is determined. The Bürkert system is also used for multi-stage capillary cleaning in which even aggressive solvents such as acetone or toluene are used. Up to 20 measurements per hour can be conducted, allowing for the discharge times.

#### Mechanical construction

The system consists of a bolted, two-layer manifold and 9 3/2-way Type 6606 valves mounted on it. The self-draining, dead volume optimized channels are produced by machining at the interface of the bases and are sealed by a fitted FFKM diaphragm. The two separate metering sections can each be supplied with vacuum or gauge pressure. Supply of two solvents is available for automatic cleaning of the viscosimeter.

#### Characteristics

- Less line-ducting, hose-connection or connection-system complexity owing to multilayer manifold
- Optimized, self-draining channel ducting with 90° suspended set-up
- Electrical contacting via plug-in contacts on a PCB.



Function-schematic, fluidic handling for automated viscosity determination of oils

- Fluidic connections in two opposite planes, mounted to the back panel of the viscosimeter which accounts for low hose connection complexity
- Media-separated valves and high-quality, inert materials allow cleaning of the system with aggressive solvents.

Description/Function	Quantity	Typical characteristic parameters
Manifold	1	2-layer multilayer base made of brass with FFKM diaphragm as seal
Solenoid valve, Type 6606	9	Medium-separated 3/2-way valve with device plug connection pointing upwards for direct PCB contacting
Hose connections for pumps, showers, solvents and viscosimeter	8	Press-fitted brass hose connectors



## 11.6. Application: “water treatment”

### Field of application

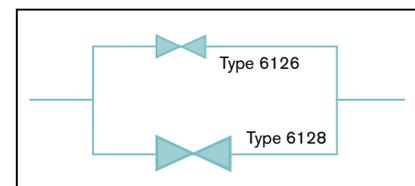
The valve manifold system is a part of a device to produce and dose deionised water. The device is primarily used in genetics and biology laboratories. In this application, it serves as an ultra-pure water source for steam generators, autoclaves and glassware washing machines. The Bürkert system is positioned as a compact unit directly in the dispenser.

### Mechanical construction

One valve, Type 6128, and one valve, Type 6126, are positioned in compact form on a manifold made of acetyl. The two valves are connected in parallel in the manifold providing the user with the option of selecting the right flow rate from three flow rates for their application. Customer-specific cartridge connections are incorporated in the manifold for connection of the inlet and outlet.

### Characteristics

- Compact mechanical construction
- 3 selectable flow rates
- Pre-assembled and pre-tested
- Customer-specific cartridge connections
- Low internal volume
- Low weight.



Parallel connection for 3 flow rates

Description/Function	Quantity	Typical characteristic parameters
Manifold	1	The material is acetyl. Both valves are connected internally in parallel.
2/2-way valve, Type 6128	1	Orifice 3 mm; $k_v$ value 0.18 m <sup>3</sup> /h PPS body and EPDM seal
2/2-way valve, Type 6126	1	Orifice 1.2 mm; $k_v$ value 0.03 m <sup>3</sup> /h PPS body and EPDM seal

## 11.7.

### **Application: “gas flow control unit”**

#### **Field of application**

The gas flow control unit was developed for use in ICP spectral analyzers. This analysis method operates with inductively coupled plasma (ICP) for determining trace elements with spectral lines in the vacuum-ultraviolet spectral band (125 to 190 nm wavelength). The system controls both the plasma flow and the cooling, in addition to two optional gas admixing processes for use of biological media (4 argon controlled systems).

#### **Mechanical construction**

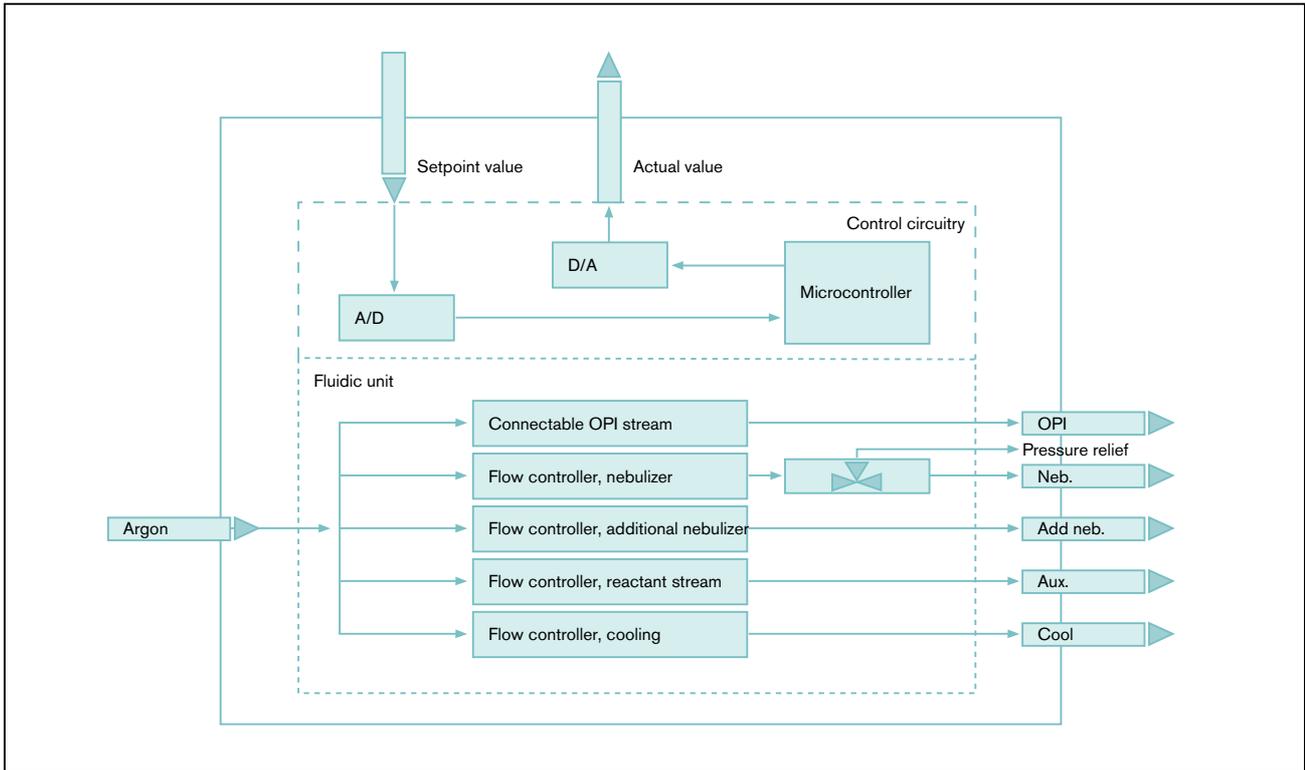
The unit consists of a microprocessor, pressure sensors and pressure switches, solenoid valves for switching the gas and for venting system sections and proportional valves for continuous closed-loop control of gas streams.

All connections of the fluidic components are integrated in an aluminum manifold. The processor kernel serves as the internal central control unit and as an AD-DA converter for set-point and actual values. The implemented firmware can be read or updated via an additional PC interface.

#### **Characteristics**

- Compact overall size (200 x 170 x 100 mm)
- 4 different control ranges for cuvette cooling, for the reactant gas, for the nebulizer and for the additional nebulizer (min. 0 to approx. 1 Sl/min., max. approx. 5 to 20 Sl/min.)
- Optionally intrinsically safe via integrated software and hardware monitoring
- Modularity allows adaptation to various control ranges and numbers of control systems
- Calibration of the sensors in accordance with measuring range and accuracy requirements
- Integrated processor kernel as communication interface and internal central control unit
- PC interface for downloading/updating the firmware
- Control of volume flows instead of mass flow rates.





Schematic, gas flow control unit

Description/Function	Quantity	Typical characteristic parameters
Manifold	1	Aluminium, channel ducting produced by machining, subsequently blocked by spherical plugs for optimum and low-dead volume functionality
Control circuitry	1	4 different controlled systems, autonomously operating system; implemented firmware can be read or updated via PC
Pilot valve, Type 375	3	3/2-way solen. valve for neutral gases, DN 1.2 mm
Proportional valve, Type 2821	1	Direct-acting solenoid proportional valve for neutral gases and fluids; DN 0.8 mm; measuring span 1:25; hysteresis <5 %; high reproducibility and good response sensitivity
Piezo-proportional valve, Type 6115S	3	Direct-act. proport. valve with piezo-ceramic bend. element for oil-free or dry neut. gases; 10 mm width per station, DN 0.4 mm; closed-circuit curr. $\leq 0.1 \mu\text{A}$
Pressure switch, Type 1045	1	Max. 100 bar overpressure safety Setting pressure range 1 ... 10 bar
Pressure switch	2	Max. 10 bar, setting pressure range 0...6 bar Very good control characteristic
Pressure sensors	4	On-chip temperature-compensated and calibrated piezo-resistive pressure sensor, $\pm 0.25$ linearity

## 12. Glossary

### **A**

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#### **Adenine**

Organic purine base, element of nucleotides

#### **Amino acid sequence**

Linear sequence of amino acids in a protein or peptide

#### **Amino acids**

Organic compounds which feature both an amino group and a carboxyl group as the characteristic feature. The 20 "natural" amino acids are joined to the ribosomes of a cell in accordance with the DNA code and form the proteins.

#### **Antibody**

The body's own immune system proteins for fighting exogenous antigens

#### **Anticodon**

Base triplet of tRNA. It serves to translate the genetic code to the ribosomes and reads the mRNA

#### **Antigen**

Foreign protein, which, once it enters the body, causes the formation of antibodies

#### **Array**

Regular two-dimensional arrangement of measuring points on a sensor. In detail: matrices made of biological material, such as DNA or proteins, which is applied in ordered patterns of high density and with equal spacings to a firm substrate. Arrays are used for efficient (parallel) investigation of a high number of biological samples.

#### **Assay**

Test system comprising targets and chemical substances evaluated by a measuring facility

#### **Autologous**

Endogenic or produced within the body

### **B**

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#### **Bacteria**

Unicellular microorganisms without nucleus. They are propagated by simple cell division. 3 basic forms: rod-shaped bacilli, ball-shaped bacilli (cocci) and curved rod-shaped bacilli (spirilla). Bacteria may be desirable (e.g. enteric bacteria or intestinal flora) or undesirable (e.g. salmonella). They are very well-suited to biotechnological production processes since they can be propagated quickly in inexpensive nutrient solutions.

#### **Base pair**

The two bases, adenine and thymine, and the two bases, cytosine and guanine; each form pairs in a DNA double strand, which are held together by weak bonds. The sum of these bonds is responsible for the coherence of the two DNA strands.

#### **Bases**

The conventional designation for the basic elements of nucleotides, the subunits of DNA and RNA. The genetic information is coded by the four DNA bases adenine (A), cytosine (C), thymine (T) and guanine (G) (see structure of DNA).

#### **Biocontainment**

See Containment

#### **Bioreactor**

A system in which microbial conversion processes of organic substances occur under controlled conditions and can be measured

#### **Biosensors**

Devices for measuring physical and chemical vital processes on and in organisms. Today, the term means a set-up in which a biological component (e.g. an enzyme, an antibody or a microorganism) is linked directly to a signal transducer (e.g. an electrode, optical fiber, piezo-crystal or transistor).

#### **Biotechnology**

Use or utilization of living organisms or their elements to produce, modify or break down substances for services (diagnostics/analytics) or to change organisms. The term "modern biotechnology" is used for innovative methods, processes or products which include the essential utilization of living organisms or their cellular constituents.

### **C**

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#### **cDNA**

(= Complementary DNA) Single or double-strand DNA copy of an RNA

#### **Cell**

The basic unit of every organism containing genetic material, an energy-producing system and other components

#### **Cell culture**

Cultivation and propagation of cells in liquid culture media or on solid culture media, in specific vessels or systems

**Cell cycle**

The cycle of growth, DNA doubling, growth and cell division in the case of eukaryotes

**Cell suspension**

Single cells in a nutrient solution

**Chromatography**

A generic term covering the processes of the physical separation of substance mixtures on the basis of their differing distribution between a solid and a liquid phase

**Chromosomes**

A sub-unit of the genome in higher organisms. It primarily consists of DNA and protein.

**Clone**

A colony of genetically identical cells. They result from divisions from a single cell.

**Cloning**

Production of cells or entire organisms with identical genotypes. The original cell originates, for example, from an early embryonic stage. Also: production of cells containing modified genotype (hereditary substance).

**Complementary DNA**

cDNA, a DNA which is synthesized in a laboratory w. the aid of mRNA matrices

**Containment**

Prevention of the uncontrolled propagation of transgenic DNA

**Codogenic strand**

The strand of the double helix which contains the genetic information

**Codon**

Sequence of three bases (triplet) on the mRNA with information for ribosomes on protein production

**Cytokines**

A generic term for numerous endogenous substances which are released by immune system cells during the immune response. They are important for repair mechanisms to repair tissue damage and specifically stimulate the growth of cells. Cytokines include interleukines (IL), interferons and the growth factors GM-CSF and G-CSF (see Chapter 7).

**Cytoplasm**

Content of a cell, with the exception of the cell nucleus, consisting of a fluid medium and numerous cell organelles

**Cytosine**

Organic pyrimidine base, an element of nucleotides

**Cytostatics**

Substances which kill off division-active eukaryotic cells, i.e. including human cells. Cytostatics are used to treat various forms of cancer.

**D**

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**Diagnostic markers**

For marking single cells which can be visualized with corresponding diagnostic methods (such as blue color when subject to UV light)

**Differentiation**

Higher organisms contain different types of cells that perform specialized functions. The human skin cell in must perform tasks different from those of a liver cell. The process that causes specialized cells to occur is referred to as differentiation. In higher organisms, all cell types are produced by division and differentiation from the fertilized egg cell.

**DNA**

Deoxyribonucleic acid. The idioplasm of all organisms (apart from viruses). DNA consists of linearly linked nucleotides, the sequence of which forms the hereditary information.

**DNA clone**

A section of DNA which is inserted into a vector (plasmide) in order to produce several copies.

**DNA polymerase**

An enzyme which catalyses the synthesis of DNA on the basis of DNA parent material (e.g. in replication). Frequently used for in-vitro synthesis of DNA sections.

**DNA replication**

The process of reproducing copies of DNA strands

**Domain**

The smallest unit of a protein with a defined and independently folded structure. Domains consist of 50 – 150 amino acids and frequently perform individual reactions whose interaction forms the overall function of a protein.

**Double helix**

Two DNA strands wound together helically

**Dots**

Measuring points with coupled biomolecules on a biochip

## **E**

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### **Electrophoresis**

A separating method based on the migration of electrically charged colloid particles (charged electrically as the result of attachment or expulsion of ions) in an electric field. It is primarily used to separate and precipitate off small quantities of substances, e.g. DNAs.

### **Electroporation**

A process for admitting DNA into cells using a brief electrical pulse

### **Elongation**

The middle phase of translation. Further reading of the mRNA.

### **Enzymes**

Proteins which act as an accelerator or promote a reaction in biological or biochemical processes

### **Exon**

Encoding section of the gene, which bears information, which is still present in the spliced mRNA and which has not been previously cut out by enzymes. Exons define the amino acid sequence of proteins.

### **Expression**

Conversion of genetic information into a corresponding protein

## **F**

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### **Fermentation**

A technical process for producing or converting substances with the aid of microorganisms

### **Flow cytometry**

Method of optical particle measurement used to identify and characterize heterogeneous cell populations

### **Fluorescence microscopy**

A process which utilizes the phenomenon of fluorescence to sharply visualize (with high contrast), specifically selected cell constituents contrasted from other components.

### **Functional genomics**

Analysis of the functional interplay of genetic information

## **G**

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### **Gel electrophoresis**

A method of separating nucleic acid molecules or proteins embedded in a gel on the basis of their mobility in an electrical field

### **Gene**

The basic unit of the hereditary information. A gene consists of a DNA section which contains the information for synthesis of an RNA. In certain cases, the RNA itself is the end product.

Generally however, it serves to transport the genetic information to the ribosomes where proteins are then formed.

### **Gene expression**

Conversion of the genetic information from the DNA via a messenger RNA (mRNA) to a protein

### **Gene probe**

A specifically marked single-strand nucleic acid (DNA, RNA) which can detect a specific gene and thus hybridize owing to its complementarity

### **Genetic code**

The translation code for translation between the information units (codons) of the genes and the amino acid elements of the proteins. Apart from very slight details, the genetic code is identical on all living beings

### **Genetic engineering**

A sub-area of biotechnology. A method which allows genes to be identified, isolated and recombined in a test tube and which allows them to be transferred in targeted manner to other organisms and produce an action there.

### **Genetic therapy**

An attempt to heal diseases, e.g. by the introduction of intact genes into the "sick" cells. Somatic gene therapy: non-hereditary; germline therapy: hereditary.

### **Genome**

The entirety of all the genes of an organism

### **Genomics**

Genome research: investigation of the genes and their functions

### **GMP**

Good Manufacturing Practices. The basic rules for production and quality inspections. Prerequisite for approval ("registration") as a medication.

## **H**

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### **Histones**

Basic proteins from which the protein complexes of the chromosomes are built

### **Horizontal gene transfer**

Transfer of genetic material to another, non-related species

### **Host cell**

Cells which fungi, bacteria and viruses require in order to live

### **Hot spots**

DNA regions susceptible to gene mutation (e.g. TT)

### **HPLC**

High-Performance Liquid Chromatography: a chromatographic method in which liquid substances (or substances dissolved in a liquid) are transported by means of a solvent (eluent) under high pressure through a separation column, where splitting into individual components occurs at the stationary phase (over the retention times). It is used frequently as a preparation for production of pure substances.

### **Hybridization**

Bonding of a nucleic acid single strand (DNA or RNA) by base pairing to another complementary single strand

## **I**

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### **Immune system**

Humankind's biological defense system for protecting against the penetration of foreign substances

### **In silico**

Literally: in silicon. Silicon is the material of which computer chips are made. Thus, in silico means: in the computer (bioinformatics).

### **In vitro**

(= in the test tube): outside of the organism

### **In vivo**

In the living organism

### **Initiation**

Starting operation of translation. The ribosome unit bonds with the mRNA.

### **Insertion**

Inserting DNA sections into another DNA molecule

### **Intron**

Section of a gene which interrupts the coded area and which is not translated into protein

## **K**

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### **Kinases**

A special class of enzyme which joins phosphate groups to proteins

## **L**

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### **LC/MS**

Coupling system comprising liquid chromatography and mass spectrometry

### **Ligand**

A natural or synthetic chemical molecule that binds selectively to a protein

### **Lymphocytes**

A specific class of white blood corpuscles which are of importance for the immune system

## **M**

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### **MALDI**

Matrix-Assisted Laser Desorption/Ionization; a combination of chromatography and mass spectrometry

### **Marker genes**

A gene in plasmids or viruses serving as a marking for detection of the genetically modified object

### **Messenger RNA**

mRNA. The RNA of a gene is transcribed to the mRNA molecules, which then serve as a blueprint for protein synthesis.

### **Monoclonal antibodies**

Structurally identical antibody clones produced by genetic engineering

### **mRNA / messenger RNA**

Ribonucleic acid molecule which is produced in the process of transcription from the DNA and which contains the information for synthesis of a protein

### **mRNA maturing/post-transcriptional processing**

Processing transcribed mRNA to form the definitive version by cutting out the introns.

### **Multi-potency, multi-potent**

The property of tissue-typical stem cells to be able to develop into different cell types of an organ

### **Mutation**

A change in the DNA sequence that can lead to the synthesis of modified, inactive protein

## **N**

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### **Nucleic acid**

The family of molecules which represent the building blocks of DNA and RNA molecules

### **Nucleoli**

**Areas in the nucleus in which rRNA is formed**

### **Nucleotide**

A single building block of DNA or RNA, consisting of a compound of phosphate with a sugar molecule comprising five carbon atoms (pentose) and a pyrimidine or purine base such as adenine, guanine, cytosine or thymine

### **Nutrient solution**

Liquid nutrient media for cultivating microorganisms and cells

## **O**

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### **Oligonucleotide**

(oligo, Greek: little or small)

A sequence of a few to several hundred interlinked nucleotides. An oligonucleotide thus corresponds to a (very) short DNA single strand.

### **Omnipotency, omnipotent**

Designation for cells that still feature all functions of the overall organism

## **P**

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### **PCR (Polymerase Chain Reaction)**

A process for propagating extremely small quantities of DNA on an in-vitro specific basis and identically without cloning

### **Peptide**

A short chain of amino acids

### **Phages**

Viruses which attack bacterial cells

### **Phenotype**

The manifestation and specific characteristics of an organism – determined by the interaction of the genotype and the ambient influences

### **Phosphatases**

A special class of enzymes that removes phosphate groups from proteins

### **Plasmides**

Small annular section of DNA of bacterial origin which is capable of independent reproduction within a host organism. Most genetic engineering experiments are performed on and with plasmids.

### **Pluripotent, pluripotency**

The potential of embryonic stem cells to mature (differentiate) in cells of different tissue types.

### **Polymerase**

An enzyme responsible for replication of DNA

### **Polymerase chain reaction**

See PCR

### **Primer**

A short RNA sequence, the starter molecule for DNA replication

### **Promotor**

A nucleotide sequence upstream of the gene which is crucial to whether the gene can be read and in what quantity it can be produced

### **Protein**

High-molecular compound (chain) of amino acids. Very diverse tools and building blocks of cells, which may have many functions (e.g. enzymes, hormones, structure proteins, transport proteins, antibodies, neurotransmitters and receptors).

### **Proteomics**

The investigation of proteins and their function

### **Protoplasts**

Cells without cell walls

### **Purine bases**

Adenine and guanine that are derived from purine

### **Pyrimidine bases**

Cytosine, thymine and uracil which are derived from pyrimidine

## **R**

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### **Recombinant antibodies**

Antibodies which are comprised of proteins whose hereditary information originates from different species

### **Recombinase**

An enzyme which attacks the DNA at specific points, cuts out sequences and connects sequences together (recombines them)

### **Recombination**

Combination of DNA of differing origin

### **Restriction enzymes**

Enzymes which detect a short, defined nucleotide sequence on the DNA and split the DNA at this point or its vicinity

### **Reverse transcriptase**

A viral enzyme that can rewrite RNA to DNA. It is used for in-vitro synthesis of cDNA.

### **Receptors**

Molecules on the cell surface which are capable of binding a precisely defined molecule (their ligands)

### **Ribosomes**

Cell organelles which serve to synthesize proteins

### **RNA (ribonucleic acid)**

Produced by transcription of the DNA and contains, amongst other things, the information for synthesis of a protein. It occurs as mRNA, tRNA and rRNA. In contrast to DNA, it contains uracil instead of thymine. It is generally single-strand.

### **rRNA**

Ribosomal RNA. It is involved in the structure of ribosomes.

## **S**

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### **Screening**

Mass testing of substance banks with an assay

### **SELDI**

Surface-Enhanced Laser

Desorption/Ionization. A combination of chromatography and mass spectrometry

### **Sequence analysis / sequencing**

Determining the base sequence of a DNA or RNA strand or the amino acid sequence in a protein

### **Somatic genetic therapy**

Genetic therapy on cells of a body, apart from the germ cells; non-hereditary

### **Specificity**

The property of active substances to differentiate between different but similar target structures

### **Splicing**

Converting a primary RNA transcript to a mature, translatable mRNA by removing intron sections

### **Starter codogene**

Triplet at the start of a gene that determines the start of transcription on the DNA

### **Starter culture**

A bacterial culture which performs the acidification process (e.g. for yogurt and cheese)

### **Structure gene**

The part of a gene which contains information for the proteins

## **T**

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### **Target**

A biological molecule, e.g. an enzyme, which plays an important role in the production or development of an illness or disease

### **Target molecule**

The point of attack of active substances (therapeutics) in sick cells

### **Terminator codogene**

The triplet at the end of a gene which determines the end of the transcription on the DNA

### **Termination**

End phase of translation. Detachment of the ribosomes from the mRNA.

### **Thymine**

Organic pyrimidine base. An element of the DNA nucleotides.

### **Tissue engineering**

An interdisciplinary area of biotechnology. It deals with processes for in-vitro reproduction of higher cells, in particular for use of specific body tissues such as skin or cartilage

### **TOF**

Time-Of-Flight mass spectrometer

### **Totipotent, totipotency**

The property of early embryonic cells (generally as far as the 8-cell stage) to be able to develop to form a complete organism even after separation from the embryo

### **Transduction**

Transfer of genes by viruses to other cells

### **Transformation**

Natural and artificial transfer of DNA to a cell

### **Transgene**

An organism containing foreign genes

### **Transcription**

The first step of gene expression. An mRNA copy of an information-bearing DNA section (gene) is synthesized in the cell nucleus by an RNA polymerase enzyme complex.

### **Translation**

Second step of gene expression. The information transferred to the mRNA is read by ribosomes and translated to the corresponding amino acid sequence (protein).

### **Triplet**

(see codon) A sequence of 3 nucleotides within the DNA. A defined amino acid in a protein is assigned to a triplet in the DNA in accordance with the rules of the genetic code.

### **tRNA**

Carrier molecule for amino acids during translation

## **U**

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### **Uracil**

Pyrimidine base comprising ribonucleic acid

## **V**

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### **Validation**

Verification of the specific action of an active substance on a target, with the aim of achieving a clear therapeutic benefit

### **Vector**

DNA molecule (e.g. a plasmid) that can be inserted into cells and which is generally forwarded to the daughter cells by the host cells during division. Vectors are used for transfer of foreign DNA sections.

### **Virus**

An infection particle (not a cell), consisting of a protein coat and a genome (DNA or RNA). Viruses may penetrate specific cells and insert their genotype (DNA or RNA). The genotype of the viruses can be integrated in the genome of the cell (always as DNA) and may remain ineffective for a long time. If it is activated, new viruses are produced and the host cell generally dies.

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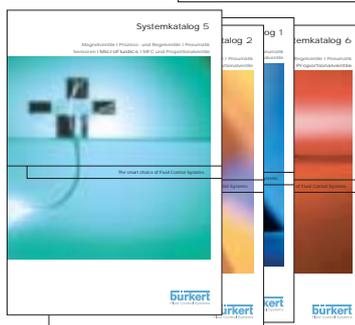
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