Science with Passion





AZURA[®] Preparative HPLC



Configure the best possible system for your purification workflow of peptides, oligonucleotides, RNA, cannabis and many other API with the marketleading AZURA preparative HPLC platform.

think LC. think KNAUER.

AZURA® Preparative HPLC Customized purification

AZURA® preparative systems are the perfect solution for frequently changing separation tasks - from milligram to kilogram scale. Design your AZURA preparative system to your needs and combine flexibility and reliability.



AZURA® Prep systems are tailor-made for you. Configure your system from injection to detection and choose between different materials, flow rates, valves and detectors.

Due to the flexible design of our devices, vou can easily change parts like pump heads or flow cells and integrate all components of the compact into the pilot-scale system.

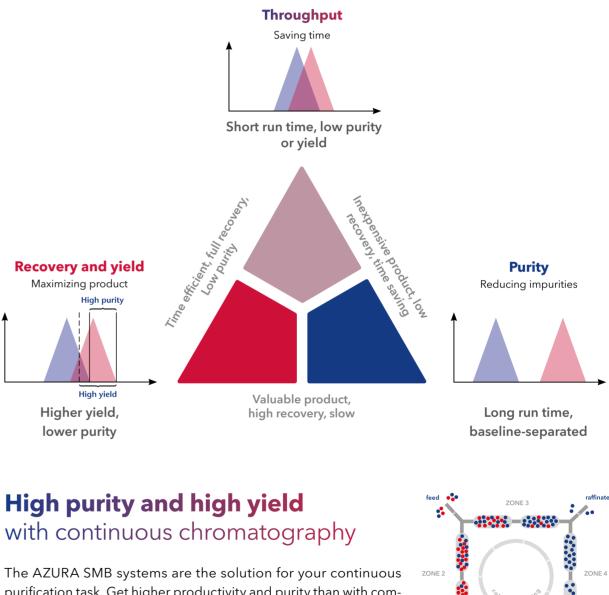
AZURA[®] Prep systems can be used for special separation modes like peak recycling and stacked injections. We help you to configure your system and choose the best software for you.

Preparative chromatography

The general objective of preparative chromatography is to isolate, purify and collect your target compounds. Preparative applications are often initially performed on an analytical level and need to be upscaled. Depending on the desired scale, the requirements for a preparative system differ in eluent supply, sample injection, column, and detection. We customize our systems to meet your chromatography scale-up and purification challenges. Benefit from our experience in preparative chromatography. For more information: www.knauer.net/prep

Purification strategy: Prioritize purity, throughput or yield?

The dependencies between throughput, purity and yield always have to be considered in HPLC purifications. Whatever priority you decide for, with the AZURA preparative systems you can successfully adapt.



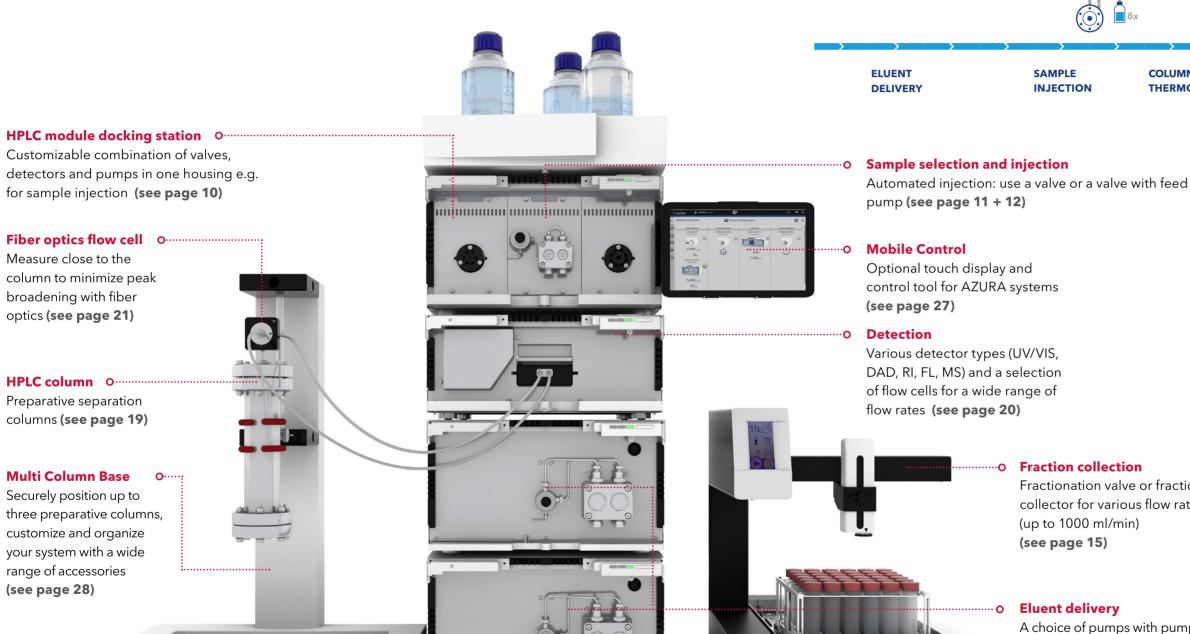
High purity and high yield with continuous chromatography

purification task. Get higher productivity and purity than with comparable batch systems. Save up to 90 % of the solvent and reduce the solid phase costs up to 80 %.

For more information see page 37.

Scheme of SMB principle

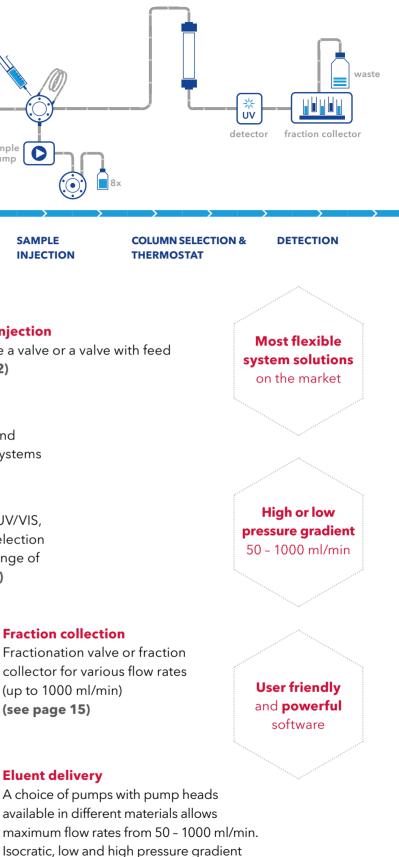
Flexibility and performance



forming are possible (see page 8)

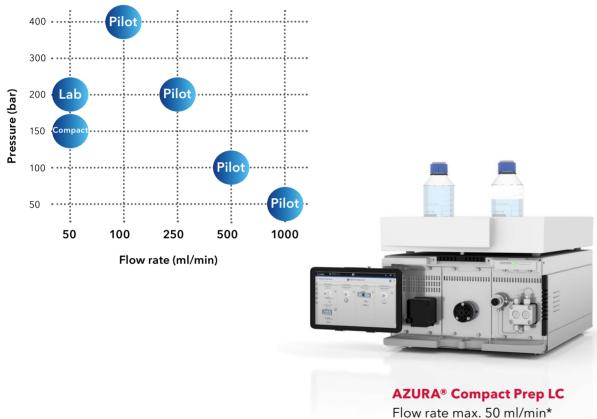
C

sample



AZURA® Preparative HPLC Upscaling from compact to pilot

The modular AZURA Preparative HPLC platform offers you the opportunity to build a purification system best suited to your needs.



Isocratic

| AZURA system | Available pump heads Max. flow rate in ml/min* | | | Gradient options | | | |
|-----------------------|---|-----|-----|------------------|------|---------------------|----------------------|
| | 50 | 100 | 250 | 500 | 1000 | LPG low pressure | HPG high pressure |
| AZURA Compact Prep LC | • | | | | | | |
| AZURA Lab Prep LC | • | | | | | | • |
| AZURA Pilot Prep LC | | • | • | • | • | • | • |

* Information on best working conditions on pages 8 - 9.



AZURA® Lab Prep LC Flow rate max. 50 ml/min* Isocratic/HPG

Scale-up from compact to pilot

The AZURA Pilot Prep LC is the ideal solution for your upscaling **For more information**: tasks. The 100 ml pump head allows you to run your system under analytical conditions before adapting your method to preparative scale.



AZURA® Pilot Prep LC

Flow rate max. 1000 ml/min* Isocratic/LPG/HPG

www.knauer.net/prep

Eluent delivery

Precise and reliable pumps covering a wide flow range for various gradient and solvent selection options.

AZURA® Pump P 2.1L

The preparative HPLC pump AZURA P 2.1L covers a wide flow rate and pressure range. It has been designed for the purification of milligram to gram samples. The integrated automatic RFID pump head recognition allows a quick adaptation to various applications.

- Flow rate up to 1000 ml/min
- LPG and HPG gradient options
- Supports constant pressure mode

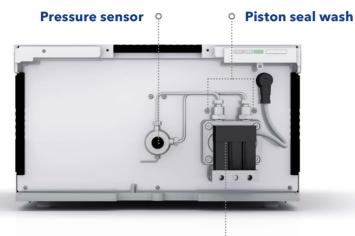
Gradient options of Pump P 2.1L

A low pressure gradient (LPG) module dynamically composes the eluent on the inlet-side or low pressure side of the pump head, by quickly switching between the different solvent channels. We offer binary or ternary LPG upgrade modules for the isocratic P 2.1L.

The eluent in a binary high pressure gradient (HPG) system is composed by combining the solvent flows of two isocratic pumps.



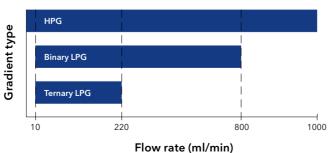
Ternary LPG valve block for the pump head



Exchangable pump heads $\dot{\circ}$ (stainless steel or titanium)

| Pump head | Max. pressure | Best working conditions |
|--------------|------------------|-------------------------|
| 100 ml | 400 bar | 1 – 80 ml/min |
| 250 ml | 200 bar | 2.5 - 200 ml/min |
| 500 ml | 100 bar | 5 - 400 ml/min |
| 1000 ml | 50 bar | 10 - 800 ml/min |

Covered flow rate



AZURA® Pump P 6.1L

The AZURA semi-preparative pump P 6.1L with 50 ml pump head is available as an isocratic or binary HPG pump. It is made for medium-size purification tasks and upscaling processes.



Solvent selection

For automated solvent change, a solvent selection valve can be attached to the pump P 2.1L.

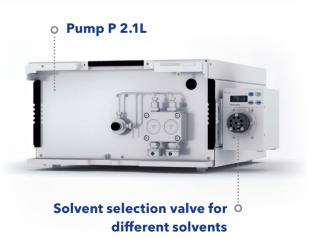
For semi-preparative purification tasks, the pump P 6.1L features a built-in 2x2 solvent selection valve (high pressure gradient version).



- > 1/8" tubing up to 80 ml/min
- > 1/4" tubing up to 1000 ml/min

o Solvent selection valve

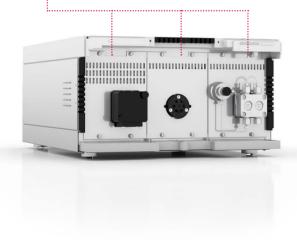
- Flow rate up to 50 ml/min
- Best working conditions: 0.1 - 40 ml/min
- Binary gradient with solvent selection valve (2x2 solvents)
- Up to **300 bar** < 10 ml/min
- Up to 200 bar max 50 ml/min



Docking station for pumps, valves and detectors AZURA® Assistant ASM 2.2L

The Assistant ASM 2.2L is a docking station for three compact devices. Valves, pumps and UV detectors can be combined in one housing. The plug-in modules are removed by loosening four screws allowing the user to exchange modules in case of service within minutes. Likewise, the configuration of the HPLC system can be adapted to new requirements. Routine maintenance work e.g. replacing the lamp of a detector is easily performed by the user.

> Freely combine pumps, valves and detectors in one housing



Depending on the integrated modules the assistant fulfills many different tasks like eluent delivery, detection, sample and solvent selection, sample injection, column switching or fraction collection. An assistant including a pump, injection valve, and detector features a complete HPLC system, like AZURA Compact Prep LC. As a part of a larger system, the ASM 2.2L allows the user to customize the system configuration according to the purification challenge.

Pumps

Choose from 15 different pumps with 10 or 50 ml pump heads and with or without pressure sensor. The material used is stainless steel, ceramic or Hastelloy C (for pumps without pressure sensors only).

Valve drive

The universal valve drive identifies valves via RFID technology and enables to read GLP data. All V 4.1 valves, independent of number of ports and position, are supported.

UV detectors

The compact single wavelength UV detector is available in a basic and fibre optics version. The wavelength can be set between 190 - 500 nm.

& Configure your assistant

Use the web-based assistant configurator to find your desired AZURA® ASM 2.2L module combination: www.knauer.net/assistantconfigurator



Sample injection

Adapt the sample injection mode to your preparative task.

Injection valve

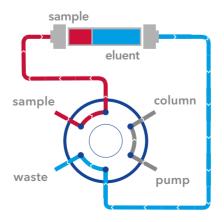
The simplest way to inject your sample into the system. Use a manual injection valve and choose from a large range of different sample loops.

KNAUER offers several injection valves for 1/16" and 1/8" tubing. The wetted parts are made of stainless steel or PEEK to cover a broad range of applications. Injection can be done either manually via hand lever or automated with a valve drive.

The Multi-Injection valve for 1/16" tubing enables the sample injection via loop and sample pump through just one valve, perfect for often changing applications.

VariLoop for sample injection

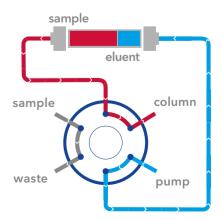
the injection of medium up to high sample volumes (up to 40 ml). The sample loop can be emptied completely or partially as well as filled completely







- The KNAUER VariLoops are the perfect solution for or partially. This allows you to work very flexible
 - and easily switch between different sample sizes
 - while keeping constant and reproducible injection
 - volumes for every sample size.



Autosampler AS 6.1L

Sample injection can be easily automated with an autosampler. The AS 6.1L can inject up to 10 ml per injection. Sample tray temperature control from 4 - 40 °C is optional available. It can handle either 30 samples in 10 ml vials or up to 768 samples in well plates.

Automated sample injection also possible with Liquid Handler LH 2.1. More information on page 14.

Sample Injection Assistant ASM 2.2L

The AZURA Sample Injection Assistant is based on the docking station for HPLC modules ASM 2.2L. It is designed to automate injection of larger sample volumes and features a sample selection valve, a sample pump, and an injection valve.

Simply attach your sample vessels via 1/8" tubing to the multiposition valve and automate injection with the integrated sample pump and injection valve.

Sample pump

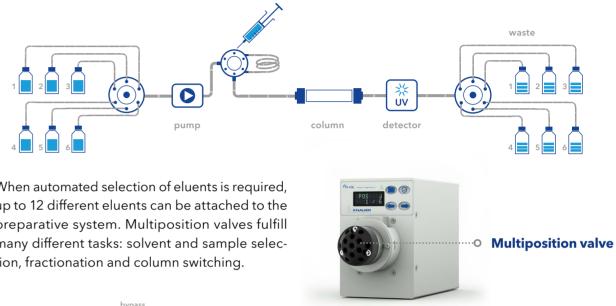
Standalone or as a plug-in module of the HPLC dockingstation ASM 2.2L: The compact pump AZURA P4.1S is perfect for feed injection.

- 10 and 50 ml exchangeable pump head
- Flow rate range: 0.01 - 50 ml/min (50 ml pump head) 0.001 - 10 ml/min (10 ml pump head)
- Pump heads available in stainless steel or ceramics
- Best working conditions: 1 - 40 ml/min (50 ml pump head) 0.1 - 8 ml/min (10 ml pump head)

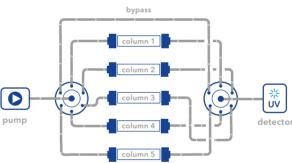




Automation for prep HPLC Eluent selection and fractionation



When automated selection of eluents is required, up to 12 different eluents can be attached to the preparative system. Multiposition valves fulfill many different tasks: solvent and sample selection, fractionation and column switching.



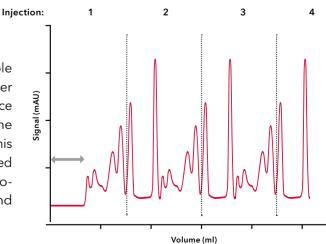
Example for the selection of 5 columns

Stacked injection

With the stacked injection function, it is possible to perform different runs automatically one after the other. The injection of the next run takes place during the current run, so that the time until the elution of the first peak can be fully exploited. This increases efficiency, saves time and eluent. Stacked injection can be operated with the chromatographic data systems (CDS) PurityChrom® and OpenLab[®].

Column switching

Switching valves are ideal for screening and scaleup. They can be easily integrated into your system at pressures up to 400 bar and maximum flow rates of 300 ml/min. Up to 16 columns can be selected.



Liquid Handler LH 2.1 - Injector & Fraction Collector

the expansion of purification processes with the of purification. The handler injects samples with ability to combine sample injection and fraction minimal loss regardless of their volume - perfect collection in one device. A high capacity of sample for working with expensive compounds. and fraction vessels meets a flexible arrangement

- **Combine** sample injection and fraction collection
- Scalable injection range up to 60 ml
- **Purify** from milligrams to several grams
- Flexible arrangement of samples and fractions via teaching option
- Reinject collected fractions
- Perform in high-throughput peptide and oligonucleotide workflows

Technical data

Fraction collection Fraction capacity Maximum vessel capacity with 5 KNAUER racks • 15 x micro titer well plates 810 x 2 ml tubes • • 490 x 15 ml tubes 160 x 50 ml tubes • **Diverter valve** yes Number of racks 5 KNAUER racks, teaching module for all racks Sample injection Sample injection standard and sandwich injection mode Sample loop up to 60 ml; larger loops on request Injection valve 1/16" or 1/8" V 4.1 injection valves and VU 4.1 supported **Temperature control** no Needle wash single needle wash step after each injection Wash solvents 4 Aluminium oxide 99.5 %, Borosilicate Glass, PTFE, FEP, Wetted materials AISI 316L, PEEK

KNAUER's new Liquid Handler LH 2.1 allows for facilitating reinjection of samples to reach new levels

KNALLER

Fraction collection

Collect large quantities or large numbers of fractions

KNAUER offers different valves for fraction collection and variations of trusted fraction collectors. Whether you are doing research and development or production, there is an appropriate solution that suits your application.

Fractionation modes:

Manually - collection by direct control Time-based - collection at defined time points Peak-based - collection according to detector signal Threshold function - collection according to any signal

Fraction collectors

LABOCOL Vario 4000 / Plus

The LABOCOL Vario 4000 fraction collectors are fining the number of fraction vessels and their characterized by their high robustness and optimal position. The wide application area makes the Vario ratio of dimensions/benefit. The user is not limited 4000 series ideal for use in research and development as well as in production. The Vario 4000 modto given rack types. The rack layout can be designed according to individual needs. Free rack els differ in the base area and the flow rate range. design. Any rack type can be integrated by de-

Rack type 80 Tubes 18 mm / max. 36 ml 125 Tubes 10.5 mm / max. 9 ml 20 Tubes 36 mm / max. 140 or 240 ml 39 Tubes 26 mm / max. 80 ml 24 Centrifuge tubes 50 ml





Foxy[®] R1

The Foxy® R1 fraction collector can be adapted to a broad spectrum of applications. Flow rates of up to 125 ml/min are possible. Fractions can be collected into 96 well microplates, standard tube sizes, bottles and many more. For essentially unlimited volumes, funnel racks can direct fluids to any collection vessel or downstream process. Both devices can be operated stand-alone or in the chromatography software PurityChrom[®].



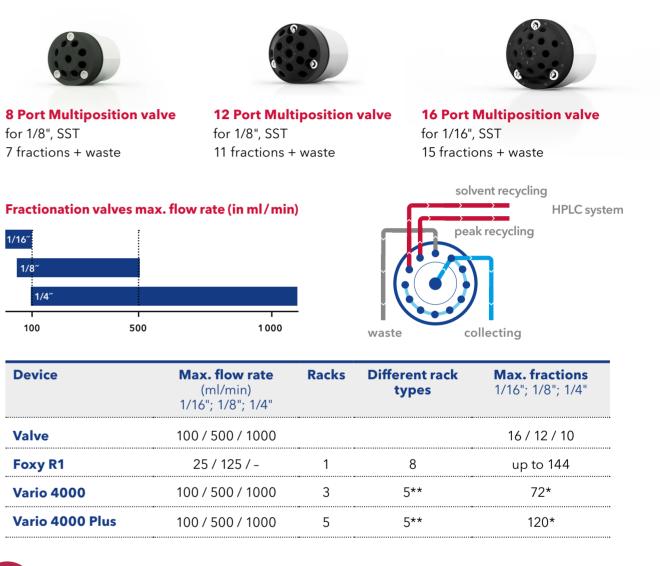
Fractionation also possible with Liquid Handler LH 2.1. For more information see page 14.

Rack type

| 144 Vials 13 mm / max. 9 ml |
|------------------------------|
| 100 Vials 16 mm / max. 20 ml |
| 36 Vials 25 mm / max. 70 ml |
| 2 Microwell plates 96 |
| 60 Tubes 1.5 ml |
| 72 Centrifuge tubes 15 ml |
| 36 Centrifuge tubes 50 ml |
| 36 Funnels with vinyl tubing |







| Device | Max. flow rate (ml/min) 1/16"; 1/8"; 1/4" |
|-----------------|--|
| Valve | 100 / 500 / 1000 |
| Foxy R1 | 25 / 125 / - |
| Vario 4000 | 100 / 500 / 1000 |
| Vario 4000 Plus | 100 / 500 / 1000 |

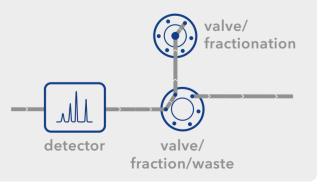


For contamination free collection, the combination of two valves is perfect

espacially as - the PurityChrom software is able to adress the matching delay volume to each valve.

* For 50 ml tubes

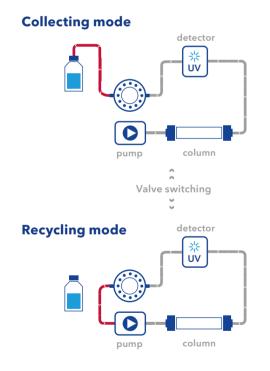
** Device supports other racks via user-defined position setting.

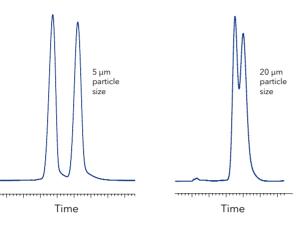


Advanced Purification Modes

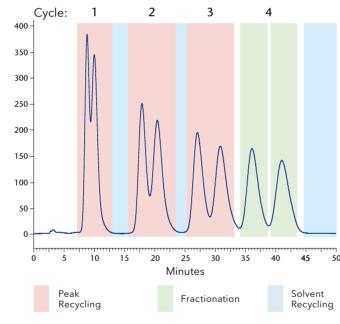
An example

A separation can be much more demanding after upscaling from analytical to preparative scale. In many cases a baseline separation is not possible anymore, so time and money consuming method development or hardware adjustments are necessary. The AZURA Prep LC system is well-suited to apply the peak recycling technique to solve demanding resolution tasks. Additionally, solvent recycling can be applied to save eluent, if it can be considered clean.









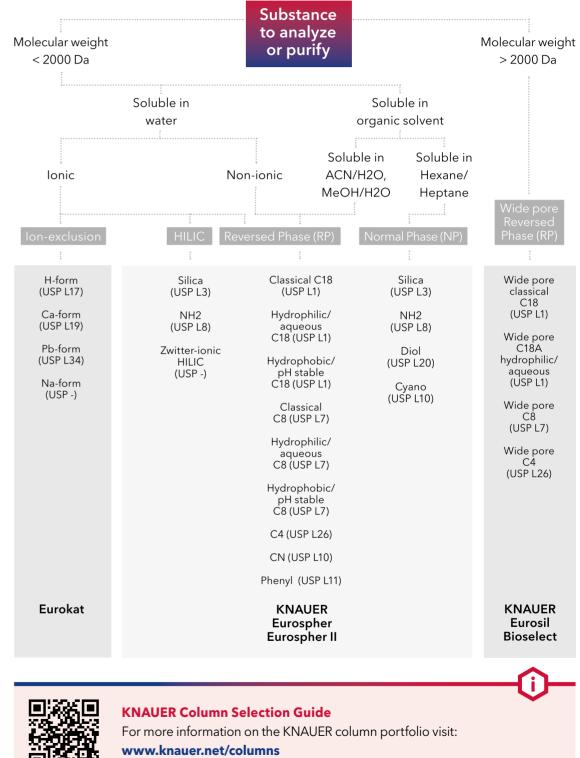
Successful peak separation with recycling mode.



KNAUER preparative columns

Find the perfect column from the large KNAUER portfolio.

This flow chart gives you a guideline how to select the right column for your application. Start at the top and follow the decision lines all the way down to find a column recommendation.





Application Note (VTN0005)

Comparing sensitivity levels for the analysis of fluorescence-labeled proteins www.knauer.net/applications

Detection

KNAUER gives you the opportunity to analyze nearly every compound due to a large portfolio of HPLC detectors. For the achievement of your analysis goals and for matching your separation scale, our detectors are flexible in the setup, including flow cells and fiber optics. Our product line of UV/VIS detectors ranges from single variable wavelength to 8-channel diode array detectors with 3D scan capability.



| Detector | UVD 2.1S | UVD 2.1L | MWD 2.1L | DAD 2.1L | DAD 6.1L |
|---------------------------|---|---|------------------------------|---|-------------|
| | Compact and versatile UV detector | Reliable UV/ VIS detector for a wide spectrum of applications | Robust multi- channel UV/ | Versatility through a wide flow cell range | High-end |
| Wavelength | 190-500 nm | 190-750 nm | 190-700 nm | 190-700 nm | 190-1000 nm |
| Channels | 1 | 1 | 4 | 8 | 8 |
| 3D scan | | | | • | • |
| Fiber optics available | • | • | • | • | • |

Flow cells for UV/VIS and DAD detectors

Select from an impressive range of easily exchangeable preparative and semi-preparative flow cells for UV/VIS and DAD detectors. With capillary connections ranging from 1/16" to 1/4" and TRI-Clamp adaptions, optional fiber optics technology and a variety of flow cell wetted materials, a wide spectrum of applications can be covered.

| Max. flow rate | Connectors | Path length | Volume | Max. pressure | available |
|-------------------|------------|---------------|----------------|---------------|-----------|
| 50 ml/min | 1/16" | 3 mm | 2 µl | 300 bar | • |
| 250 ml/min | 1/16" | 0.5 mm | 3 µl | 200 bar | • |
| 1000 ml/min | 1/8″ | 0.5/1.25/2 mm | 1.7/4.3/6.8 μl | 200 bar | • |
| 10000 ml/min | 1/4′′ | 0.5/1.25/2 mm | 1.7/4.3/6.8 μl | 200 bar | • |

Fiber optics technology

More flexibility

Fiber optic cables offer the possibility to separate the flow cell from the detector. This enables demanding applications such as measuring directly after a heated LC column or in hazardous environments, allowing safe operation of the instrument while maintaining performance.

Safe operation

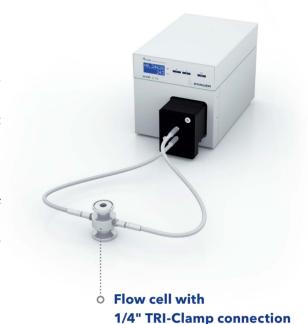
When working at high flow rates, separation of the flow cell and the detector is a safety feature. In case of leakages, no damage to the detector occurs. Fiber optics are available in a customized length of up to 10 meters.

AZURA® RID 2.1L HighFlow Preparative refractive index detector

The AZURA RID 2.1L HighFlow is a sensitive and competitively priced differential refractometer. It is suitable for detecting compounds with little or no UV activity such as alcohols, sugars, lipids or polymers in high concentrations. This instrument is designed for use in semi-preparative and preparative HPLC for flow rates up to 100 ml/min. Optional are higher flow rates possible with a flow splitter. The intelligent temperature control guarantees fast baseline stabilization and stable operation.

Application Note (VPH0068)

Cyclodextrin purification Part 2 - Method transfer and purification **www.knauer.net/applications**





Special detection

Choice of specialized detection technology, fully integrated in PurityChrom[®]. Suitable for preparative LC with the help of a flowsplitter.

Light Scattering Detector Sedex LC

Sensitive universal detection with the possibility to run gradients

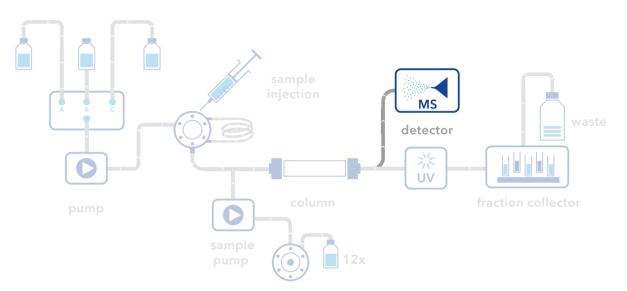
As a universal detector, an ELSD detector offers numerous possibilities for detecting substances that have few or no chromophores. Since the eluents are evaporated, the use of non-UV-compatible solvents poses no problems and the ELSD is gradient compatible.

Target analytes: Carbohydrates and similar compounds, detergents, ionic and non-ionics, artificial sweeteners, antioxidants, amino acids, lipids, peptides, polymers, pestizides, proteins, steroids.

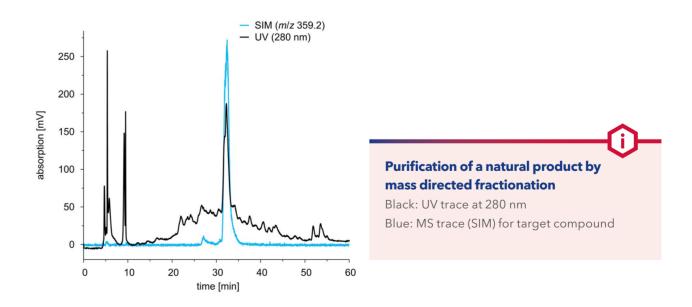




KNAUER offers several system solutions for mass triggered preparative chromatography applications. Systems can be operated using the Chromeleon[™] or PurityChrom[®] software.



The mass-controlled fractionation allows the selective isolation of substances that are not UV active. Furthermore, in combination with a UV detector and intelligent and/or algorithms, high purities can be achieved even with very difficult preparative separations.





AZURA® Conductivity Monitor **CM 2.1S**

The Conductivity Monitor CM 2.1S can monitor salt gradients with flow rates of up to 100 ml/min and a maximum pressure of 100 bar. It supports a wide measurement range of 0.01 mS/cm - 999 mS/cm. Flow cells in PEEK for both analytical and preparative scale are available.

Mikron 81 Conductivity Monitor

The mikron 81 is a reliable in-line conductivity monitor with a very low footprint and measures with high linearity in the range from 0.002 to 500 mS/cm. Its cutting-edge temperature sensor technology enables highly precise automated temperature correction of the conductivity signal. The intelligent flow cell design allows for a broad flow rate regime from microliter to lower liter per minute scale. It comes pre-calibrated and readyto-use with all accessories needed. Accordingly, the mikron 81 can be used to monitor salt gradients during purification or cleaning-in-place procedures as well as for upscaling (eg. from 1 ml/min to 1 l/min).



Temperature control

Increase performance. Minimize solvent viscosity.



Column Heating Sleeve

Our column heating sleeves are the perfect solution for thermostating your preparative column hardware. Available for all preparative KNAUER column dimensions at temperatures up to 100 °C. Custom dimensions, clean room compatible and autoclavable materials are available on request.



Eluent and Column Heater

When performing preparative LC at temperatures above 40°C in air-conditioned laboratories e.g. in

RNA purification processes, a uniform temperature

distribution is essential. With the Eluent Heater,

solvent temperature can be precisely controlled

using the integrated touchscreen. It supports flow rates of up to 500 ml/min and is cleanroom com-

Pump Head Heater

Electrical heating element for pump heads. Temperature can be controlled using the eluent heater or a single device control unit.

Column Oven

patible.

This oven can heat up to 80 °C. It can accommodate up to 8 KNAUER columns with up to 250 x 50 mm inner dimensions.



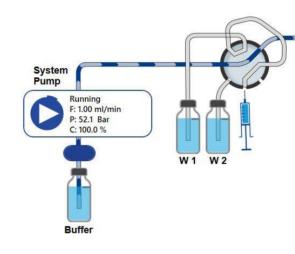
Software solutions

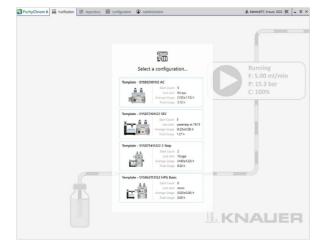
PurityChrom[®] 6

PurityChrom is a powerful software to control your purification system. With its revised, modern user interface, new powerful features and improved useability, PurityChrom 6 is a new generation of

Animated flow path

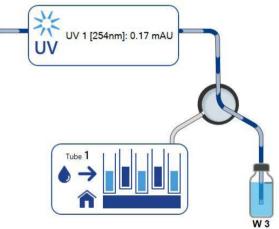
See exactly what you are doing and easily avoid mistakes with the animated flow path. Device functions for example starting the pumps, setting





our purification software PurityChrom. The software is developed according to GAMP 5 guidelines and is 21 CFR part 11 compliant.

the flow rates, switching the valves, change the wavelength of your UV detector are available via pop-up menus in the system visualization.



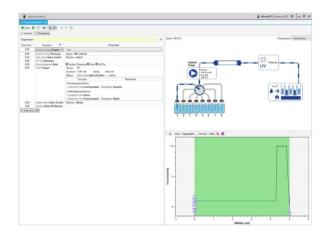
Configuration of multiple systems

PurityChrom 6 is as flexible as our KNAUER hardware. Multiple set ups can be configured in the software, e.g. configure the system with heating devices and without them, for all purification tasks that do not require temperature control.

Method writing with just a couple of clicks

In PurityChrom 6, methods can be written by simply clicking in the system visualization. Furthermore, a graphical editor enables a comparison between the current method and a previous chromatogram.

Methods can be created based on volume, column volume or time, depending on your preference. All solvent, waste and sample bottles can be configured and fill levels and solvent consumption are calculated, offering extra security during operation.



Advanced user administration

In PurityChrom 6, users can be assigned to a role simplifying the administration of a high number of users. An audit trail summarizes all actions of all users. Additionally, each chromatogram contains a run protocol summarizing the functions and events occurring during the method

| PurityChrom 6 W Purification | Repository Configuration | | | | |
|------------------------------|--|--|--|--|--|
| Users | Role: Assistant | | | | |
| Admin | | | | | |
| & Flaine | Configuration Create | | | | |
| | Delete | | | | |
| & Cosmo | Edit Rename | | | | |
| 🔏 Lab Manager | Kename | | | | |
| + Add | Environment Edit Settings | | | | |
| | File Sustant Copy Files | | | | |
| loles | File System | | | | |
| & Administrator | Create Macro | | | | |
| Auministrator | Create Method | | | | |
| 🚨 Laboratory Manager | Create Sequence | | | | |
| Assistant | Delete Directory | | | | |
| Assistant | Delete Macro Delete Method | | | | |
| + Add | Delete Result | | | | |
| | Delete Sequence | | | | |
| Actions | Rename Files | | | | |
| Print | Rename Directories | | | | |
| | Edit Macro | | | | |
| | ✓ Edit Sequence | | | | |
| | Automatic Reintegration | | | | |
| | Integration Cite Peak Windows | | | | |
| | Manual Reintegration | | | | |
| | Edit Check List | | | | |
| | Method Edit Information | | | | |
| | Cdit Procedure | | | | |
| | Edit Result Naming | | | | |
| | Administration | | | | |
| | Section Configuration | | | | |
| | Repository | | | | |
| | ✓ Purification | | | | |
| | Allow Device Control During Run | | | | |
| | Purification Allow Device Control During Run | | | | |
| | Allow Direct Commands | | | | |
| | Sign Reconstruction Documents | | | | |
| | Signature Save Signed Reconstruction Documents | | | | |
| | Validate Reconstruction Documents | | | | |

ClarityChrom[®] CDS

ClarityChrom is an easy-to-use chromatography data system (CDS) for workstations. Besides support of all KNAUER devices, components and systems from more than 45 manufacturers are also supported. ClarityChrom® includes the drivers for several fraction collectors and supports peak recognition by level and/or slope. The manual fraction control and the option to use the KNAUER electric valves for fractionation give you even more flexibility.

- Fraction collecting via peak recognition (level only, slope only, level AND / OR slope - incl. self-learning) or single event (unconditional, timed event)
- Easy to collect: waste, collect to position / collect to next, solvent recycling
- Direct control during a run manually switch to: collect, waste, solvent recycling
- Consecutive runs: easily find your chromatogram by clicking on your fraction

OpenLab

OpenLAB CDS EZChrom Edition provides support of devices from KNAUER and many other manufacturers. The KNAUER fraction collector control option includes the drivers of several fraction collectors and supports fractionation by time, the peak recognition by level and/or slope, also with spectral confirmation. Collect Slices allows for setting a desired volume for each fraction, within the defined fraction vial volume. The manual fraction control and the option to use the KNAUER electric valves for fractionation gives you more flexibility. The combination of virtual detector and virtual fraction collector allows for optimizing the fractionation settings from an existing chromatogram of your separations without any physically existing device and, therefore, without the loss of solvent or target substance.

Mobile Control (Chrom)

The hand-held Mobile Control (Chrom) allows a complete overview of all devices of the AZURA systems on one screen. Remotely check important parameters or control and monitor devices. The touch-optimized user interface facilitates navigation using just your fingers. The display software Mobile Control provides full access to AZURA devices. Change device settings, set operating parameters, automate device control or check the system status and GLP data... Mobile Control features all functionalities of a device display.

Do you want to acquire data without the overhead of a chromatographic data system? Mobile Control Chrom features data acquisition from AZURA detectors in addition to full device control. Basic purification tasks can be addressed by the function for threshold based fraction collection. Depending on the value of the detector signal, the target substances are automatically collected in different vessels. For each fraction collection block in a program an individual fraction volume can be defined. Choose Mobile Control as a basic, easy-to-use and cost-effective software solution!

Chromeleon[™] 7

Chromeleon is one of the most wide-spread chromatography data systems. It offers a broad range of third-party drivers and can be easily used with existing HPLC systems. Chromeleon drivers for many KNAUER devices are available.



Accessories

Improve system performance, organize your lab bench, and work more conveniently with the right accessories.

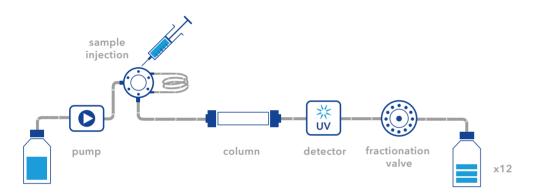
| Accessory | Features | Benefit |
|------------------------------|--|--|
| Pump head inlet | Connect one 1/4" tube to the AZURA Pump P2.1L Adapters for other diameters available | For high flow rates and viscous eluent |
| Mass flow controller | Unmatched accuracy at flow rates up to 833 ml/min Compatible with PurityChrom[®] | Precisely monitor the eluent flow |
| Dynamic Mixing Chamber | Effective homogenization of eluents Choose 1/16" version for flowrates up to 100 ml/min Choose 1/8" version for flowrates above 100 ml/min | Better performance |
| VariLoop | Variable injection volume and multiple injections | Adapt the sample volume to your application |
| Interface Box IFU 2.1 LAN | Highly precise analog data acquisition 4-channel input/output Sample rates of up to 50 Hz (one channel only) | Add any detector with analog output to your system |
| Pulse Damper | high damping performance membrane-free assembly easy implementation into the HPLC system | reduces pulsationimproves performance |



| | Benefit |
|---|---|
| eparative columns | Flexible operation with up to three columns |
| at space-limited sites, ns. | Space-saving solultion for AZURA system setup |
| ces and one fraction y 8 AZURA L devices | Mobile and compact arrangement of a prepara- tive HPLC system |
| or end of sample s per system ngs with 1/16" or ameter | Protect column from air damage and support automation (e.g. sample injection) |
| essure control, your interface box our AZURA L | Organize your system. |
| n 5 mm to con tubes, | Organize accessories directly at the system and reduce dead volume |
| precise direct os ne fluidic desig | Collect fractions while using your preferred detection method |
| RA L device holes apillaries | Sort the capillaries coming from the fraction collection valve |

AZURA® Compact Prep HPLC System

The AZURA® Prep Compact system is the perfect start into preparative chromatography. With the complete, semi-preparative HPLC system you master your isocratic purification tasks.



Compact Prep System

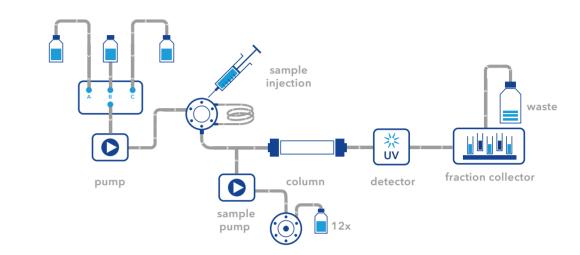
milligrams at up to 50 ml/min. Detection takes place via a versatile UV/VIS detector. The intuitive preparative software PurityChrom controls the compact its place in every laboratory.

One manual injection can purify several hundred system and regulates the fraction collection via a 12-port fractionating valve. Thanks to its compact design, the AZURA Prep Compact system finds

- Complete semi-preparative isocratic HPLC system with **low** space requirements
- Injection valve incl. 500 μl sample loop
- UV/VIS detector with one variable wavelength
- Intuitive **PurityChrom**[®] software
- Compact and expandable



Easy upgrade without big investment

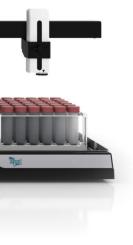


Pilot Prep System

After starting preparative chromatography with the space-saving prep system, the requirements for your purification tasks can quickly increase. The existing Compact System (50 ml/min) can be

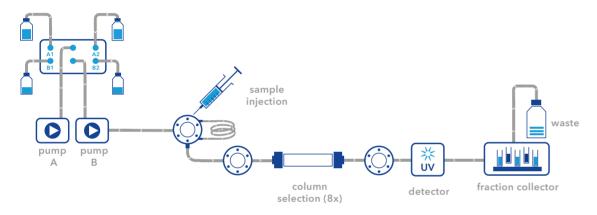


- expanded to a Pilot System (220 ml/min) by investing in a fraction collector and a preparative pump. All components of the Compact System are fully integrated into the Pilot System.
 - Pilot Ternary gradient HPLC system
 - Injection valve incl. 500 µl sample loop
 - Sample pump with automatic sample selection
 - UV/VIS detector with one variable wavelength
 - Intuitive **PurityChrom**[®] software
 - Fraction collector



AZURA® Lab Prep **HPLC System**

The Lab Prep LC system is designed for your more demanding semipreparative separations. You can customize a highly flexible LC system with the freely combinable components. With a maximum flow rate of 50 ml/min it is possible to separate up to several hundred milligrams per run.



- Lab Prep HPLC system with **binary high** pressure gradient
- Column selection
- Injection valve incl. 500 µl sample loop
- UV/VIS detector with one variable wavelength
- Intuitive **PurityChrom**[®] software
- Fraction collector



Method transfer from analysis of chamazulen to preparative scale

Chamomile plants are known for their medical properties, having among others anti-inflammatory, analgesic and sedative effects. These are due to the various phenolic compounds, one of them matricine is converted during the distillation process to chamazulene. The characteristic blue

Results

Fractionation/Purification

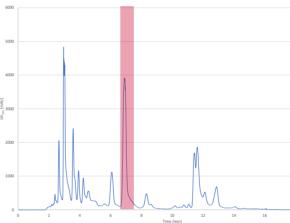
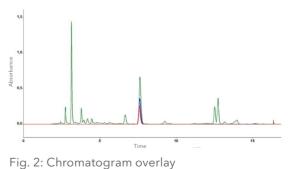


Fig. 1: Chromatogramm of preparative separation of chamazulene blue, collected fraction highlighted in red, 1 ml sample injection

Fraction analysis



Application Note (VPH0071)

Purification of chamazulene by preparative HPLC and its scale-up www.knauer.net/applications

- color of chamomile essential oils as "chamomile blue" is due to chamazulene. It has anti-inflammatory and anti-oxidant activity. The present application tested preparative HPLC to purify chamazulene from commercialy available "chamomile
- blue" oil.
- The separation of chamazulene was optimized in analytical scale and the two step gradient method transferred to preparative scale. Chamazulene purification was performed on C18 250x20 mm column, 25 ml/min. Fractionation of chamazulen was conducted by threshold function of Purity-Chrom software.
- The collected fraction was analysed by analytical HPLC and revealed nearly 100 % purity. Chromatogram overlay of the fraction, chamazulen standard and the sample clearly showed the succesfull purification of chamazulen (Fig. 2).
- Further the comparison of the fraction spectra and chamazulen spectra (Fig. 3) revealed that the purified fraction was chamazulen.

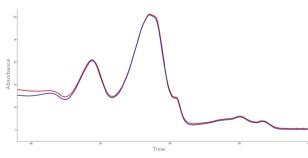
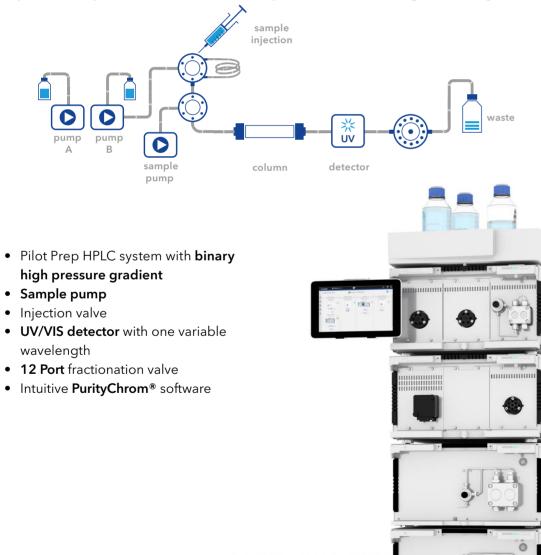


Fig. 3: Spectral view of fraction (-) and of chamazulen standard (-)



AZURA[®] Pilot Prep HPLC System

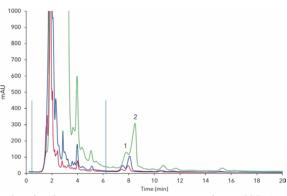
Choose the Pilot Prep LC system if you want to increase your productivity even more. As for the AZURA Lab Prep LC system you can freely build up your system. Flow rates up to 1000 ml/min and loads up to several grams are possible. Optional peak and solvent recycling can be set up to increase separation power and reduce separation costs significantly.



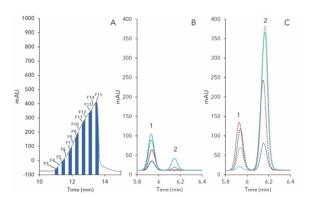
Improved purity by combining online SPE with preparative LC

Steviol glycosides are the main sweetening compounds in Stevia rebaudiana and are often used as natural sugar substitutes. To enable a commercial usage, the plant extracts need to be purified. In this work preparative online SPE (solid phase extraction) with the AZURA Pilot Prep LC was investigated for improvement of overall purity due to reduction of matrix contamination.

Results



Overload experiments on preparative column, 200 μ L (red), 500 μ L (blue), 2000 μ L (green); 1) rebaudioside A, 2) stevioside, blue bars - matrix, 25°C, 22 ml/min

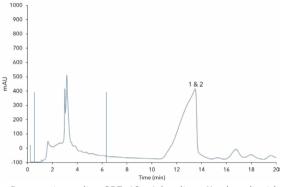


Fraction analysis of preparative online-SPE purification (Fig.2) of rebaudioside A (1) and stevioside (2); a) F3 (blue), F4 (red), F5 (green), F6 (light blue); b) F7 (red dashed), F10 (blue dashed), F12 (green sashed), F15 (light blue dashed); c) fractionation of target peak, 5 mL fractions

Application Note (VFD0171)

Evaluating preparative online SPE for the purification of stevia leave extracts **www.knauer.net/applications**

- The steviol glycoside rebaudioside A is the main
- d compound of interest as it is the sweetest and less
- bitter compound of the extract. Often Stevia prod-
- ucts contain a mixture of rebaudioside A and ste vioside. The development of a purification meth-
- od with high yield of rebaudioside A, only few
- stevioside impurities, and high throughput in-
- creases the economic output of Stevia production.



Preparative online SPE, 10 mL loading; 1) rebaudioside A, 2) stevioside, blue bars - matrix, 25°C, 22 mL/min

Fig. 1 shows the batch LC without online SPE. The matrix peak (1-5min) negatively affect the separation abilities. In comparision Fig 2 shows that the automated SPE process significantly decreased the matrix. The fraction analysis revealed that only a small part of the overlapping peak contained nearly pure rebaudioside A; fractions 3-5 approx. 15 mL with >90 % rebaudioiside A and <10% stevioside (Fig. 3, B). The later fractions contained high amounts of stevioside but also still rebaudioside A (Fig. 3 C). The results showed that purification of highly pure rebaudioside A is possible by an additional online-SPE.

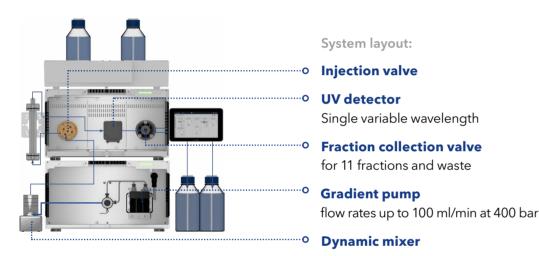
1) tion of stevia leave extracts

Preparative HPLC for cannabinoid purification

Whenever highly pure cannabinoids should be method to choose. Take a look at the exemplary produced from Cannabis plants or raw extracts, developed method to purify cannabidiol (CBD) preparative chromatography is the most versatile from CBD oil.

Cannabis Purifier

Purification of single cannabinoids from extracts of cannabis flos or cannabinoid oil.



Easy scale-up by changing the pump head, valves, flow cell, and capillaries. No exchange of devices necessary.

SMB for cannabinoid purification

like simulated moving bed (SMB) chromatography, depends on the specific customer's needs.

Whether to favor batch LC or a continuous process KNAUER has extensive experiences in customized solutions for both types of applications for the cannabis industry.

More information

For more information about analyzing and purifying Cannabis: www.knauer.net/cannabis



AZURA® SMB systems



What is the difference between batch LC and SMBC?

| Batch chromatography (single-column) |
|--------------------------------------|
| Unlimited number of fractions |
| Recovery typically below 80% |
| EITHER high purity OR high yield |
| lsocratic or gradient |
| High solvent consumption |
| Very diluted product |

Simulated moving bed chromatography (SMBC) is increasingly applied as a separation technique in the pharmaceutical industry, production of fine chemicals and in the field of bioengineering. SMB is a method in process chromatography that enables substance mixtures to be continuously separated and extracted in two fractions. By repeated use of the SMB process each partial fraction can be separated into a further fraction - down to binary substance mixtures.

Typically, the SMB process is set up in advance for a two component mixture. Following this, both substances can be immediately extracted in pure form.

For more information about SMB: www.knauer.net/smb

| SMB chromatography (multi-column) |
|--|
| Two fractions, no waste |
| Recovery up to 100% |
| High purity AND high yield |
| lsocratic |
| Can be as low as 10% of batch consumption |
| Product concentration comparable with input concentration (feed) |

Science with Passion



π. Worldwide partner in

science since 1962

includes systems and components for analytical

HPLC / UHPLC, preparative HPLC, fast protein

liquid chromatography (FPLC), multi-column

chromatography / simulated moving bed (SMB),

of the company since 2000. Several awards for

outstanding products and innovations as well

as entrepreneurial excellence make KNAUER a

KNALER

Based in Berlin, KNAUER is a medium-sized, owner-managed company that has been serving the sciences since 1962. We develop and manufacture scientific instruments of superior quality for liquid chromatography. The range



"leading employer".

and osmometry.

Independent and family owned

The founder Dr. Herbert Knauer and his wife Roswitha are still active as advisers in the company to this day. The couple's daughter, Alexandra Knauer, has been managing director and owner

THAT'S WHAT A CUSTOMER SAYS

"At Numaferm, we use proprietary recombinant technologies to identify, optimize and produce peptides. Our customers receive peptides of the highest quality at significantly reduced production costs, being produced sustainable. As an ISO 9001:2015 certified company, reliability and customer satisfaction are our top priorities. We have been working together with KNAUER for many years and successfully use the HPLC systems for purification."

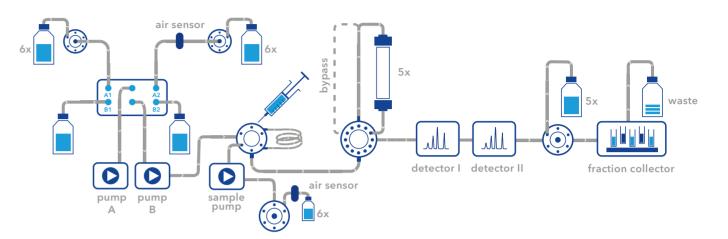




System configurator Preparative HPLC by KNAUER

MAKE YOUR PRESELECTION

Stainless steel



| SOLVENT SELECTION & DELIVERY | SAMPLE INJECTION | COLUMN S & THERMO |
|---|--|--|
| 50 ml/min binary gradient pump P 6.1L x 100 ml/min pump P 2.1L x 250 ml/min pump P 2.1L x 500 ml/min pump P 2.1L x 1000 ml/min pump P 2.1L Ternary gradient module for pump P 2.1L Binary gradient module for pump P 2.1L x solvent selection valve | Injection valve Sample pump module Sample selection valve: x inlets Autosampler AS 6.1L | Column (two col one byp Column high flov (5 colum bypass) |
| | | |

| x Airsensor main pump | x Airsensor feed pump | □ Mass flow |
|---------------------------------|---------------------------------|-----------------|
| x Tubing 1/16" | x Tubing 1/8" | x Tubing |
| SOFTWARE | | |
| □ ClarityChrom [®] | □ OpenLAB® | 🛛 PurityCh |
| □ Chromeleon™ | Mobile Control | |

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Biocompatible

| SELECTION | |
|-----------|--|
| OSTAT | |

| n selectior |
|-------------|
| olumns or |
| pass) |

Column selection high flow (5 columns, one bypass)

DETECTION

- UV/VIS single wavelength
- UV/VIS multiwave length
- DAD 2.1L
- □ Conductivity
- 🛛 рН
- □ Refractive index
- □ Light Scattering
- □ 4000 MiD
- A/D-converter (integration of further detectors)

FRACTION COLLECTION

- LABOCOL fraction collector with individual rack types
- □ KNAUER Liquid Handler 2.1
- Foxy fraction collector with fixed rack types
- □ Fractionation valve
- Rack for fraction collector
- □ Flow splitter

 ow controller
 AZURA Click
 AZURA Organizer

 g 1/4"
 Workstation (Windows)

 COMMON APPLICATIONS

 Chrom®
 Reversed phase
 Normal phase

 I other...
 System Qualification

KNAUER is the proud winner of the German Innovation Award 2022 in the category of medium-sized businesses.

(U)HPLC • FPLC • SMB • Osmometry and units for the production of lipid nanoparticles (LNP)

Y

Innovation

Own hardware and software development



Customized solutions

Pumps, detectors, valves and systems adapted to your needs

\bigcirc

Made in Germany

Independent and familyowned since 1962

think LC. think KNAUER.

KNAUER Wissenschaftliche Geräte GmbH

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