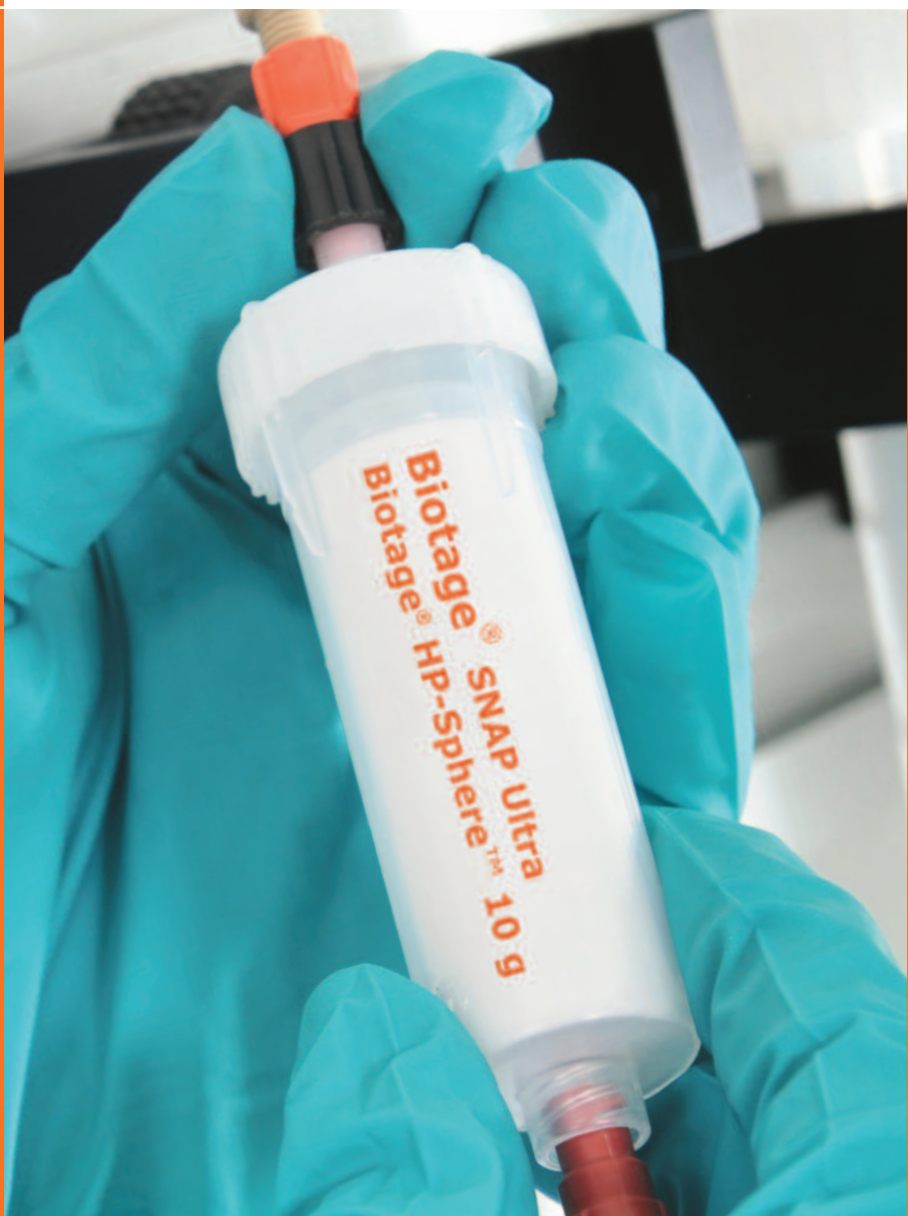


# Flash Purification Cartridges & Method Development



## Flash Cartridge / Instrument Compatibility Table

### Cartridges & Columns

Biotage® SNAP  
Biotage SNAP Ultra  
Biotage SNAP XL  
Biotage ZIP™  
Samplet®  
Dry Load Vessels  
Flash 75 and Flash 150  
Flash 400

### Accessories

TLC Plates  
Liquid Sample Injection  
Valves & Adapters  
Cartridge Adapters & Holders  
FlashPack Plunger Assemblies

### Stationary Phases for Flash Purification

### Flash Purification Method Development

Normal-Phase  
Reversed-Phase

# Flash purification cartridges for better, more reproducible separations

From routine to highly specialized applications, Biotage® has your solution

In 1994, Biotage was the first company to develop pre-packed cartridges for flash purification, and has been the leader in quality, performance and innovation ever since.

## Biotage Flash Purification

Flash purification is a technique developed by Professor W.C. Still that uses a column or cartridge filled with an insoluble solid support (stationary phase) and elution solvent mixture (mobile phase) to separate and purify a mixture of organic compounds. The stationary phase and the mobile phase typically have very different polarities, which work in tandem to separate compound mixtures. The separated molecules can then be collected in a purified state for use in a subsequent synthesis or as a final product.

In 1994, Biotage was the first to develop and introduce pre-packed cartridges for flash purification. A broad selection of distinct cartridge styles now enable professionals to choose the cartridge which best suits the purification need and purification system:

<b>Biotage SNAP</b>	<b>Flash 75</b>
<b>Biotage SNAP Ultra</b>	<b>Flash 150</b>
<b>Biotage SNAP XL</b>	<b>Flash 400</b>
<b>Biotage ZIP</b>	

All Biotage cartridges are designed to meet the requirements of HPFC (high-performance flash chromatography) systems including Biotage Isolera Prime™, Biotage Isolera™ One, Four and LS, Biotage SP1, Biotage SP4, and Biotage FlashMaster. Automated cartridge-packing techniques ensure efficiently packed Biotage flash cartridges minimize performance variability. Cartridge sizes range from 5 g to 40+ kg for purification scales ranging from milligrams to kilograms. Manufactured from polypropylene to meet the extractable requirements in US 21 CFR 177.1520 and packed using proprietary methods that strictly adhere to ISO 9001:2000 quality standards, Biotage cartridges also undergo QC testing and production control. Each Biotage SNAP cartridge is laser-etched with a unique lot number to ensure traceability.

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## Flash Cartridge/Instrument Compatibility

Cartridge	Media Mass	Automated Systems				Manual Systems		Scale-up Systems			
		Biotage Isolera™ One	Biotage Isolera Four	Biotage Isolera LS	Biotage Isolera Prime™	Biotage FlashMaster Personal	Biotage FlashMaster PersonalPlus	Biotage Isolera LS	Flash 75	Flash 150	Flash 400
<b>Biotage SNAP and Biotage SNAP Ultra</b>	10 g	●	●		●	●	●				
	25 g	●	●		●	●					
	50 g	●	●	●	●	●	●	●			
	100 g	●	●	●	●	●	●	●			
	340 g	●	●	●	●	●	●	●			
<b>Biotage SNAP XL</b>	750 g	●	●	●	●			●	●	●	
	1500 g			●				●	●	●	
<b>Biotage ZIP™</b>	5 g	●	●		●	●	●				
	10 g	●	●		●	●	●				
	30 g	●	●		●	●	●				
	45 g	●	●		●	●	●				
	80 g	●	●	●	●	●	●	●			
	120 g	●	●	●	●	●	●	●			
											For more information on Biotage flash purification instruments please see our Flash Systems brochure.
<b>Flash 75S</b>	200 g	●	●	●				●	●		
<b>Flash 75M</b>	400 g	●	●	●				●	●		
<b>Flash 75L</b>	800 g	●	●	●				●	●		
<b>Flash 150M</b>	2.5 kg			●				●		●	
<b>Flash 150L</b>	5 kg			●				●		●	
<b>Flash 400M</b>	20 kg									●	
<b>Flash 400L</b>	40 kg									●	

● = Attaches directly

● = Requires external cartridge stand or adapter



## Biotage® SNAP

Flexibility, performance, and high-value

The Biotage SNAP family of flash cartridges is the recognized industry standard for professionals worldwide. Available in a wide range of sizes for practically any situation, Biotage SNAP cartridges deliver the purification efficiency and favorable separation time to meet the growing demands of a global economy.

The Biotage SNAP cartridge design enables seven types of loading techniques including three internal dry loading options. They are packed to provide the highest resolution of any 50 µm particle cartridge on the market providing increased loading capacity and better resolution. Biotage SNAP cartridges are constructed using USP Class VI plastics (medical-grade) for lower extractables and cleaner fractions.

Biotage SNAP cartridges are ready-to-use flash columns designed to withstand 100 psi (7 bar) without the use of compression modules. Optimal cartridge packing protocols produce cartridges that deliver larger loading capacities, tighter elution bands, and higher purity fractions. Biotage SNAP cartridges are tested to ensure they meet stringent performance standards including efficiency (plate count) and peak shape (symmetry).

Available in sizes from 10 g to 340 g with the choice of silica, C18, or NH functionalized media, Biotage SNAP cartridges will purify your valuable compounds whether you have low mg samples or multi-gram samples.

For maximum sample loading and the best separation, dry loading the sample using Biotage SNAP Samplet® cartridges gives the best performance by eliminating any dissolution solvent effects.

*See page 8 for more information on Biotage SNAP Samplet cartridges.*

## Advantages

- Universal fit onto all flash systems
- 3000 N/M minimum performance provides narrower elution bands increasing sample load
- Up to 100 psi / 7 bar pressure rating enables faster flow rates and use with viscous solvents
- 20% more loading capacity than competitive cartridges
- Higher peak resolution than competitive cartridges
- Luer-lock connections eliminate adapters and provide compatibility with most flash systems
- Removable cap allows internal sample loading
- 3000 N/M minimum performance provides narrower elution bands increasing sample load
- Up to 100 psi / 7 bar pressure rating enables faster flow rates and use with viscous solvents
- Translucent barrel provides assurance that solvent is flowing and separation is occurring
- USP Class VI construction materials minimize extractables providing cleaner purified products (**Figure 1**)
- Removable cap with extra column head space allows Samplet and bulk dry-loading to improve purification performance
- Three standard silica-based media provide selectivity choices for optimal purification
- Pre-packed Samplet cartridges increase loading capacity and produce tighter elution bands by removing the injection solvent effect
- Individually lot numbered for traceability

## Available Media

- Biotage KP-Sil — 50 µm
- Biotage KP-C18-HS — 50 µm
- Biotage KP-NH — 50 µm

## Optional Accessories

- Dry loading vessels
- 3-way injection valves
- Samplet cartridges
- Adapters

## Recommended Instruments

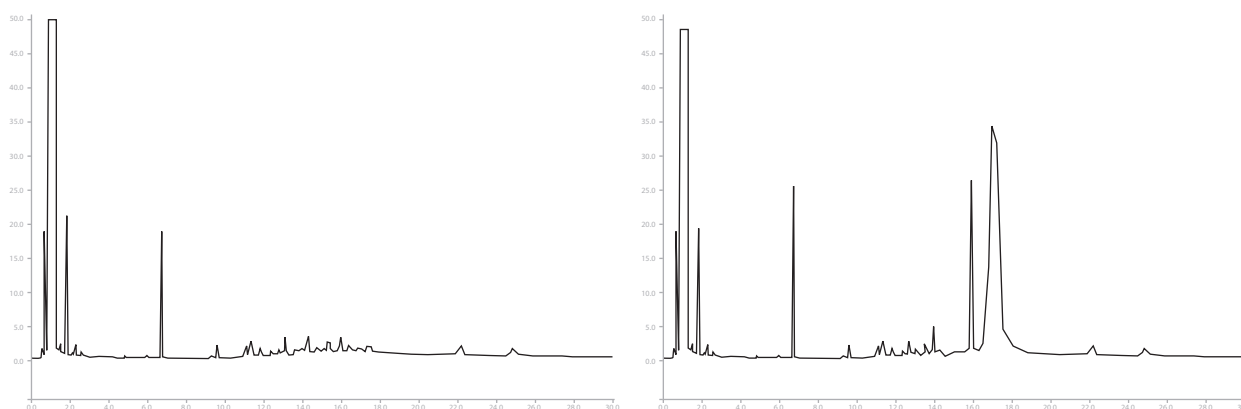
- Biotage Isolera™ flash purification family
- Compatible with instruments requiring Luer-lock connections

See **Flash Cartridge/Instrument Compatibility Chart** for the Biotage SNAP product that best fits your flash purification needs.

## Cartridge Selection / Performance

Biotage SNAP	10 g	25 g	50 g	100 g	340 g*	750 g	1500 g	Efficiency (N/m)	Symmetry
<b>Normal-phase</b> (Biotage KP-Sil)	1	1	1	1	1	1	1	≥3000	0.7–1.5
<b>Reversed-phase</b> (Biotage KP-C18-HS)	1	1	1	1	1	1	1	≥3000	0.7–1.5
<b>Amine Purification</b> (Biotage KP-NH)	1	1	1	1	1	1	1	≥3000	0.7–1.5

\*Maximum pressure of 75 psi (5 bar)



**Figure 1.** Biotage SNAP cartridges, manufactured from materials meeting USP Class VI extractables requirements, reduce the amount of extractables and leachables generated by cartridge construction materials. The gas chromatography analysis above show the results of DCM washes (4 liters) of a Biotage SNAP 1500 g cartridge (left) and a competitive 1500 g cartridge (right). Lower extractables ensure higher purity fractions.



## Biotage® SNAP Ultra

### The highest purification performance

Biotage SNAP Ultra flash chromatography cartridges deliver the highest purification performance available with the highest column loading capacity utilizing Biotage HP-Sphere™, spherical silica delivering 40% more surface area.

#### The highest performance available

Precision engineered Biotage SNAP Ultra cartridges deliver double the purification capacity of other flash cartridges by utilizing Biotage HP-Sphere, small particle, 25 µm spherical silica with a 40% increase in surface area. The higher surface area provides twice the loading capacity of lower surface area silicas. This improved load capacity means that a smaller SNAP Ultra cartridge can be used to replace a more expensive and larger competitive cartridges.

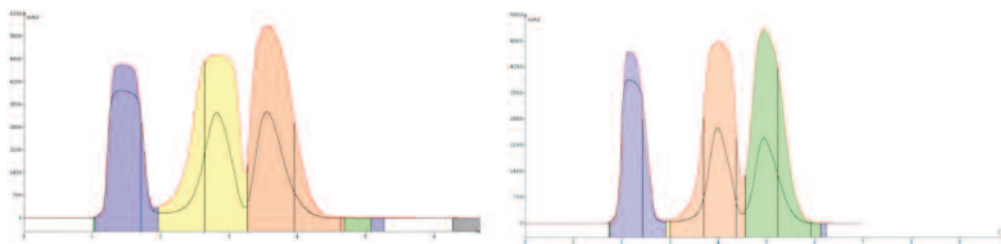
The small, 25 µm spherical particles minimize band-spreading and back-pressure enabling the use of faster flow rates and steeper gradients to speed purification and improve productivity.

#### Designed for the Biotage Isolera™ and other flash chromatography systems

Isolera software and Biotage SNAP Ultra cartridges work together as a system to produce pure compounds the first time. Luer lock inlet and outlet ports ensure robust connection to the Biotage Isolera or any other flash chromatography system.

#### Wide range of sizes for scalability

Biotage SNAP Ultra cartridges are available from 10 g to 340 g sizes providing simple and direct scale-up of samples from low milligrams to multi-grams



**Figure 2.** The very high surface area of the spherical, 25 µm Biotage HP-Sphere silica enables the use of smaller Biotage SNAP Ultra cartridges in place of a larger flash cartridge. In this example, a Biotage SNAP Ultra 50 g cartridge (far left) actually purifies more sample than a competitive 80 g high-performance cartridge saving 38% in solvent use and run time

## Advantages

- Biotage HP-Sphere, spherical silica with 40% more surface area delivers the highest sample loading available to reduce purification costs
- Smaller, 25 µm silica delivers the best peak resolution and higher concentration fractions reducing solvent evaporation time
- Removable cap provides internal sample loading, both liquid and dry to improve purification performance
- Samplet® cartridge compatibility to pre-concentrate samples and eliminate solvent dissolution
  - Pre-packed Samplet cartridges increase loading capacity and produce tighter elution bands by removing the injection solvent effect
- Low back pressure enables faster flow rates and use with viscous solvents
- Medical-grade polypropylene reduces leachables that can contaminate purified compounds
- Individually lot numbered for traceability
- Designed to be the perfect complement to Biotage Isolera chromatography systems
- Universal fit onto all flash systems with Luer inlet and outlet connections

## Specifications

<b>Materials</b>	Tubes: inert, medical-grade polypropylene (USP Class VI) Frits: inert polyethylene
<b>Pressure limit</b>	100 psi (7 bar) 340 g max pressure 75 psi (5 bar)
<b>Resolution</b>	Minimum 7000 N/m (plates per meter)
<b>Quantity</b>	20/case, 6/case for 340 g
<b>Compatibility</b>	Universal Luer connections
<b>Sizes</b>	10, 25, 50, 100 and 340 g cartridges

## Recommended Instruments

- Biotage Isolera™ flash purification family
- Compatible with instruments requiring Luer-lock connections

## Optional Accessories

- Dry loading vessels
- 3-way injection valves
- Samplet® cartridges
- Adapters

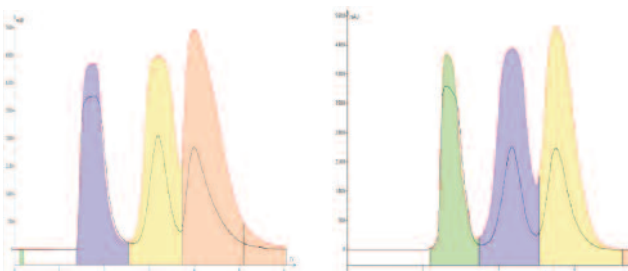
See **Flash Cartridge/Instrument Compatibility Chart** for the Biotage SNAP Ultra product that best fits your flash purification needs.

Performance like this means that cartridges can be loaded with twice the amount of sample to be purified. That means you can use a cartridge that is half the size that you would normally use. A smaller, high surface area cartridge takes less time and less solvent to do the job of a cartridge twice the size.

- Higher loading per gram of silica means smaller cartridges can be used to perform any separation
- Smaller cartridges use less solvent and require less time
- Chemists produce more compounds in less time
- Less solvent use also means less generated waste – a major ecological improvement

	Biotage ZIP™	SNAP	SNAP Ultra
Features			
Performance			
Price			

Biotage® Flash Cartridge Comparison



**Figure 3.** Biotage SNAP Ultra silica has 40% higher surface area than other spherical flash silica and delivers increased loading capacity. At 4% load (4 grams sample per 100 grams of silica) a 25 g Biotage SNAP Ultra cartridge (far left) provides an equivalent separation as a competitive 25 g, 30 µm silica cartridge with a 2% sample load.



## Biotage<sup>®</sup> SNAP XL

### Cartridges for development-scale purification

SNAP 750 g and 1500 g cartridges address a growing need for optimized gram-to-150+ g-scale purification, increasing purification efficiency while reducing costs.

SNAP XL cartridges are rugged and laser welded for safe operation at pressures up to 100 psi, enabling higher flow rates and the use of higher viscosity solvents.

#### **High Productivity**

Operating at flow rates up to 500 mL/min, Biotage SNAP XL cartridges allow you to quickly purify large sample masses, saving hours or even days of purification runtime.

#### **Direct Scale-up**

With greater than 10% sample loading capacity possible, Biotage SNAP XL cartridges provide seamless scale-up from milligrams to >150 g simply and efficiently without method modification see (**Figure 4**).

Biotage SNAP XL cartridges are available in three media – Biotage KP-SIL, Biotage KP-C18-HS, and Biotage KP-NH enabling direct scale-up from smaller Biotage SNAP cartridges.



## Advantages

- Laser welded for strength and safety
- High loading capacity increases purification throughput
- Seamless scale up from other Biotage flash cartridges
- Up to 100 psi / 7 bar pressure rating enables faster flow rates
- Translucent barrel provides assurance that solvent is flowing and separation is occurring
- USP Class VI construction materials minimize extractables providing cleaner purified products

## Available Media

- Biotage KP-Sil — 50  $\mu$ m
- Biotage KP-C18-HS — 50  $\mu$ m
- Biotage KP-NH — 50  $\mu$ m

## Recommended Instruments

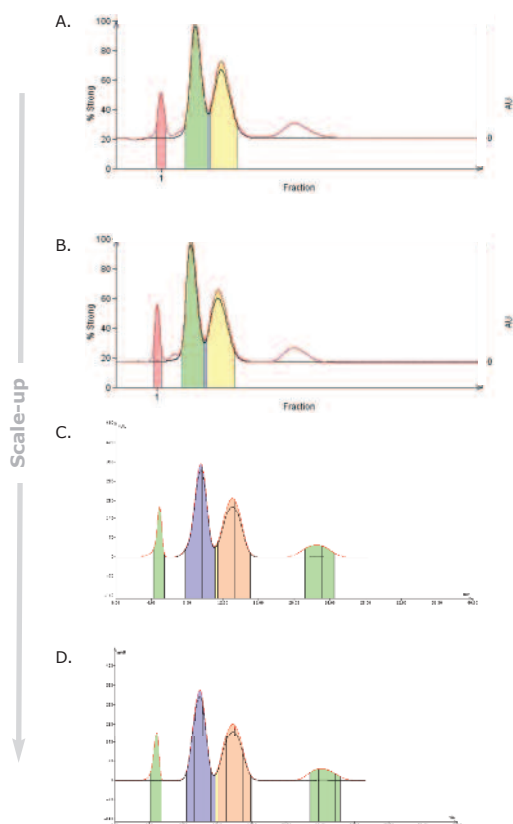
- Biotage Isolera™ One
- Biotage Isolera Four
- Biotage Isolera LS
- Compatible with most manufacturers' development-scale instruments

## Optional Accessories

- 3-way, large-bore injection valve
- Inlet and outlet Luer fittings
- Cartridge holder for Biotage Isolera systems
- External dry load vessel (DLV-500)

## Cartridge Selection / Performance

Biotage SNAP XL	750 g	1500 g	Efficiency (N/m)	Symmetry
<b>Normal-phase</b> (Biotage KP-Sil)	1	1	3000	0.7–1.5
<b>Reversed-phase</b> (Biotage KP-C18-HS)	1	1	3000	0.7–1.5
<b>Amine Purification</b> (Biotage KP-NH)	1	1	3000	0.7–1.5



**Figure 4.** Successful scale-up using Biotage SNAP KP-C18-HS 12 g, 60 g, 950 g, and 1850 g cartridges shows direct scalability even with closely eluting compounds.

(A.) 12 g Biotage SNAP C18 1% load mixed probes 10 mL/min 60:40 MeCN:H<sub>2</sub>O

(B.) 60 g Biotage SNAP C18 1% load mixed probes 50 mL/min 60:40 MeCN:H<sub>2</sub>O

(C.) 950 g Biotage SNAP XL C18 1% load mixed probes 250 mL/min 60:40 MeCN:H<sub>2</sub>O

(D.) 1850 g Biotage SNAP XL C18 1% load mixed probes 500 mL/min 60:40 MeCN:H<sub>2</sub>O



## Biotage ZIP™

### Simplicity, performance and high-value

Biotage ZIP flash cartridges are an evolution in Biotage's long history of manufacturing flash purification solutions. Biotage ZIP cartridges deliver industry-leading performance that meet the difficult challenges of day to day purification. Plug and play Biotage ZIP cartridges are the perfect compliment to all Biotage Isolera™ flash chromatography systems.

#### High performance at a value price

Biotage ZIP cartridges are engineered to meet the needs of chemists requiring high performance and consistency from a purification cartridge to ensure delivery of pure compounds. Biotage ZIP cartridges are laser-welded to ensure leak-free operation even at elevated pressures.

#### Plug and play design

Constructed with industry-standard Luer lock inlet and Luer tip outlet ports, Biotage ZIP cartridges are truly universal in their use with flash systems from around the world. No adapters are needed, so installation is quick and easy.

#### Wide range of sizes and scalable

Biotage ZIP cartridges are packed with Biotage KP-Sil brand silica which is used in all Biotage silica flash cartridges. Available in sizes from 5-120 g, Biotage ZIP cartridge purifications can be scaled up to larger Biotage cartridges including Biotage SNAP 340 g, 750 g and 1500 g, Flash 75 and 150, and even Flash 400 without method modification.

## Advantages

- 20% more loading capacity than competitive cartridges
- Up to 150 psi / 10 bar pressure rating enables faster flow rates and use with viscous solvents
- Translucent barrel provides assurance that solvent is flowing and separation is occurring
- Consistent performance provides reliable results
- Laser-welded inlet fitting provides leak-free operation and high pressure tolerance
- Luer tip outlet is compatible with all flash systems
- Automated manufacturing process provides better cartridge reliability, reproducibility and performance over lesser quality competitive products

## Optional Accessories

- Dry loading vessels
- 3-way injection valves
- Adapters

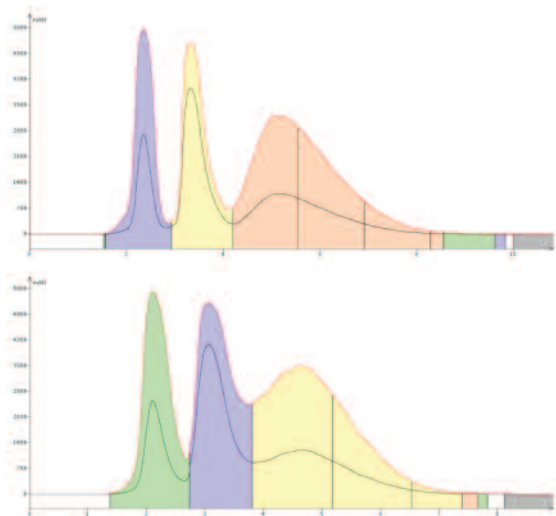
## Specifications

<b>Materials</b>	Tubes: inert polypropylene Frits: polyethylene 40-63 µm Biotage KP-Sil
<b>Pressure limit</b>	Stable to 150 psi (10 bar) 80 g max pressure 120 psi (8 bar)
<b>Quantity</b>	Packages of 20
<b>Compatibility</b>	Universal Luer connections, compatible with all flash systems
<b>Sizes</b>	5, 10, 30, 45, 80 and 120 g cartridges

## Recommended Instruments

- Biotage Isolera™ flash purification family
- Compatible with instruments requiring Luer connections

See **Flash Cartridge/ Instrument Compatibility Chart** for the Biotage ZIP product that best fits your flash purification needs.



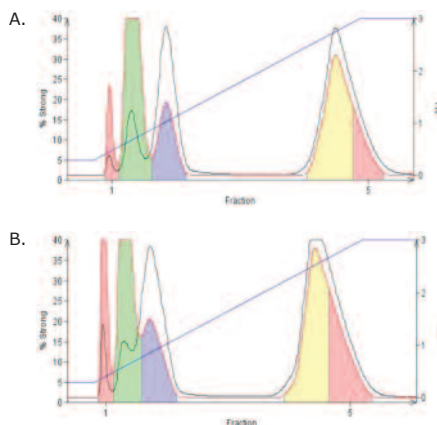
**Figure 5.** A 30 g Biotage ZIP cartridge (top) and a competitive 30 g flash cartridge with spherical silica (bottom) are used to isocratically separate a 3-component mix. The Biotage ZIP cartridge provides more compound resolution and sharper peaks for the same sample load.



## Samplet® Cartridges

### Internal dry loading improves separations

Samplet cartridges are developed for quick and convenient sample introduction acting as sample pre-concentrators and guard cartridges. After the sample has been applied to the Samplet cartridge and the solvent evaporated, the Samplet cartridge is inserted into the flash cartridge, providing a dry, concentrated sample for purification. Dried, adsorbed samples enhance the separation and improve compound recovery, purity, and loading capacity (**Figure 6**). Samplet cartridges are available for both Biotage® SNAP and FLASH+® cartridges.



**Figure 6.** Increased performance of a Samplet-loaded cartridge (A) versus a liquid-loaded cartridge (B) is seen by the improved peak resolution. By allowing the sample to dry in the Samplet cartridge, the sample diluent's strong displacement effects are eliminated and improved purification is achieved.



DLV-030

DLV-500

## Dry Loading Vessels

### Improve purification results with higher sample loads

One of the most common flash purification challenges is dealing with hard-to-dissolve crude samples. Because polar solvents cause poor chromatographic results when used as injection solvents in normal-phase flash chromatography, other sample load options are needed.

A commonly used option is dry loading, which involves dissolving the sample in a suitable solvent, mixing with an inert, clean, dry adsorbent, such as silica, alumina, or diatomaceous earth, and drying the slurry. The dried sample is then loaded into an empty vessel and inserted ahead of the purification cartridge. By drying the mixture, the polar solvent is removed and will have no impact on the purification. Dry load vessels are available for use with all cartridges. For Flash 75 and Flash 150, repackable Biotage SIM modules in sizes from 100 mL to 2 L are available.

#### Advantages

- Improved sample separation
- Higher loading capacity
- Increased fraction purity

#### DLV Accessories

- Replacement barrels and frits
- DLV holders

#### Biotage SIM Accessories

- Replacement frits
- Replacement filter

## Accessories



### Thin-Layer Chromatography (TLC) Plates

TLC is a commonly used method development tool for flash purification. Product mixtures separated using TLC generally can be purified by flash chromatography using the identical solvent system. TLC to flash method transfer accuracy is improved when TLC plates and flash cartridges made with identical silica from Biotage are used. For synthetic chemists, the benefits of matched TLC plates and flash cartridges are better purification throughput, increased compound purity and yield, and reduced solvent cost.



### Liquid Sample Injection Valves and Adapters

For liquid samples, Biotage offers two 3-way injection valves that attach directly to Biotage cartridges and compression modules. These stainless steel valves come complete with finger tight fittings and a Luer adapter for syringe injection. The straight-through injection design minimizes wash volume and minimizes precipitation potential.



### Cartridge Holders

Designed for simple attachment onto Biotage Isolera™ systems, these cartridge holders accommodate 10 g, 25 g, 50 g, 100 g, 340 g, 750 g and 1500 g Biotage SNAP cartridges. A FLASH+ cartridge holder is also available.



### Cartridge Adapters

Luer-lock to Luer tip adapters enable Biotage SNAP cartridges to be used with Biotage FlashMaster, Isco and other competitive flash systems. The very easy to install adapter attaches directly to the Biotage SNAP cartridge outlet converting the Biotage SNAP cartridge from Luer lock to Luer tip.



### FlashPack Plunger Assemblies

ISOLUTE® flash chromatography columns use a plunger assembly for compatibility with Biotage FlashMaster flash chromatography systems. The plungers are adjustable to accommodate virtually any bed mass including dry loaded samples.



## Flash 75 and Flash 150

The industry standard original for development-scale cartridges

The original flash cartridges invented by Biotage® deliver excellent purification, ease of use and are available in 75 mm and 150 mm ID. Additionally, these development-scale cartridges provide a straightforward purification scale-up path from other flash cartridges.

Routinely operating at a flow rate of 250 mL/min, Flash 75 systems and Flash 75 cartridges allow you to scale-up and quickly complete runs, saving hours or even days of purification time compared to using glass-column flash purification.

### Radial Compression Improves Separations

Flash 75 and 150 cartridges operate in radially-compressed barrels that squeeze the cartridge walls improving bed density and minimizing the chance of voiding and channeling

### Reliable Gram-scale Purification

Flash 75 and 150 cartridges provide a direct scale-up pathway from smaller scale Biotage flash cartridges. Five cartridge sizes — 200 g, 400 g, and 800 g (Flash 75), and 2.5 kg and 5 kg (Flash 150) enable purification from tens of grams to hundreds of grams of crude reaction mixture.

### Media Options

Flash 75 and 150 cartridges are available with many media choices to enable direct scale-up from smaller-scale Biotage cartridges and chemisorption resins. Choose from Biotage KP-Sil, Biotage KP-C18-HS, Biotage KP-NH, activated carbon, HP20, or HP20SS.

## Advantages

- Compression modules radially compress Flash 75 and Flash 150 cartridges to maximize sample contact with silica (higher sample load) and separation performance (greater purity, recovery)
- Compression modules seal up to 100 psi, ensuring leak-free operation even with high flow rates and reversed-phase solvents
- Knife-edge sealing mechanism in the Flash 75 and Flash 150 compression modules seals to 100 psi without o-rings

## Recommended Instruments

- Flash 75
- Flash 150

## Optional Accessories

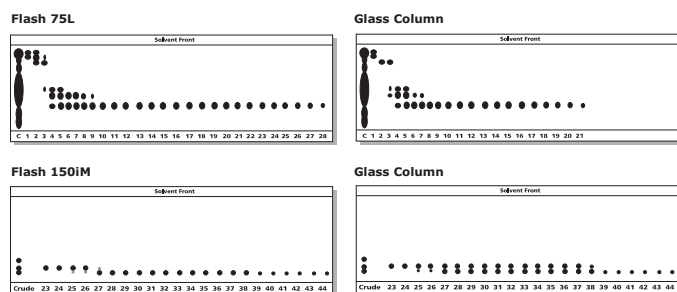
- Biotage SIMs (Sample injection module)
- 3-way injection valve

## Cartridge Selection

Development Scale	Biotage KP-Sil	Biotage KP-C18-HS	Flash-AC	Diaion® HP20	Diaion HP20SS
75 mm	1	1	1	1	1
150 mm	1	1	1	1	1

## Specifications

	Diameter (mm)	Bed Length (cm)	Packing Weight (g) (nom)	Column Volume (mL)	Flow Rate (mL/min)	Easy $\Delta CV = 4+$	Typical $\Delta CV = 2-3.9$	Difficult $\Delta CV = 0.1-1.9$
Flash 75S	75	9	200	320	250	5-10	1-5	0.2-1
Flash 75M	75	15	400	535	250	10-20	2-10	0.4-2
Flash 75L	75	30	800	1070	250	20-40	4-20	0.8-4
Flash 150M	150	300	2500	4300	500-1000	80-160	16-80	3-16
Flash 150L	150	600	5000	8600	500-1000	160-320	32-160	6-32



**Figure 7.** The TLC of the collected fractions in the Flash 75L and 150M vs. glass column comparisons show more pure fractions were collected vs. the glass column. That improvement along with a faster flow rate yielded a throughput enhancement of two times over the glass column.

Routinely operating at a flow rate of 250 mL/min, the Flash 75 systems and Flash 75 cartridges allow you to scale-up and quickly complete runs, saving hours, even days, of purification time compared to using glass-column flash purification.

## Flash 150 vs. Glass Comparison

	Flash 150M	Glass Column
Column Size (mm)	150 x 300	120 x 660
Silica Volume (kg)	2.5	3
Sample Load (g)	180	450
Flow Rate (mL/min)	500	70
Number of Fractions	45	30
Purification Time (min)	90	430
Pure Compound (g/run)	70.5	45
Recovery (%)	87.0	22.2
Purification Throughput (g/hr)	120	63

In this application the Flash 150 system saved nearly four weeks of purification time on a 1 kg scale synthesis project.



### Advantages

- Can purify up to 4+ kg of crude reaction mixture
- Minimize exposure to highly active or toxic compounds
- Improves safety by eliminating the need to handle kilogram amounts of media
- Shipped in a re-sealable container for disposal and incineration

### Recommended Instruments

- Flash 400

### Available Media

- Biotage KP-Sil
- Biotage KP-C18-HS (reversed-phase)
- Biotage KP-NH (amine-functionalized)
- Mitsubishi Diaion® HP20 and HP20SS
- Activated carbon

## Flash 400

### The industry standard original for process-scale purification

Biotage® Flash 400 cartridges are pre-packed cartridges (400 mm ID) designed for large-scale chromatography, and radially compressed to provide maximum performance and reliability. Flash 400 cartridges are faster, safer and easier to use than traditional glass and steel columns. Built to last and engineered to perform, Flash 400 systems and cartridges are the first choice of companies around the world for critical adsorption purification.

*For more information on Flash 400 systems, please refer to the Biotage flash systems brochure.*

Flash 400 systems have interchangeable barrels for use with either 400 mm x 30 cm (Flash 400M) or 400 x 60 mm (Flash 400L) cartridges. A system includes a compression module of one size, and the module of the other size can be ordered as an interchangeable option.

- Ensures highest purity, yield and throughput
- Designed for both chromatographic purification and adsorption processing of a variety of organic synthetic and natural products
- Minimizes exposure to highly active or toxic compounds
- Available with normal- and reversed-phase silica, activated carbon, polymeric adsorbents or custom-packed media

### Specifications

	Flash 400M	Flash 400L
Diameter (mm)	400	400
Bed length (cm)	30	60
Flow rate (L/min)	1–7	1–7
Column Void Volume (L)	28	56
Biotage KP-Sil Silica Packing Weight (kg)	20	40
Biotage KP-C18-HS Packing Weight (kg)	24	48
Flash-AC Activated Carbon Packing Weight (kg)	13	26
Mitsubishi HP20 Packing Vol (L)	38	76



# Stationary Phases for Flash Purification

## Normal-phase Purification

Most organic synthesis products are organic solvent soluble — e.g. dichloromethane (DCM), acetone, ethyl acetate (EtOAc), etc. — and are lipophilic. Lipophilic molecules are typically separated using a polar stationary phase and a non-polar mobile phase. This technique is called normal-phase purification. Common mobile phases for normal-phase purification include hexane/EtOAc and DCM/MeOH. DCM/MeOH is primarily for organic molecules with highly polar functional groups (acids, amines, alcohols), which have a higher affinity for silica than esters, amides, ethers, etc.

In normal-phase purification, Biotage cartridges packed with Biotage KP-Sil, Biotage HP-SIL, or Biotage KP-NH are typically used.

## Four main purification modes available with Biotage cartridges

### Normal-phase

#### Biotage HP-Sphere™

Biotage HP-Sphere is a true high-performance silica. Its spherical shape, 25 µm particle size, and extremely high surface area combine to make Biotage HP-Sphere the most efficient flash silica available. Its spherical shape reduces backpressure while the 25 particle diameter and 700 m<sup>2</sup>/g surface area combine to provide maximum resolving power and sample loads twice that of other silicas.

#### Biotage KP-Sil and Biotage HP-SIL

The most frequently used silica for flash purification features high surface area (500 m<sup>2</sup>/g), moderate porosity (60 Å), a tight uniform particle distribution (40–63 µm), neutral pH, and low metals content. These factors combine to provide high loading capacity, efficiency, and reproducibility. Biotage HP-SIL in a smaller particle size range (20–30 micron average) is made of exactly the same silica as the larger particle Biotage KP-Sil. Methods developed on Biotage KP-Sil can easily be transferred to the higher resolution, smaller particle Biotage HP-SIL silica.

### Normal-phase — amine functionalized

#### Biotage KP-NH

Biotage KP-NH chemistry shields synthetic organic amines from acidic silanols providing improved selectivity, peak shape, purity and yield. Unlike traditional silica and 1° amine (propyl amine) bonded silica, Biotage KP-NH does not require the use of chlorinated solvents or amine additives. Biotage KP-NH flash cartridges and matching TLC plates separate 2°, 3°, and heterocyclic amines using non-chlorinated solvents. Biotage KP-NH TLC plates are made using the same chemistry as Biotage KP-NH flash cartridges. Methods developed using Biotage KP-NH TLC plates accurately transfer to Biotage KP-NH flash cartridges simplifying flash purification.

### Physical Properties of Biotage Flash Chromatography Media

	Chemistry	Particle Size (µm)	Surface Area (m <sup>2</sup> /g)	Pore Volume (mL/g)	Pore Diameter (Å)
Biotage HP-Sphere	Silica	25	700	0.9	50
Biotage KP-Sil	Silica	50	500	0.7	60
Biotage KP-C18-HS	C18	50	400	0.95	90
Biotage KP-NH	amine	50	230	0.6	100
ISOLUTE® NH2	1° amine	50	500	0.7	60
HP20	S-DVB	250–600	600	1.3	300–600
HP20SS	S-DVB	75–150	600	1.3	300–600
Activated Carbon	Carbon	45–105	—	—	—

## Reversed-phase Purification

Some synthetic mixtures and natural products are soluble in more polar solvents such as MeOH, DMSO, DMF, MeCN, or water. Polar solvents are very disruptive to normal-phase mass-transfer kinetics and should only be used with reversed-phase purification. Reversed-phase purification uses a hydrophobic stationary phase with a polar mobile phase. Reversed-phases consist of a hydrocarbon, typically 18 carbons long, covalently bonded to silica. The bonding of this moiety creates a highly lipophilic environment, separating compounds on the basis of hydrophobic partitioning, similar to liquid-liquid extraction.

Unlike silica cartridges, reversed-phase cartridges can be re-used many times.

### Reversed-phase

#### Biotage KP-C18-HS

Reversed-phase flash chromatography is a very effective purification technique. Its main application areas include polar, ionizable and highly lipophilic compounds which cannot easily be separated by normal-phase techniques. Unlike normal-phase chromatography, reversed-phase uses a hydrophobic stationary phase (e.g. C18 or ODS) and hydrophilic mobile phases (methanol/water, acetonitrile/water). By converting silica's active, polar silanols sites to neutral, lipophilic sites, compounds that will either aggressively stick to silica or not stick at all can be retained, separated and eluted using water-based solvent systems.

### Chemisorptive

#### Diaion HP20, HP20SS

HP20 and HP20SS are styrene-divinyl benzene copolymers primarily used as traps for organic molecules from aqueous fermentation solutions. HP20 has a particle size of 250–600 µm, while HP20SS has a particle size of 75–150 µm. Each has a high 600 m<sup>2</sup>/g surface area, which allows a large concentration of organic materials to accumulate from aqueous solutions.

### Activated carbon

For color, by-product, catalyst, or liposaccharide (LPS) pyrogen removal from active compound solutions prior to crystallization Biotage offers activated carbon packed into cartridges. Use of pre-packed cartridges is generally safer and more efficient than batchmode scavenging. Biotage activated carbon is scalable to Flash 400 and can be used in a cGMP environment when API cleanliness is required.

# Flash Chromatography Method Development

This guide covers the stages of optimizing flash chromatography for three popular types of flash purification methods — isocratic normal-phase, gradient normal-phase, and reversed-phase. These guidelines address important issues related to achieving successful flash purification.

## Normal-phase

- Stage 1:** Converting TLC (thin-layer chromatography)  $R_f$  (retention factors) to CV (column volumes)
- Stage 2:** Determining the best solvent selectivity using TLC
- Stage 3:** Determining the best solvent strength using TLC
- Stage 4:** Determining the optimum cartridge size and sample load based on TLC data

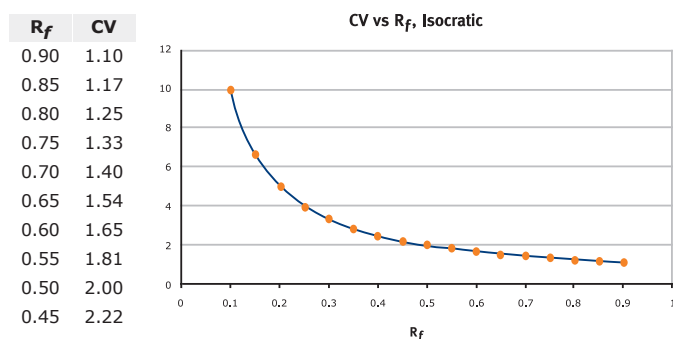
## Reversed-phase

- Stage 1:** Converting HPLC (high-performance liquid chromatography) retention times and gradient methods to CV
- Stage 2:** Determining the optimum cartridge size and sample load based on TLC data

## Developing Normal-phase Methods — Isocratic

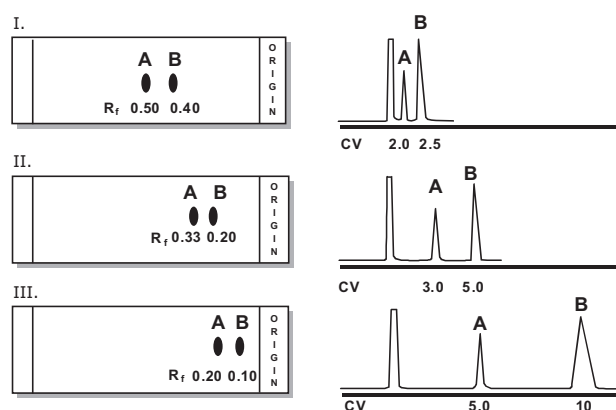
### Stage 1: Converting TLC $R_f$ to CV

Biotage recommends method development using Biotage TLC plates since the silica on the plates matches with cartridge flash silica. Retention on TLC plates is denoted by  $R_f$ . In flash purification, retention is usually measured in CV. Methods developed using TLC are generally transferable to flash chromatography because the relationship between  $R_f$  and CV is reciprocal,  $CV=1/R_f$  (**Figure 8**).



**Figure 8.**  $R_f$  to CV correlation, isocratic elution.

When scouting TLC solvent systems, it is important to realize a low  $R_f$  (0.15–0.35) is preferred because a lower  $R_f$  means a greater CV. Large CVs indicate increased compound-silica contact time, improving the chances of component resolution. Since CV is a measure of compound retention, then  $\Delta CV$  is the measure of compound resolution, (see **Figure 9**). In flash purification,  $\Delta CV$  dictates the sample load range possible for any given cartridge size, (see **Table 3** on page 21). For two adjacent components, a large  $\Delta CV$  is desirable.



**Figure 9.** The  $R_f$ -CV relationship is illustrated in this graphic.

- (I) Although compounds A and B are well resolved on TLC with  $R_f$  of 0.5 and 0.4, respectively, flash purification with the same solvent conditions provides low retention and low resolution ( $\Delta CV = 0.5$ ) for A and B, respectively.
- (II) Lowering the  $R_f$  (A = 0.33, B = 0.20) provides increased retention and resolution ( $\Delta CV = 2$ ).
- (III) Extremely high resolution ( $\Delta CV = 5$ ) is obtained by further reducing the compounds'  $R_f$ .

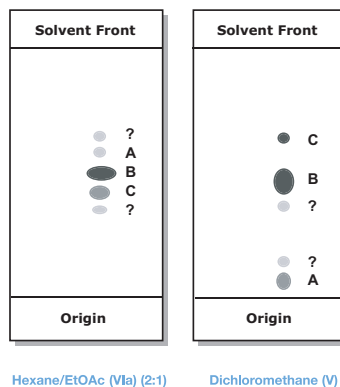
## Stage 2: Determining the best solvent selectivity using TLC

The first step in successful flash purification is maximizing  $\Delta CV$ . Accomplish this by evaluating various solvent mixtures by TLC. Look for a binary mixture that provides the largest  $\Delta CV$  between the compound of interest and all the impurities. All solvents fall into one of the selectivity groups below. Each group has a different impact on a sample component's relative retention to another compound (selectivity). **Table 1** shows the most frequently used flash solvents and their selectivity groups.

Solvent	Selectivity Group
Ether	I
Methanol	II
Ethanol	II
Isopropanol	II
Tetrahydrofuran	III
Dichloromethane	V
Acetone	VIa
Ethyl acetate	VIa
Acetonitrile	VIb
Toluene	VII
Chloroform	VIII
Hexane	—
Heptane	—
Isooctane	—

**Table 1.** Commonly used flash chromatography solvents.

When possible, optimization should include mixtures of hexane with ethyl acetate (VIa), methylene chloride (V), toluene (VII), tetrahydrofuran (III), and ether (I). For more polar compounds and amines, mixtures of methylene chloride (V) with methanol (II) or acetonitrile (VIb) should be evaluated. These solvent combinations provide a broad range of separation selectivity and will help define the correct solvents for purification (**Figure 10**).



**Figure 10.** Impact of solvent selectivity on a chromatographic separation. In hexane/ethyl acetate the compound of interest (**B**) is poorly resolved from its major impurities (**A** and **C**). In dichloromethane, retention of impurities **A** and **C** has been dramatically altered, providing a better purification of **B**.

## Stage 3: Determining the best solvent strength using TLC

When the correct solvents have been determined, the next step is to adjust the solvent composition (solvent strength) so the compound of interest elutes within the  $R_f$  range 0.15–0.35 (6.7–2.8 CV). By adjusting solvent strength to provide elution within this window, the chances for optimal purification are greatly enhanced. As with selectivity, each solvent has its own polarity (**Table 2**). Each solvent mixture or mobile phase then has its own unique solvent strength. Calculation of a solvent mixture's strength is useful for comparison to other solvent mixtures. Solvent mixtures with the same strength but different selectivity can then be evaluated.

Solvent	Solvent Strength
Methanol	0.95
Ethanol	0.88
Isopropanol	0.82
Acetonitrile	0.65
Ethyl acetate	0.58
Tetrahydrofuran	0.57
Acetone	0.56
Dichloromethane	0.42
Chloroform	0.40
Ether	0.38
Toluene	0.29
Hexane	0.01
Heptane	0.01
Isooctane	0.01

**Table 2.** A solvent mixture's strength is calculated using volume proportions and the individual solvent's strength. In this example, diluting a solvent mixture with a less polar solvent (hexane) from 50% to 60% reduces solvent strength, increasing compound retention and resolution ( $\Delta CV$ ). Also, solvent combinations of similar strength but different selectivity can also be compared. Both hexane/ethyl acetate (50:50) and hexane/dichloromethane (30:70) have solvent strength of 0.3, but ethyl acetate and dichloromethane provide different selectivity.

## Flash Chromatography Method Development

### Stage 3: Determining the best solvent strength using TLC (cont'd)

To bring the  $R_f$  of the compound of interest into the optimal range, reduce the amount of polar solvent in the mobile phase. As an example, in **Figure 11**, the results of a solvent selectivity study show a mobile phase of 50% hexane and 50% ethyl acetate (solvent strength = 0.30), providing adequate selectivity for a crude sample (**Figure 11**, top). The  $R_f$  for the compound of interest (B) is 0.4 (2.5 CV) and the  $R_f$  of the impurity (A) is 0.55 (1.8 CV), providing a  $\Delta CV$  of 0.7. With a  $\Delta CV$  this low, only a small sample amount can be flash purified before overload (resolution loss, low purity fractions) occurs. By weakening the solvent strength to 60% hexane and 40% ethyl acetate (solvent strength 0.24) (**Figure 11**, middle); the  $R_f$  of compound B falls to 0.2 (5 CV) and impurity A's  $R_f$  is lowered to 0.3 (3.3 CV) with a resulting  $\Delta CV$  of 1.7, enabling a potential fivefold increase in sample load on a flash cartridge (**Table 3**).

#### Formula:

$$\frac{(\text{Solvent A\%} \times \text{solvent A strength})}{100} + \frac{(\text{Solvent B\%} \times \text{solvent B strength})}{100}$$

#### Examples:

Hexane/ethyl acetate (50:50)

$$\text{Solvent strength} = (0.5 \times 0.01) + (0.5 \times 0.58) = 0.30$$

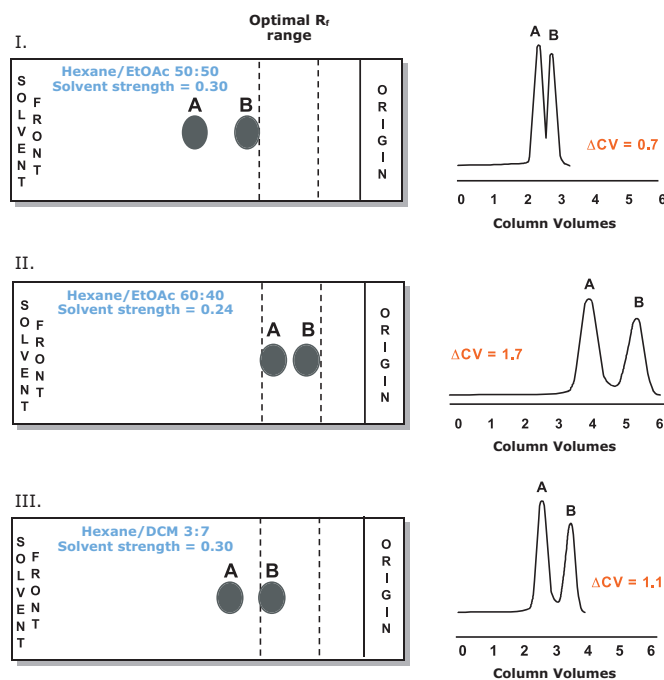
Hexane/ethyl acetate (60:40)

$$\text{Solvent strength} = (0.6 \times 0.01) + (0.4 \times 0.58) = 0.24$$

Hexane/dichloromethane (30:70)

$$\text{Solvent strength} = (0.3 \times 0.01) + (0.7 \times 0.42) = 0.30$$

If you find adequate component retention with a particular solvent mixture, you can prepare other solvent mixtures of similar strength but different selectivity for comparison (**Figure 11**, bottom).



**Figure 11.** Examples of solvent strength on compound retention and resolution.

- (I) TLC shows two sample components resolved with a 50:50 hexane/ethyl acetate solvent system ( $\Delta CV = 0.7$ ). Neither the component of interest (B) nor the impurity (A) has an  $R_f$  value within the optimal 0.15–0.35 range. This leads to poor flash purification.
- (II) After adjusting the solvent to 60% hexane/40% ethyl acetate, the  $R_f$  values for both A and B fall into the optimal zone. Flash chromatography with these conditions shows increased compound retention and greatly improved resolution ( $\Delta CV = 1.7$ ).
- (II) Replacing 50:50 hexane/ethyl acetate with 30:70 hexane/dichloromethane (both 0.30 solvent strength) alters both selectivity and resolution ( $\Delta CV = 1.1$ ).

Once a solvent system has been selected,  $R_f$  values measured, and  $\Delta CV$  values calculated, use **Table 3** to select the correct cartridge for your sample size and  $\Delta CV$ . The data generated from your TLC method development efforts are applicable to any sized Biotage cartridge.

#### Stage 4: Determining the optimum cartridge size and sample load based on TLC data

Having calculated CVs, **Table 3** may be used to help in selecting the most appropriate Biotage cartridge size.

##### Step 1:

Look for the CV that is the closest match to your system

##### Step 2:

Scan down the table based on scale of reaction

##### Step 3:

Select cartridge

	Dimensions (mm)	Column Volume	Flow Rate (mL/min)	Load		
				$\Delta CV = 0.1-1.9$	$\Delta CV = 2.0-3.9$	$\Delta CV = 4.0+$
<b>Biotage ZIP™ 5 g</b>	15.5 x 61	8 mL	5–20	<50 mg	50–250 mg	250–500 mg
<b>Biotage ZIP 10 g</b>	20 x 69	15 mL	10–20	<100 mg	100–500 mg	500–1000 mg
<b>Biotage ZIP 30 g</b>	27 x 116	45 mL	20–40	<300 mg	300–1500 mg	1.5–3 g
<b>Biotage ZIP 45 g</b>	32 x 107	60 mL	30–50	<450 mg	450–2250 mg	2.25–4.5 g
<b>Biotage ZIP 80 g</b>	38 x 130	102 mL	30–50	<800 mg	800–4000 mg	4–8 g
<b>Biotage ZIP 120 g</b>	42 x 176	170 mL	50–75	<1.2 g	1.2–6 g	6–12 g
<b>Biotage SNAP Ultra 10 g</b>	21 x 55	17 mL	10–50	<200 mg	200–1000 mg	1–2 g
<b>Biotage SNAP Ultra 25 g</b>	30 x 72	45 mL	20–100	<500 mg	500–2500 mg	2.5–5 g
<b>Biotage SNAP Ultra 50 g</b>	39 x 81	85 mL	30–150	<1 g	1–5 g	5–10 g
<b>Biotage SNAP Ultra 100 g</b>	39 x 157	164 mL	30–150	<2 g	2–10 g	10–20 g
<b>Biotage SNAP Ultra 340 g</b>	71 x 168	582 mL	65–325	<6.8 g	6.8–34g	34–68 g
<b>Biotage SNAP 10 g</b>	21 x 55	15 mL	10–20	<100 mg	100–500 mg	500–1000 mg
<b>Biotage SNAP 25 g</b>	30 x 72	33 mL	20–40	<250 mg	250–750 mg	750–2500 mg
<b>Biotage SNAP 50 g</b>	39 x 81	66 mL	30–50	<500 mg	500–2500 mg	2.5–5 g
<b>Biotage SNAP 100 g</b>	39 x 157	132 mL	30–50	<1.0 g	1–5 g	5–10 g
<b>Biotage SNAP 340 g</b>	71 x 168	470 mL	65–100	<3.4 g	3.4–17 g	17–34 g
<b>Biotage SNAP 750 g</b>	82 x 291	990 mL	100–300	<7.5 g	7.5–40 g	40–75 g
<b>Biotage SNAP 1500 g</b>	107 x 328	1980 mL	300–500	<15.0 g	15–80 g	80–150 g
<b>Flash 75S</b>	75 x 90	300 mL	100–250	<2.0 g	2–10 g	10–20 g
<b>Flash 75M</b>	75 x 150	500 mL	100–250	<4.0 g	4–20 g	10–40 g
<b>Flash 75L</b>	75 x 300	1000 mL	100–250	<8.0 g	8–40 g	40–80 g
<b>Flash 150M</b>	150 x 300	4.3 L	500–1000	<25.0 g	25–125 g	125–250 g
<b>Flash 150L</b>	150 x 600	8.6 L	500–1000	<50.0 g	50–250 g	250–500 g
<b>Flash 400M</b>	400 x 300	28 L	7000	<200 g	0.2–1.0 kg	1–2 kg
<b>Flash 400L</b>	400 x 600	56 L	7000	<400 g	0.4–2.0 kg	2–4 kg

**Table 3.** Gradient loading table

## Flash Chromatography Method Development

### Normal-phase Gradients

Gradient elution enables chemists to speed purification, improve recovery and yield, and even increase fraction purity. Gradients are slightly different to Isocratic elution modes since the stronger eluting solvent concentration is increased over time relative to the weaker solvent, increasing the solubility of more highly retained compounds. This causes compounds to elute sooner and in tighter bands compared to isocratic systems. As solvent strength increases during the purification, the classic isocratic  $CV=1/R_f$  relationship is not applicable. In a gradient, compounds elute with fewer column volumes than predicted by the isocratic equation. The exact number of elution CV depends on the gradient slope.

Biotage has developed an algorithm to help chemists transfer TLC  $R_f$ s to gradient CVs. This algorithm is incorporated all Biotage Isolera and Isolera LS instrumentation. Using these conditions, a compound with an  $R_f$  of 0.4 elutes near the middle of the purification and is separated from other compounds within the  $R_f$  range of 0.1 to 0.9 (see **Table 3**).

#### TLC-to-gradient

To convert TLC data into a gradient elution method uses the same optimization tools as previously discussed.

After determining the best selectivity and best solvent strength you can use the patented Biotage algorithm to create the best gradient. If you use a Biotage Isolera™ or SP system, the instrument will do this for you.

### Developing Reversed-phase Methods

#### Stage 1: Converting HPLC (high-performance liquid chromatography) retention times and gradient methods to CV

As a technique used for purification of water-soluble compounds, reversed-phase flash purification method development is different to normal-phase. The recommended approach for reversed-phase includes developing and optimizing methods using HPLC and a Biotage KP-C18-HS scaling column (4.6 x 250 mm). The scaling column is packed with the identical C18 phase as the Biotage KP-C18-HS flash cartridges. Begin by creating a gradient on the HPLC from 10–90% acetonitrile (or methanol) in water at 3 mL/min (1 CV/min) with this gradient. A recommended starting point is:

**Segment 1: 10% ACN (or MeOH) for 1 min**

**Segment 2: 10-90% ACN (MeOH) over 10 min**

**Segment 3: Hold 90% ACN for 2 min**

Continue to modify this until the compound of interest is fully separated from its impurities and has a retention time of at least five minutes. On the HPLC, optimal load can be

**First Determine the  $R_f$  values for the compound of interest**

**Second Determine the  $R_f$  value for the compound preceding the compound of interest**

**Third Determine the  $R_f$  value for the compound eluting just after the compound of interest**

Then, convert all  $R_f$  values to CV using the equation  $CV=1/R_f$ . Whichever pair has the lowest  $\Delta CV$  dictates sample load.

The algorithm has three steps for the three gradient segments:

- 1. Use 1/4 of the strong solvent % used with the TLC run, hold for 1 CV**
- 2. Create a linear segment from the step 1 % to 2x the % used with the TLC run over 10 CV**
- 3. Hold this % for 2 CV**

Then, using **Table 3**, locate the  $\Delta CV$  from your calculation above and the cartridge you plan on using and the result will be the recommended sample load.

#### Scale-up

Scaling up flash purification methods is easy and straight forward. Any method developed on a Biotage flash cartridge can be scaled-up to a larger cartridge simply by referring to **Table 4** in the following page.

determined by increasing the sample amount until resolution has been lost. To transfer the HPLC method to flash, convert compound retention time ( $T_r$ ) to column volume using the following equation:

$$\text{Compound CV} = \text{compound } T_r / T_o, \text{ where } T_o = \text{the void time (about 1 min at 3 mL/min)}$$

Use the same formula to convert the gradient program from time to CV:

$$\text{Gradient segment length (time)} / T_o = \text{flash segment length (CV)}$$

By using these formulas and the same solvents, reproducible reversed-phase flash gradients can be developed. In many cases flash methods can be developed from HPLC style methods using scaling columns. Some caution must be noted when transferring methods from commercial HPLC columns (typically 3–5 micron silica) to flash (typically 40–60 micron silica).

## Stage 2: Determining the optimum cartridge size and sample load based on TLC data

Flash scale-up is based on consolidating the solvent's linear velocity and sample load in cartridges of different sizes/volumes. Small scale flash purification is easily scaled with scale-up factors based, which take into account the cartridge differences.

Scaling up a method performed on a Biotage scaling column or flash cartridge is easily accomplished by using **Table 4** below.

Locate the column or cartridge used for method development (Left column) and then locate the scale factor required. Read up that column to determine which cartridge is needed.

### Step 1:

Find your current flash cartridge in the left column

### Step 2:

Read across the row to obtain the scale-up factor for your larger sample

### Step 3:

Read up to find the appropriate Biotage flash cartridge for that scale factor

**Example:** A 1-gram purification on 25 g Biotage SNAP requires scale-up to 30 g, the appropriate scale-up cartridge, according to the table below, is a Biotage SNAP XL (750 g cartridge).

Media Mass	4.6 x 250	5 g	10 g	25 g	30 g	45 g	50 g	80 g	100 g	120 g	200 g	340 g	400 g	750 g	800 g	1.5 kg	2.5 kg	5 kg	20 kg	40 kg	
4.6 x 250	1	2	4	10	12	18	20	32	40	48	80	136	160	300	320	600	1000	2000	8000	16000	
5 g		1	2	5	6	9	10	16	20	24	40	68	80	150	160	300	500	1000	4000	8000	
10 g			1	2.5	3	4.5	5	8	10	12	20	34	40	75	80	150	250	500	2000	4000	
25 g				1	1.2	1.8	2	3	4	5	8	14	16	30	32	60	100	200	800	1600	
30 g					1	1.5	2	3	3.5	4	7	12	14	25	27	50	83	167	667	1333	
45 g						1	1.1	1.8	2.2	3	4.5	7.5	9	17	18	33	56	111	444	888	
50 g							1	1.6	2	2.4	4	6.8	8	15	16	30	50	100	400	800	
80 g								1	1.25	1.5	2.5	4.3	5	9.5	10	19	31	62	250	500	
100 g									1	1.2	2	3.4	4	7.5	8	15	25	50	200	400	
120 g										1	1.7	2.8	3.5	6.5	7	12	21	42	167	333	
200 g											1	1.7	2	3.8	4	7.5	13	25	100	200	
340 g												1	1.2	2.2	2.4	4.4	7.4	15	60	120	
400 g													1	1.9	2	3.8	6.3	13	50	100	
750 g														1	1.1	2	3.5	7	27	54	
800 g															1	1.9	3.1	6.2	25	50	
1.5 kg																1	1.7	3.3	13.5	27	
2.5 kg																	1	2	8	16	
5 kg																		1	4	8	
20 kg																				1	2
40 kg																					1

**Table 4.** Scale-up table for use with flash cartridges

## Ordering Information

### Biotage® SNAP Cartridges

Product	Part Number
<b>Biotage KP-Sil</b>	
10 g, 20/case	FSK0-1107-0010
25 g, 20/case	FSK0-1107-0025
50 g, 20/case	FSK0-1107-0050
100 g, 20/case	FSK0-1107-0100
340 g, 6/case	FSK0-1107-0340
750 g, 2/case	FSK0-1107-0750
1500 g, 2/case	FSK0-1107-1500
<b>Biotage HP-SIL</b>	
10 g, 20/case	FSHP-1207-0010
25 g, 20/case	FSHP-1207-0025
50 g, 20/case	FSHP-1207-0050
100 g, 20/case	FSHP-1207-0100
340 g, 6/case	FSHP-1207-0340
<b>Biotage KP-C18-HS</b>	
12 g, 2/case	FSL0-1118-0012
30 g, 2/case	FSL0-1118-0030
60 g, 2/case	FSL0-1118-0060
120 g, 2/case	FSL0-1118-0120
400 g, 1/case	FSL0-1118-0400
950 g, 1/case	FSL0-1118-0950
1800 g, 1/case	FSL0-1118-1800
<b>Biotage KP-NH</b>	
11 g, 10/case	FSN0-0909-0011
28 g, 10/case	FSN0-0909-0028
55 g, 10/case	FSN0-0909-0055
110 g, 10/case	FSN0-0909-0110
375 g, 1/case	FSN0-0909-0375
900 g, 1/case	FSN0-0909-0900
1800 g, 1/case	FSN0-0909-1800

### Biotage SNAP Ultra Cartridges

Product	Part Number
10 g, 20/case	FSUL-0442-0010
25 g, 20/case	FSUL-0442-0025
50 g, 20/case	FSUL-0442-0050
100 g, 20/case	FSUL-0442-0100
340 g, 6/case	FSUL-0442-0340

### Biotage SNAP XL Cartridges

Product	Part Number
<b>Biotage KP-Sil</b>	
SNAP 750 g, 2/case	FSK0-1107-0750
SNAP 1500 g, 2/case	FSK0-1107-1500
<b>Biotage KP-C18-HS</b>	
SNAP 950 g, 1/case	FSL0-1118-0950
SNAP 1850 g, 1/case	FSL0-1118-1850
<b>Biotage KP-NH</b>	
SNAP 900 g, 1/case	FSN0-0909-0900
SNAP 1800 g, 1/case	FSN0-0909-1800

### Biotage ZIP™ Cartridges

Product	Part Number
5 g, 20/case	440-0500-DZ-20
10 g, 20/case	440-1000-EZ-20
30 g, 20/case	440-3000-FZ-20
45 g, 20/case	440-4500-SZ-20
80 g, 20/case	440-8000-JZ-20
120 g, 20/case	440-120G-UZ-20

### FLASH+® Cartridges

Product	Part Number
<b>Biotage KP-Sil</b>	
FLASH 12+S, 12 x 75 mm, 20/case	FPK0-1107-15026
FLASH 12+M, 12 x 150 mm, 20/case	FPK0-1107-15046
FLASH 25+S, 25 x 75 mm, 20/case	FPK0-1107-16026
FLASH 25+M, 25 x 150 mm, 20/case	FPK0-1107-16046
FLASH 40+S, 40 x 75 mm, 20/case	FPK0-1107-17026
FLASH 40+M, 40 x 150 mm, 20/case	FPK0-1107-17046
<b>Biotage KP-C18-HS</b>	
FLASH 12+S, 12 x 75 mm, 2/case	FPL0-1118-15025
FLASH 12+M, 12 x 150 mm, 2/case	FPL0-1118-15045
FLASH 25+S, 25 x 75 mm, 2/case	FPL0-1118-16025
FLASH 25+M, 25 x 150 mm, 2/case	FPL0-1118-16045
FLASH 40+S, 40 x 75 mm, 2/case	FPL0-1118-15025
FLASH 40+M, 40 x 150 mm, 1/case	FPL0-1118-17020
<b>Biotage KP-NH</b>	
FLASH 12+S, 12 x 75 mm, 20/case	FPNH-12S
FLASH 12+M, 12 x 150 mm, 20/case	FPNH-12M
FLASH 25+S, 25 x 75 mm, 10/case	FPNH-25S
FLASH 25+M, 25 x 150 mm, 10/case	FPNH-25M
FLASH 40+S, 40 x 75 mm, 5/case	FPNH-40S
FLASH 40+M, 40 x 150 mm, 5/case	FPNH-40M



## Biotage SNAP Samplet® Cartridges

Product	Part Number
<b>Biotage KP-Sil</b>	
1 g, 20/case	SAS-1107-0010
3 g, 20/case	SAS-1107-0025
10 g, 20/case	SAS-1107-0100
34 g, 6/case	SAS-1107-0340
<b>Biotage HP-SIL</b>	
1 g, 20/case	SAS-1207-0010
3 g, 20/case	SAS-1207-0025
10 g, 20/case	SAS-1207-0100
34 g, 6/case	SAS-1207-0340
<b>Biotage KP-C18-HS</b>	
1 g, 20/case	SAS-1118-0012
3 g, 20/case	SAS-1118-0030
12 g, 20/case	SAS-1118-0120
40 g, 6/case	SAS-1118-0400
<b>Biotage KP-NH</b>	
1 g, 20/case	SAS-0909-0011
3 g, 20/case	SAS-0909-0028
10 g, 20/case	SAS-0909-0110
37 g, 6/case	SAS-0909-0375

## Biotage SNAP Ultra Samplet Cartridges

Product	Part Number
1 g, 20/case	SAS-0442-0010
3 g, 20/case	SAS-0442-0025
10 g, 20/case	SAS-0442-0100
34 g, 6/case	SAS-0442-0340

## FLASH+ Samplet Cartridges

Product	Part Number
<b>Biotage KP-Sil</b>	
FLASH 12+, 48/case	SAM-1107-1421J
FLASH 25+, 20/case	SAM-1107-16016
FLASH 40+, 20/case	SAM-1107-17016
<b>Biotage KP-C18-HS</b>	
FLASH 12+, 48/case	SAM-1118-1421J
FLASH 25+, 20/case	SAM-1118-16016
FLASH 40+, 20/case	SAM-1118-17016
<b>Biotage KP-NH</b>	
FLASH 12+, 48/case	SAM-NH12
FLASH 25+, 20/case	SAM-NH25
FLASH 40+, 20/case	SAM-NH40

## ISOLUTE® Flash Cartridges

Product	Part Number
<b>ISOLUTE Flash Silica II</b>	
2 g Silica II, 25 mL cartridges, 20/case	440-0200-D
5 g Silica II, 25 mL cartridges, 20/case	440-0500-E
10 g Silica II, 70 mL cartridges, 16/case	440-1000-F
20 g Silica II, 70 mL cartridges, 16/case	440-2000-F
25 g Silica II, 150 mL cartridges, 8/case	440-2500-J
50 g Silica II, 150 mL cartridges, 8/case	440-5000-J
70 g Silica II, 150 mL cartridges, 8/case	440-7000-J
<b>ISOLUTE Flash C18</b>	
2 g C18, 15 mL cartridges, 20/case	451-0200-D
5 g C18, 25 mL cartridges, 20/case	451-0500-E
10 g C18, 70 mL cartridges, 16/case	451-1000-F
20 g C18, 70 mL cartridges, 16/case	451-2000-F
25 g C18, 150 mL cartridges, 8/case	451-2500-J
50 g C18, 150 mL cartridges, 8/case	451-5000-J
70 g C18, 150 mL cartridges, 8/case	451-7000-J
<b>ISOLUTE Flash NH</b>	
2 g NH, 15 mL cartridges, 20/case	454-0200-D
5 g NH, 25 mL cartridges, 20/case	454-0500-E
10 g NH, 70 mL cartridges, 16/case	454-1000-F
20 g NH, 70 mL cartridges, 16/case	454-2000-F
25 g NH, 150 mL cartridges, 8/case	454-2500-J
50 g NH, 150 mL cartridges, 8/case	454-5000-J
70 g NH, 150 mL cartridges, 8/case	454-7000-J
<b>ISOLUTE Flash SCX-2</b>	
2 g SCX-2, 15 mL cartridges, 20/case	456-0200-D
5 g SCX-2, 25 mL cartridges, 20/case	456-0500-E
10 g SCX-2, 70 mL cartridges, 16/case	456-1000-F
20 g SCX-2, 70 mL cartridges, 16/case	456-2000-F
25 g SCX-2, 150 mL cartridges, 8/case	456-2500-J
50 g SCX-2, 150 mL cartridges, 8/case	456-5000-J
70 g SCX-2, 150 mL cartridges, 8/case	456-7000-J

## Dry Loading Vessels (DLV)

Product	Part Number
Dry load vessel kit for external sample loading, 1–30 g capacity	DLV-030
Dry load vessel kit for external sample loading, 10–70 g capacity	DLV-070
Dry load vessel kit with holder, 1 empty cartridge and frit, 50–500 g capacity	DLV-500
Replacement DLVs and frits for DLV-030, 20/case	DLV-035
Replacement DLVs and frits for DLV-070, 20/case	DLV-075
Replacement DLVs and frits for DLV-500, 4/case	DLV-505

## Ordering Information (cont'd)

### Flash 75/Flash 150 Cartridges

Product	Part Number
<b>Biotage® KP-Sil</b>	
Flash 75S, 200 g, 75 x 90 mm, 2/case	FK0-1107-19165
Flash 75M, 400 g, 75 x 150 mm, 2/case	FK0-1107-19045
Flash 75L, 800 g, 75 x 300 mm, 2/case	FK0-1107-19075
Flash 75S (Jumbo), 200 g, 75 x 190 mm, 10/case	FK0-1107-19163
Flash 75M (Jumbo), 400 g, 75 x 150 mm, 10/case	FK0-1107-19043
Flash 75L (Jumbo), 800 g, 75 x 300 mm, 10/case	FK0-1107-19073
Flash 150M, 2500 g, 150 x 300 mm, 2/case	FK0-1107-25075
Flash 150L, 5000 g, 150 x 600 mm, 2/case	FK0-1107-25155
<b>Biotage KP-C18-HS</b>	
Flash 75S, 250 g, 75 x 90 mm, 1/case	FL0-1118-19160
Flash 75M, 480 g, 75 x 150 mm, 1/case	FL0-1118-19040
Flash 75L, 960 g, 75 x 300 mm, 1/case	FL0-1118-19070
Flash 150M, 3000 g, 150 x 300 mm, 1/case	FL0-1118-25070
Flash 150L, 6000 g, 150 x 600 mm, 1/case	FL0-1118-25150
<b>Biotage KP-NH</b>	
Flash 75S, 225 g, 75 x 90 mm, 1/case	FPNH-75S
Flash 75M, 440 g, 75 x 150 mm, 1/case	FPNH-75M
Flash 75L, 880 g, 75 x 300 mm, 1/case	FPNH-75L
Flash 150M, 2750 g, 150 x 300 mm, 1/case	FPNH-150M
Flash 150L, 5500 g, 150 x 600 mm, 1/case	FPNH-150L
<b>Diaion® HP20</b>	
Flash 75S, 75 x 90 mm, 2/case	FT6-2030-19165
Flash 75M, 75 x 150 mm, 2/case	FT6-2030-19045
Flash 75L, 75 x 300 mm, 2/case	FT6-2030-19075
Flash 75S (Jumbo), 75 x 190 mm, 10/case	FT6-2030-19163
Flash 75M (Jumbo), 75 x 150 mm, 10/case	FT6-2030-19043
Flash 75L (Jumbo), 75 x 300 mm, 10/case	FT6-2030-19073
Flash 150M, 150 x 300 mm, 2/case	FT6-2030-25075
Flash 150L, 150 x 600 mm, 2/case	FT6-2030-25155
<b>Diaion HP20SS</b>	
Flash 75S, 270 g, 75 x 90 mm, 1/case	FT6-2530-19160
Flash 75M, 450 g, 75 x 150 mm, 1/case	FT6-2530-19040
Flash 75L, 900 g, 75 x 300 mm, 1/case	FT6-2530-19070
Flash 150M, 3600 g, 150 x 300 mm, 1/case	FT6-2530-25070
Flash 150L, 7200 g, 150 x 600 mm, 1/case	FT6-2530-25150

### Flash 400 Cartridges

Product	Part Number
<b>Biotage KP-Sil</b>	
Flash 400M, 20 kg, 2/case	FK0-1107-50075
Flash 400L, 40 kg, 2/case	FK0-1107-50155
<b>Biotage KP-C18-HS</b>	
Flash 400M, 24 kg, 1/case	FL0-1118-50070
Flash 400L, 48 kg, 1/case	FL0-1118-50150
<b>Flash-WAC</b>	
Flash 400M, 13 kg, 2/case	C1YR-4021-50075
Flash 400L, 26 kg, 2/case	C1YR-4021-50155

## Accessories

Product	Part Number	Product	Part Number
<b>Thin-Layer Chromatography (TLC)</b>		<b>ISOLUTE® FlashPack Plunger Assemblies</b>	
Biotage KP-NH TLC plate, 5 x 10 cm, 50/box	TLC-KPNH-0510-FI	Plunger for 6 mL cartridges	00-012-000
Biotage KP-Sil TLC plate, 5 x 10 cm, glass, 50/box	TLC-0510-FI	Plunger for 15 mL cartridges	00-016-000
Biotage KP-Sil TLC plate, 10 x 10 cm, glass, 25/box	TLC-1010-FI	Plunger for 25 mL cartridges	00-020-000
Biotage KP-Sil TLC plate, 2.5 x 7.5 cm, glass, 100/box	TLC-2575-FI	Plunger for 70 mL cartridges	00-027-000
		Plunger for 150 mL cartridges	00-037-000
<b>Liquid Sample Injection Valve and Adapters</b>		<b>Cartridge Holders</b>	
Adapter to attach a Biotage 3-way injection valve to a Biotage SNAP cartridge	411081	Cartridge holder for Biotage Isolera systems	
		Biotage SNAP 10 g cartridge	411922
<b>Cartridge Adapters</b>		Biotage SNAP 25 g cartridge	411776
Biotage SNAP to Biotage FlashMaster II and PersonalPlus adapter	411069	Biotage SNAP 50 g and 100 g cartridge	411923
Biotage SNAP 10 g cartridge adapter ring and connecting tubing for SP systems	410792	Biotage SNAP 340 g cartridge	411924
Biotage SNAP 25 g cartridge adapter ring and connecting tubing for SP systems	411824	Biotage ZIP 5 g and 10 g cartridge	413902
Biotage SNAP 50 g/100 g cartridge adapter ring and connecting tubing for SP systems	410797	Biotage ZIP 30 g cartridge	413302
Biotage SNAP 340g cartridge adapter kit for non-Biotage SP systems	410805	Biotage ZIP 45 g cartridge	413303
Female Luer inlet fitting for SNAP 750 g and 1500 g cartridges, 1/pk	412358	Biotage ZP 80 g cartridge	413304
Male Luer outlet fitting for SNAP 750 g and 1500 g cartridges, 1/pk	412537	Biotage ZP 120 g cartridge	413305
Luer lock to Luer tip adapter	120-1110	DLV-030 and -070 holder for use with Biotage SNAP 10-100 g cartridges, 1/pk	413127
		DLV-030 and -070 holder for use with Biotage SNAP 340 g cartridges, 1/pk	413128
<b>Samplet® Accessories</b>		Biotage SNAP 340g cartridge holder for SP systems, 1/pk	410800
Empty Samplet kit for Biotage SNAP		Biotage SNAP 750 g and 1500 g cartridge holder for Biotage Isolera systems, 1/pk	412422
1 g Samplet, 20/case	SES-0010		
3 g Samplet, 20/case	SES-0025	<b>Scaling Column</b>	
10 g Samplet, 20/case	SES-0100	Biotage KP-C18-HS 4.6 x 250 mm HPLC column	S1L0-1118-93050
34 g Samplet, 6/case	SES-0340		
Dry load frits for		<b>Biotage SIMs (Sample Injection Module)</b>	
10 g Biotage SNAP cartridges, 20/case	SLF-0010	Stainless steel Biotage SIM for Flash 75, 100 mL	SIM-0102
25 g Biotage SNAP cartridges, 20/case	SLF-0025	Stainless steel Biotage SIM for Flash 75, 500 mL	SIM-0502
100 g Biotage SNAP cartridges, 20/case	SLF-0100	Stainless steel Biotage SIM for Flash 150, 1000 mL	SIM-1002
340 g Biotage SNAP cartridges, 6/case	SLF-0340	Stainless steel Biotage SIM for Flash 150, 2000 mL	SIM-2002
Frit insertion tool for			
1 g empty Biotage SNAP Samplet cartridges	SFS-0010		
3 g empty Biotage SNAP Samplet cartridges	SFS-0025		
10 g empty Biotage SNAP Samplet cartridges	SFS-0100		
34 g empty Biotage SNAP Samplet cartridges	SFS-0340		

## Tools for Discovery and Development Chemistry

### Discovery Chemistry

- Flash purification
- Microwave synthesis
- Work-up and sample preparation
- Evaporation
- Polymer supported reagents

### Process Chemistry

- Silica and polymer metal scavengers
- Genotoxin removal
- Catalyst screening
- Purification scale-up

### Peptide Synthesis and Purification

- Automated, semi-automated and manual synthesizers
  - Microwave peptide synthesis
  - Room temperature peptide synthesis
  - Solution phase peptide synthesis
- Resins for solid phase peptide synthesis
- HPLC columns

### Analytical Chemistry / Sample Preparation

- Supported liquid extraction columns and plates
- Automated SPE systems
- Evaporation instrumentation
- Molecularly imprinted polymers
- Silica and resin based SPE columns and plates
- Processing tools for SPE columns and plates

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