SOLUTIONS FOR SCAVENGING OF METAL AND ORGANIC IMPURITIES









Founded in 1995, SiliCycle is specialized in the development, manufacturing and commercialization of high value silica gels and specialty products for chromatography, purification and synthesis.

Scavenging Solutions

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SiliCycle Scavenging Solutions

- Broadest portfolio of scavengers with associated applications
- Great variety of formats for all purification scales: from laboratory to plant scale
- Excellent scavenging efficiency for a wide range of metal and organic impurities
- Green and environmentally friendly technology
- Compatible with various experimental conditions, solvents, pH, and temperatures

SiliaMetS and SiliaBond Silica-Based Scavengers

Chemists have been searching for techniques and tools to separate, isolate and purify chemical substances from one another to improve the quality of the synthesized molecules. SiliCycle scavenging technologies enable powerful purification processes to help reach new purity standards.

Silica-based scavengers have been proven to be the most effective method for removing metal or organic impurities without contaminating drug candidates. A silica matrix offers many advantages over polymers:

- Reduced purification steps
- No swelling
- More general solvent compatibility

- Higher mechanical and thermal stabilities
- Easily scalable applications
- Availability in different product formats, such as bulk, E-PAK flow cartridges, SPE or flash cartridges, etc.



Typical structure of our functionalized silicas with various organic groups

What makes Silia*MetS* metal scavengers and Silia*Bond* organic scavengers so easy to use is their heterogeneous nature. When used in bulk in a contaminated solution, all you need to do is filter off the scavenger with the bound impurity and dispose of it.

When used in SPE or flash cartridges for batch reactions, the filtration process – with adapted experimental conditions – will leave the contaminant in the cartridge, rendering the solution free of impurities.





Different Formats for Different Applications

Please refer to our Ordering Information section to learn more about all formats available and the corresponding part numbers.

Scavengers as Bulk Silica

All our scavengers can be used in bulk directly in your reaction flask or reactor and are available from 5 g to 25 kg formats, up to multi-ton scale.

- All our scavengers have, by default, the same silica backbone: our SiliaFlash R10030B.
 - Particle Size: 40 63 μm
 - Pore Size: 60 Å
- All our Silia*Flash* silica gels of various particle sizes and pore sizes are available as silica backbones upon request.

Scavengers in SiliaPrep SPE and SiliaSep OT Cartridges

All our scavengers are available in pre-packed cartridges.

- SiliaPrep SPE cartridges From 3 mL / 200 mg to 12 mL / 2 g
- SiliaSep OT (Open Top) flash cartridges From 25 mL / 5 g to 150 mL / 70 g

Scavengers in SiliaSep Flash Cartridges

All our scavengers are available pre-packed in flash cartridges.

• SiliaSep flash cartridges From 4 g to 1.6 kg bed weight

Packings can also be tailored to your available equipment and scales.

Scavengers in SiliaChrom Guard Cartridges

Silia*Chrom* HPLC guard cartridges are designed to effectively protect both analytical and preparative HPLC columns. The usage of this shorter column is highly recommended to prolong column lifetime and does not alter chromatography. All metals can be prejudicial and very damaging to your column and detector, complicating purification steps, often making them longer, more laborious and less effective.

Crude reaction mixtures can now directly be injected without further metal removal, which will save precious time for the chemist. Another great benefit is that there is much less risk of corroding the equipment by injecting dirty samples.

Silia*Chrom* guard cartridges are available in lengths of 10, 20, and 30 mm and four internal diameters (*ID: 4.0, 10, 21.2, and 30 mm*).









SILICYCLE

Scavengers in E-PAK Flow Cartridges

E-PAK is a family of radial flow adsorption cartridges developed specifically for pharmaceutical processings.

Created with proprietary technology, E-PAK cartridges provide rapid adsorption kinetics at flow rates and processing capacities suitable for laboratory, pilot and commercial operations. They are designed for use with both organic and aqueous solvents, and incorporate design features useful for the production of active pharmaceutical ingredients (*API*).

Features and Benefits

- Proven cartridge design ensures rapid, simple and reliable technology
- High adsorption capacity and flow rate
- Fixed-bed design ensuring safer handling, clean-up and disposal
- · Large adsorbent capacity in small area footprint increases product recovery and reduces solvent requirements

Sorbents for E-PAK Cartridges

E-PAK cartridges are available in a range of sorbents to accommodate the broad range of processing requirements. Other adsorbents are available under request.

	Sorbents for E-PAK	
Active Adsorbents (Typical Loading)	For Removal and / or Recovery of:	pH Operation
SiliaMetS Thiol (1.2 mmol/g)	Pd, Ag, Au, Hg, Os, Ru, Cu, Ir, Pb, Rh, Se, Sn & U	
SiliaMetS DMT (0.5 mmol/g)	Pd, Ag, As, Au, Bi, Ir, Ni, Os, Pt, Rh, Ru, Se, U, Cd, Co, Cu, Fe, Sc & Zn	
SiliaMetS Imidazole (0.96 mmol/g)	Cd, Co, Cu, Fe, Ir, Li, Mg, Ni, Os, U, W, Zn, Cr, Pd & Rh	
SiliaMetS AMPA (0.8 mmol/g)	Al, Ce, Dy, Er, Eu, Gd, Ho, La, Lu, Mg, Mn, Nd, Ni, Pm, Pr, Sb, Sm, Tb, Tm, V, Yb, Zr, Co, Cu, Fe & Zn	
SiliaBond Amine (1.2 mmol/g)		
SiliaMetS Diamine (1.28 mmol/g)	Pd, Cr, Pt, Rh, W, Zn, Cd, Co, Cu, Fe, Hg, Ni, Pb, Ru, Sc, Se & U	2 to 9
SiliaMetS Triamine (1.11 mmol/g)		
SiliaMetS TAAcONa (0.41 mmol/g)	Bi, Ca, Cd, Cs, Cu, Fe, Ir, La, Li, Mg, Ni, Os, Rh, Ru, Sc, U Cr, Pd, Se & Sn	
SiliaBond Cyano (1.38 mmol/g) and Florisil®	Various organic molecucles	
SiliaBond Propylsulfonic Acid SCX-2 (0.63 meq/g)	Amines & anilines, ion exchange	
Silia <i>Flash</i> Bare Silica	Very vast range of organic impurities, metals, pigments, etc.	
Activated Carbon SiliaCarb C-CA		
Activated Carbon SiliaCarb C-HA		1 to 12
Activated Carbon SiliaCarb C-VA	- Precious metal catalysis & colors	1 10 13
Activated Carbon SiliaCarb C-VW		

Best scavenger for the removal of a particular metal is indicated in Navy Blue while good scavenger is indicated in Pale Blue.

Solvent Compatibility

E-PAK cartridges are manufactured using a proprietary technology and chemically stable materials in most common organic solvents. They have been tested and found satisfactory for use with the following commonly used solvents:

- 2-Butanone
- Heptane
- Tetrahydrofuran (at room temperature)

- DichloromethaneEthanol
- MethanolMTBE
- Toluene (at room temperature)

- Ethyl acetate
- N-ethyl-2-pyrrolidone

Solutions for Scavenging of Metal and Organic Impurities



Scavenging

Analysis



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Various Ways of Using E-PAK

Depending on the application and how you prefer to work, E-PAK cartridges can be used in different ways as shown below on all housing scales (*lab, pilot and commercial scale*). Click on the picture to access the video on E-PAK methods.

For a single pass usage, we suggest to run at very low flow rate compared to recirculation process, which can be run at higher flow rate.



E-PAK Portfolio

Lab Scale

Lab scale cartridges are designed to facilitate small samples evaluation. Testing with loose media can be done with samples as small as a few milliliters and is normally done before cartridge testing to identify the formula with the highest capacity to remove contaminants with the highest recovery.

	Lab Scale Cartridges						
Cartridge Size Diameter × Height (cm)	Typical Flow Rate Range (mL/min)	Pressure Drop with w/1 cP Fluid (psig)	Media We Silia <i>Flash</i> , Silia <i>MetS</i> and Silia <mark>Bond</mark>	ight (g) SiliaCarb			
5×1	1 - 20	≤ 5	8	5			
5 × 10	10 - 200	≤ 5	75	50			
5 × 25	25 - 500	≤ 5	200	125			



Note: Faster flow rates can be used for the lab scale cartridges depending on the application or the scavenging difficulty (1 cm up to 100 mL/min, 10 cm up to 500 mL/min and 25 cm up to 1 L/min).

Pilot and Commercial Scale

E-PAK pilot scale cartridges provide rapid processing for volumes from 10 to hundreds of liters, and can establish the parameters upon moving to larger scales, since E-PAK achieve true linear scalability.

E-PAK commercial scale cartridges provide rapid processing for manufacturing operations needing to process batch sizes of > 10,000 L or can be adapted for continuous operation using a duplex design.

Pilot and Commercial Scale Cartridges							
Cartridge Size Diameter × Height (cm)	Typical Flow Rate Range (L/min)	Pressure Drop with w/1 cps Fluid (psig)	Media Wei Silia <i>Flash</i> , Silia <i>MetS</i> and Silia <mark>Bond</mark>	ght (kg) SiliaCarb			
Pilot Scale							
16.5 × 12.5	0.10 - 2.5	≤ 10	0.87	0.55			
16.5 × 25	0.25 - 5	≤ 10	1.75	1.10			
Commercial Scale							
16.5 × 50	0.50 - 10	≤ 10	3.50	2.10			
16.5 × 100	1 - 20	≤ 10	7.00	4.10			



Both pilot and commercial cartridges are provided with a Code 8 (*closed top and open bottom end caps-bottom with double 2-222 Teflon® encapsulated Viton® o-ring*) cartridge sealing configuration. To meet commercial processing requirements, E-PAK cartridges can be operated in parallel for increased capacity.



Housings

Lab Scale

- Various housing lengths available (for 1 cm, 10 cm, and 25 cm cartridges)
- Made in stainless steel 316L or Hastelloy C276
- Pressure rating for housing 150 psi (10 bar)
- · Easy housing conversion for all lengths by changing the bowl
- · Operated with standard pump, low pressure and peristatic

Pilot and Commercial Scale

- Various housings available for simultaneous operation of 1 to 12 cartridges (for 12.5 cm, 25 cm, 50 cm and 100 cm cartridges)
- Made in stainless steel 316L or Hastelloy C276
- Pressure rating for housing 150 psi (10 bar)
- Can be operated in parallel to process batch sizes of \geq 1,000 L
- · ASME and PED 2014/68/EU, CE compliant

Scale-Up Calculation

Although there are always exceptions, scale-up projections based on a linear extrapolation of adsorbent mass have proven to be quite accurate when test conditions including contact time, temperature, solvent type, and contaminant and compound levels are held constant. The following table shows the scale-up / relative change in mass between lab, pilot and commercial size E-PAK cartridges available with activated carbons and scavengers.

\mathcal{Q}	Scale-Up Calculation for Various Adsorbents						
Cartridge Sizes (cm)	5×1	5 × 10	5 × 25	16.5 × 12.5	16.5 × 25	16.5 × 50	16.5 × 100
Typical Scale-up Factor	-	10	25	110	220	440	875
Mass of Activated Carbon (g)	5	50	125	550	1,100	2,100	4,100
Mass of Silica (g)	8	75	200	875	1,750	3,500	7,000
# mmol SiliaMetS AMPA (0.8 mmol/g)	6.4	60	160	700	1,400	2,800	3,600
# mmol SiliaMetS Diamine (1.28 mmol/g)	10.2	96	256	1,120	2,240	4,480	8,960
# mmol SiliaMetS DMT (0.50 mmol/g)	4.0	38	100	438	875	1,750	3,500
<pre># mmol SiliaMetS Imidazole (0.96 mmol/g)</pre>	7.7	72	192	840	1,680	3,360	6,720
# mmol SiliaMetS TAAcONa (0.41 mmol/g)	3.3	31	82	359	717	1,435	2,870
# mmol SiliaMetS Triamine (1.11 mmol/g)	8.9	83	222	971	1,942	3,885	7,770
# mmol SiliaMetS Thiol (1.20 mmol/g)	9.6	90	240	1,050	2,100	4,200	8,400
# mmol SiliaBond Amine (1.20 mmol/g)	9.6	90	240	1,050	2,100	4,200	8,400
# mmol SiliaBond Cyano (1.38 mmol/g)	11.0	103	276	1,207	2,415	4,830	9,660
# mmol SiliaBond SCX-2 (0.63 meq/g)	5.0	47	126	551	1,102	2,205	4,410
Bed Volume (<i>cm</i> ³)	18.8	188	470	2,375	4,750	9,500	19,000
Recommended Flow Rate (mL/min)	7.5	75	190	950	1,900	3,800	7,600
Typical Flow Rate Range (<i>mL/min</i>)	1 - 20	10 - 200	25 - 500	100 - 2,500	250 - 5,000	500 - 10,000	1,000 - 20,000

Measurement Methodology:

Approximated Tank Volume (mL)

Minimum System Flushing (mL)

- Scale-up Factor: mass of silica / 8 g (smallest size)
- Mass of Activated Carbon (g) / Mass of Silica (g): amount of SiliaCarb / SiliaMetS / SiliaBond in the cartridge

50

150

- # mmol SiliaMetS / SiliaBond XXX (X.X mmol/g): silica mass x typical loading of SiliaMetS / SiliaBond
- Bed Volume (*cm*³): total volume of the cartridge without the volume of the hole
- Recommended Flow Rate (mL/min): for residence times of 2.5 minutes
- Flow Rate Range (mL/min): for residence times from 1 to 20 minutes. Faster flow rate can be used for some applications.
- Minimum System Flushing (*mL*): corresponds to 3 tank volumes measured experimentally with the cartridges inside the housing. The volume of solvents needed for conditioning can vary depending on experimental conditions. It is recommended to do a minimum of 3 system flushing before use.

200

600

Click to watch our video: from lab to

commercial scale E-PAK housings





450

1,350

2,500

7,500

5,000

15,000

11.600

35,000



23,300

70,000

Quality and Regulatory Documentation

The SiliCycle scavengers have been used for decades in pilot plants and production units under GMP conditions by the pharmaceutical industry as well as CMOs and CROs. They have ran their own analysis proving that silica-based scavengers can safely be used, both in reactors and cartridges to purify their compounds.

SiliCycle is committed to high quality standards and all products are manufactured in an ISO 9001:2015 compliant facility and subjected to stringent quality control.

For any inquiries, please contact: scavenger@silicycle.com

All products are shipped with the following information:

- Certificate of Analysis (CoA)
- Safety Data Sheet (SDS)
- Technical Information

BSE / TSE Declaration (non animal-derived)
GMO-Free Certificate

Other statements available upon request:

Melamine-Free Certificate, etc.

Our Regulatory Support Files (*RSF*) are documents that include both proprietary and non-proprietary information on performance, chemical / thermal / mechanical stability, extractable and leachable compounds, SOPs, scale-up procedures, batch history, analytical methods and more. RSF documentation can be obtained through a Non-Disclosure Agreement (*NDA*).

SiliCycle can also work with you to provide customized regulatory documents, including specific analytical tests in line with your needs.

Stability and Leaching Studies

Since our metal and organic scavengers are being used by many pharmaceutical companies, each Silia*MetS* and Silia*Bond* manufactured by SiliCycle is submitted to an extensive washing procedure to ensure the product exhibits extremely low levels of extractables and leachables.

Manufacturing Capability

Silia*MetS* and Silia*Bond* are manufactured at our headquarters in Quebec City, where we can meet all customers production needs.

Our state of the art facilities include (but are not limited to):

- 1,000 L to 10,000 L reactors (total capacity of 38,000 L)
- Stainless steel and hastelloy nutsche filters (3 m²)
- Bulk solvent tank farm (60,000 L capacity)



Enjoy a virtual tour of SiliCycle's facility









Metal and Organic Scavenging Screening Services

Scavenging screening services provide solutions to quickly develop the most efficient metal scavenging process providing both time and cost savings. Confidentiality is assured and we will be working with you, in function of your needs.



Screening and Optimization Services

Batch: Lab and Process Development

Step 1: Screening

- Different Silia*Carb* activated carbons and Silia*MetS* metal scavengers will be tested on the crude product at two different temperatures.
- Influence of contact time will also be evaluated.

Step 2: Optimization

- After this initial screening, we will select the most efficient carbon and scavenger to optimize the ratio vs the API.
- Pairwise combinations of metal scavengers could also be tested, if satisfactory results are not obtained. Scavenging yields as well as recoveries will be evaluated.

Step 3: Scale-up Confirmation and Optimization

- A new amount of crude product will then be treated with Silia*Carb* (*if relevant*), using the best obtained conditions to get enough product to work on the next step.
- Finally, we will test the best conditions on the API solubilized in two other solvent systems (*depending on API solubility and customer's requirement*). Scavenging yields as well as recoveries will be evaluated.
- Best conditions will be transposed on larger scale (to be determined based on the remaining crude product).

Batch to E-PAK: Including Process Development

Includes all points performed with "batch" service plus the following:

- Silia*Carb* and Silia*MetS* best conditions will be then transfered to E-PAK radial flow cartridges.
- Contact time's effect can be evaluated as well as parameters optimization on E-PAK for both Silia*Carb* and Silia*MetS* reaction.
- Scalability study based on customer final equipment can be performed if enough crude is available.

→ Main objective is to accelerate research, optimize production costs, while remaining fully in compliance with new regulations and environmental challenges

→ Scavenging strategies imply lower volumes of solvent, less person-days and higher yields by minimizing loss of product via classical purification processes

Two Plans for R&D Services		
Includes	BATCH Scavenger Screning	BATCH TO E-PAK Scavenger Screening
SiliaCarb and SiliaMetS screening in bulk		
Optimization in different solvents		 Image: A start of the start of
Metal concentration tracked by ICP-MS or ICP-OES		
Scale-up confirmation and optimization (will need to be confirmed by our researchers)		
Transfer of the optimized conditions to E-PAK radial flow cartridge		>
Evaluation of contact time in E-PAK radial flow cartridge		\checkmark
Scalability based on customer final equipment can be performed if enough product is available		
Complete report provided		 Image: A start of the start of
Expected screening service delay after reception of customer's sample	3 - 4 weeks	4 - 6 weeks
Typical crude quantity needed (will need to be confirmed by our researchers)	50 to 200 g of crude	100 to 200 g of crude

Analysis



SiliaMetS Metal Scavengers Portfolio

	SiliaMetS Metal Scavengers Technical Information						
	Scavengers	Structure	Brief Description	Metals Removed ¹	Typical Characteristics ^{2, 3}		
PGI-S	SiliaMetS Thiol PN: R51030B Loading: ≥ 1.20 mmol/g Endcapping: Yes	Зульн	Silia <i>MetS</i> Thiol is our most versatile and robust metal scavenger for a variety of metals under a wide range of conditions.	Ag, Au, Hg, Os, Pd & Ru Cu, Ir, Pb, Rh, Sn & U	Color: White Density: 0.682 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 4 Years		
PGI-S	SiliaMetS DMT PN: R79030B Loading: ≥ 0.50 mmol/g Endcapping: Yes	SH N N N N N N SH	Silia <i>MetS</i> DMT is the silica-bound equivalent of 2,4,6-trimercaptotriazine (<i>trithiocyanuric acid</i> , <i>TMT</i>). It is a versatile metal scavenger for a variety of metals and the preferred metal scavenger for ruthenium catalysts and hindered Pd complexes (<i>i.e.</i> $Pd(dppf)Cl_2$).	Ag, As, Au, Bi, Ir, Ni, Os, Pd, Pt, Rh, R, Se & U Cd, Co, Cu, Fe, Sc & Zn	Color: Light brown Density: 0.732 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years		
	SiliaBond Amine PN: R52030B Loading: ≥ 1.20 mmol/g Endcapping: Yes	S NH2		Cd, Cr, Pd, Pt, Rh, Re & Ru Co, Cu, Fe, Hg, Pb, U, W & Zn	Color: Off-white Density: 0.700 g/mL Solvent Compatibility: 2 Prolonged Storage: 2 Shelf Life: 2 Years		
-	SiliaMetS Diamine PN: R49030B Loading: ≥ 1.28 mmol/g Endcapping: Yes		Also known for their electrophile scavenging efficiencies and their base reagent qualities, Silia <i>MetS</i> Amine, Diamine and Triamine have also proven to be very useful for the scavenging of the following metals: Pd, Pt, Cr, W and Zn.	Cr, Pd, Pt, W & Zn Cd, Co, Cu, Fe, Hg, Ni, Pb, Ru, V & Sc	Color: Off-white Density: 0.728 g/mL Solvent Compatibility: 2 Prolonged Storage: 2 Shelf Life: 2 Years		
PGI-S	SiliaMetS Triamine PN: R48030B Loading: ≥ 1.11 mmol/g Endcapping: Yes	Correction of the second secon		Cr, Pd, Pt, W & Zn Ag, Cd, Co, Cu, Fe, Hg, Ni, Os, Pb, Rh, Ru & Sc	Color: Off-white Density: 0.736 g/mL Solvent Compatibility: 2 Prolonged Storage: 2 Shelf Life: 2 Years		
	SiliaMetS AMPA PN: R85130B Loading: ≥ 0.80 mmol/g Endcapping: Yes	CH N POH CH N OF CH CH PO	Silia <i>MetS</i> AMPA is an aminomethyl-alkylphosphonic acid ligand known for its excellent metal-bonding properties. It is particularly efficient to remove Al, Sb, Ni, La, and also very effective for Co, Cu, Fe, Mg and Zn scavenging from reaction intermediates or final APIs.	Al, Ce, Dy, Er, Eu, Gd, Ho, La, Lu, Mg, Mn, Nd, Ni, Pm, Pr, Sb, Sm, Tb, Tm, V, Yb & Zr Co, Cu, Fe, Mg & Zn	Color: Yellow Density: 0.707 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 1 Year		
	SiliaMetS Cysteine PN: R80530B Loading: ≥ 0.30 mmol/g Endcapping: Yes	SH CNa	Silia <i>MetS</i> Cysteine is the silica-bound equivalent of the amino acid cysteine. It is a versatile scavenger for a variety of metals and the preferred metal scavenger for tin residues. By attaching the molecule to the backbone via the amino group, the thiol group remains free and accessible for higher metal scavenging efficiency.	Au, Cd, Fe, Ir, Os, Ru, Sc, Sn & U Ca, Cr, Cs, Cu, La, Mg, Pd, Pt, Rh & Zn	Color: Orange Density: 0.665 g/mL Solvent Compatibility: 2 Prolonged Storage: 3 Shelf Life: 1 Year		
pgl-s	SiliaMetS DEAM PN: R54430B Loading: ≥ 0.85 mmol/g Endcapping: Yes	Сон сон	Silia <i>MetS</i> DEAM is a versatile scavenger designed to remove trace metal of Ti, Zn, Fe and Ag as well as boronic acids from reaction intermediates or final APIs.	Ag, Au, Fe, Sn, Ti, Zn & Zr	Color: Off-white Density: 0.691 g/mL Solvent Compatibility: 1 Prolonged Storage: 2 Shelf Life: 2 Years		



SiliaMetS Metal Scavengers Technical Information							
Scavengers	Structure	Brief Description	Metals Removed ¹	Typical Characteristics ^{2, 3}			
SiliaMetS DOTA PN: R91030B Loading: ≥ 0.38 mmol/g Endcapping: Yes	Contraction of the second seco	Silia <i>MetS</i> DOTA is a silica-supported tetracarboxylic acid and its various conjugate bases. DOTA molecule is a well-adopted complexing agent. Linked to various metals, so formed-complexes are used as contrast agents in cancer treatments or other medical applications.	Ca, Cu, Gd, La, Ni & Zn Co, Fe, Mg, Pd, Pt & Rh	Color: Light yellow Density: 0.681 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 1 Year			
SiliaMetS Imidazole PN: R79230B Loading: ≥ 0.96 mmol/g Endcapping: Yes		Silia <i>MetS</i> Imidazole is a versatile metal scavenger for a variety of metals including Cd, Co, Cu, Fe, Ni, Os, Pd and Rh.	Cd, Co, Cu, Fe, Ir, Li, Mg, Ni, Os, U, W & Zn Cr, Pd & Rh	Color: Off-white Density: 0.681 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years			
SiliaMetS TAAcOH PN: R69030B Loading: ≥ 0.41 mmol/g Endcapping: No		Silia <i>MetS</i> TAAcOH & TAAcONa are supported versions of EDTA in their acid and sodium salt forms. These two products are effective metal scavengers for Ca, Mg, Li, Ir, Cs, Os, Sn, Pd, Ni	Au, Ca, Co, Ir, Li, Mg, Ni, Os, Ru, Sc & U Cr, Cs, Fe, Pd, Rh & Sn	Color: Off-white Density: 0.635 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years			
SiliaMetS TAAcONa PN: R69230B Loading: ≥ 0.41 mmol/g Endcapping: No	Contouri Contouri	and Cu. Silia <i>MetS</i> TAAcOH is effective for metals in low or zero oxidation states, compared to Silia <i>MetS</i> TAAcONa which is useful for metals in higher oxidation states (≥ 2).	Bi, Ca, Cd, Cs, Cu, Fe, Ir, La, Li, Mg, Ni, Os, Rh, Sc, Sn & U Cr, Pd, Ru & Zn	Color: Off-white Density: 0.712 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years			
SiliaMetS Thiourea PN: R69530B Loading: ≥ 1.07 mmol/g Endcapping: Yes	a م الب الب	Silia <i>MetS</i> Thiourea is a versatile metal scavenger for all forms of palladium and is widely used in the pharmaceutical industry. Once complexed with a transition metal, it has been reported to be an effective catalyst.	Au, Pd & Ru Ag, Cu, Fe, Os, Rh, Sc & Sn	Color: Off-white Density: 0.767 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years			
SiliaBond Tosic Acid PN: R60530B Capacity: ≥ 0.54 meq/g Endcapping: Yes		Silia <i>Bond</i> Tosic Acid is in a class of strong acids used in different fields of synthetic organic chemistry. The aromatic ring makes it slightly more acidic than other supported sulfonic acids.	Fe, Rh & Sn Ag, Cu, Ni, Pd, Pt, Ru & Zn	Color: Off-white Density: 0.698 g/mL Solvent Compatibility: 2 Prolonged Storage: 1 Shelf Life: 2 Years			

¹ Scavenging Efficiency:

- Best scavenger for the removal of a particular metal is indicated in Navy Blue
- Good scavenger indicated in Pale Blue

² Solvent Compatibility:

- 1- All solvents, aqueous and organic
- 2- All organic solvents

³ Prolonged Storage:

- 1- Keep dry
- 2- Keep cool (< $8^{\circ}C$) and dry
- 3- Keep cool (< $8^{\circ}C$), dry, and under inert atmosphere



Potentially Genotoxic Impurities (*PGI*) Scavenger See pages 16 - 17 or contact us for more information



Synthesis

Chromatography

Sample Preparation

Analysis

R&D Services

SiliaMetS Metal Scavengers Selection Table

This table will help in selecting the most efficient scavenger for a specific metal and application. Since some parameters may affect the efficiency of the scavenging, we recommend performing a preliminary screening experiment using the Silia*MetS* Metal Scavenger Kit.

Best scavenger: G

Good scavenger:

Y	SiliaMetS Metal Scavengers Selection Table								
Sca	venger	Silia <i>MetS</i> Thiol (<i>Si-Thiol</i>)	Silia <i>MetS</i> DMT (<i>Si-DMT</i>)	Silia <mark>Bond</mark> Amine (Si-WAX)	Silia <i>MetS</i> AMPA (<i>Si-AMPA</i>)	Silia <i>MetS</i> Cysteine (<i>Si-CYS</i>)	Silia <i>MetS</i> DEAM (<i>Si-DEAM</i>)	Silia <i>MetS</i> Diamine (<i>Si-PSA</i>)	
PN		R51030B	R79030B	R52030B	R85130B	R80530B	R54430B	R49030B	
Loa	ding (<i>mmol/g</i>)	≥ 1.20	≥ 0.50	≥ 1.20	≥ 0.80	≥ 0.30	≥ 0.85	≥ 1.28	
Тур	ical Density (g/mL)	0.682	0.732	0.700	0.707	0.665	0.691	0.728	
	Ag								
	AI								
	Ar								
	As								
	Au								
	Bi								
	Ca								
	Cd		•						
	Се								
	Co			•					
	Cr								
	Cs								
	Cu								
	Fe								
	Gd								
	Hg								
þ	lr								
enge	La								
scav	Li								
be s	Mg								
ls to	Mn								
leta	Ni								
2	Os								
	Pb								
	Pd								
	Pt								
	Re								
	Rh								
	Ru								
	Sb								
	Sc								
	Se								
	Sn								
	Ti								
	U								
	v								
	W								
	Zn								
	Zr								



When selecting a metal scavenger, every parameter must be considered: metal catalyst, solvent, residual reagents, by-products, structure of the API (*or molecule of interest*) and temperature.

SiliaMetS Metal Scavengers Selection Table								
	Silia <i>MetS</i> Imidazole (Si-IMI)	Silia <i>MetS</i> TAAcOH (<i>Si-TAAcOH</i>)	Silia <i>MetS</i> TAAcONa (<i>Si-TAAcONa</i>)	Silia <i>MetS</i> Thiourea (<i>Si-THU</i>)	Silia <i>MetS</i> Triamine (Si-TRI)	Silia <mark>Bond</mark> Tosic Acid (Si-SCX)	Scavenger	
	R79230B	R69030B	R69230B	R69530B	R48030B	R60530B	PN	
	≥ 0.96	≥ 0.41	≥ 0.41	≥ 1.07	≥ 1.11	≥ 0.54 meq/g	Loading (<i>mmol/g</i>)	
	0.681	0.635	0.712	0.767	0.736	0.698	Typical Density (g/mL)
							Ag	
							AI	
							Ar	
							As	
							Au	-
							Bi	
							Ca	
							Cd	
							Ce	
							Co	
							Cr	
							Cs	
							Cu	
							Fe	-
							Gd	
							Hg	
							lr	σ
							La	nge
							Li	cave
							Mg	be so
							Mn	tok
							Ni	etals
							Os	ž
					•		Pb	
							Pd	_
							Pt	
						_	Re	
		•					Rh	
							Ru	
_							Sb	
							Sc	
							Se	
					-		Sn	
							Ti	
							U	
			-				v	
							w	
							Zn	
						-	Zr	
			1					



SiliaBond Organic Scavengers Portfolio

SiliaBond Organic Scavengers Technical Information							
Scavengers	Structure	Nature	Molecules Removed	Typical Characteristics 1,2			
Silia <mark>Bond</mark> Amine		Scavenger for Electrophiles (Covalent Bonding)	Acyl Chlorides, Aldehydes, Anhydrides, Chloroformates, Isocyanates, Ketones & Sulfonyl Chlorides	Color: Off-white Density: 0.700 g/mL			
Loading: ≥ 1.20 mmol/g Endcapping: Yes		Scavenger for Acids (<i>Ionic Bonding</i>) Catch & Release	Acids & Acidic Phenols	Prolonged Storage: 2 Shelf Life: 2 Years			
SiliaBond Carbamate PN: R50130B Loading: ≥ 1.16 mmol/g Endcapping: Yes	S. Hyo.	Scavenger for Nucleophiles (Covalent Bonding)	Alcohols, Alkoxides, Amines, Anilines, Hydrazines, Organometallics, Thiols & Thiolates	Color: Off-white Density: 0.741 g/mL Solvent Compatibility: 3 Prolonged Storage: 2 Shelf Life: 2 Years			
SiliaBond Carbonate PN: R66030B Loading: ≥ 0.46 mmol/g Endcapping: Yes		Scavenger for Acids (<i>Ionic Bonding</i>) Catch & Release	Acids, Acidic Phenols & Boronic Acids	Color: Off-white Density: 0.608 g/mL Solvent Compatibility: 3 Prolonged Storage: 2 Shelf Life: 1 Years			
SiliaBond Carboxylic Acid PN: R70030B Loading: ≥ 0.92 mmol/g Endcapping: Yes	Сартин	Scavenger for Bases (<i>Ionic Bonding</i>) Catch & Release	Primary / Secondary Amines & Anilines	Color: Off-white Density: 0.687 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years			
SiliaMetS DEAM PN: R54430B Loading: ≥ 0.85 mmol/g Endcapping: Yes	Сон	Scavenger for Electrophiles & Lewis Acids (Covalent & Ionic Bonding) Catch & Release	Boronic Acids	Color: Off-white Density: 0.691 g/mL Solvent Compatibility: 1 Prolonged Storage: 2 Shelf Life: 2 Years			
SiliaMetS Diamine PN: R49030B	Garren Mer	Scavenger for Electrophiles (Covalent Bonding)	Acyl Chlorides, Aldehydes, Anhydrides, Chloroformates, Isocyanates, Ketones & Sulfonyl Chlorides	Color: Off-white Density: 0.728 g/mL Solvent Compatibility: 2			
Loading: ≥ 1.28 mmol/g Endcapping: Yes		Scavenger for Acids (<i>Ionic Bonding</i>) Catch & Release	Acids & Acidic phenols	Prolonged Storage: 2 Shelf Life: 2 Years			
SiliaBond Diol PN: R35030B Loading: ≥ 0.97 mmol/g Endcapping: No	С стран	Scavenger for Electrophiles & Lewis Acids (Covalent & Ionic Bonding) Catch & Release	Boronic Acids	Color: Off-white Density: 0.687 g/mL Solvent Compatibility: 1 Prolonged Storage: 1 Shelf Life: 2 Years			
SiliaBond DMAP PN: R75630B Loading: ≥ 0.53 mmol/g Endcapping: Yes		Scavenger for Electrophiles (<i>Covalent Bonding</i>)	Acyl Chlorides & Sulfonyl Chlorides	Color: Light brown to brown Density: 0.674 g/mL Solvent Compatibility: 1 Prolonged Storage: 3 Shelf Life: 1 Years			
SiliaBond Guanidine PN: R68230B Loading: ≥ 0.80 mmol/g Endcapping: Yes		Scavenger for Acids (<i>Ionic Bonding</i>) Catch & Release	Acids, Acidic Phenols & Boronic Acids	Color: Light yellow Density: 0.732 g/mL Solvent Compatibility: 1 Prolonged Storage: 2 Shelf Life: 2 Years			



	SiliaBond Organic Scavengers Technical Information								
	Scavengers	Structure	Nature	Molecules Removed	Typical Characteristics ^{1, 2}				
pgl-S	SiliaBond Piperazine		Scavenger for Electrophiles (Covalent Bonding)	Acyl Chlorides, Aldehydes, Anhydrides, Chloroformates, Isocyanates, Ketones & Sulfonyl Chlorides	Color: Off-white Density: 0.671 g/mL				
	Loading: ≥ 0.83 mmol/g Endcapping: Yes	U VIII	Scavenger for Acids (<i>lonic Bonding</i>) Catch & Release	Acids & Acidic Phenols	Prolonged Storage: 2 Shelf Life: 2 Years				
pGI-S	SiliaBond Propylsulfonic Acid PN: R51230B Loading: ≥ 0.63 meq/g Endcapping: Yes	Э с с с с с с с с с с с с с с с с с с с	Scavenger for Bases (lonic Bonding)	Aminos & Apilipos	Color: Off-white Density: 0.728 g/mL Solvent Compatibility: 2 Prolonged Storage: 1 Shelf Life: 2 Years				
pGI-S	SiliaBond Tosic Acid PN: R60530B Loading: ≥ 0.54 meq/g Endcapping: Yes		Catch & Release	Animes & Animes	Color: Off-white Density: 0.698 g/mL Solvent Compatibility: 2 Prolonged Storage: 1 Shelf Life: 2 Years				
	SiliaBond TMA Acetate PN: R66430B Loading: ≥ 0.71 mmol/g Endcapping: No	Government of the second	Scavenger for Acids (<i>lonic</i> <i>Bonding</i>) Catch & Release	Carboxylic Acids	Color: Off-white Density: 0.665 g/mL Solvent Compatibility: 1 Prolonged Storage: 2 Shelf Life: 2 Years				
pgl-5	SiliaBond Tosyl Chloride PN: R44030B Loading: ≥ 0.63 mmol/g Endcapping: Yes		Scavenger for Nucleophiles (Covalent Bonding)	Alcohols, Alkoxides, Amines, Anilines, Hydrazines, Organometallics, Thiols & Thiolates	Color: Off-white Density: 0.761 g/mL Solvent Compatibility: 4 Prolonged Storage: 3 Shelf Life: 6 months				
PGI-S	SiliaMetS Triamine	anni	Scavenger for Electrophiles (<i>Covalent Bonding</i>)	Acyl Chlorides, Aldehydes, Anhydrides, Chloroformates, Isocyanates, Ketones & Sulfonyl Chlorides	Color: Off-white Density: 0.736 g/mL Specification				
	oading: ≥ 1.11 mmol/g indcapping: Yes	υ μ · · · ΝΗ ₂	Scavenger for Acids (<i>Ionic</i> Bonding) Catch & Release	Acids & Acidic Phenols	Prolonged Storage: 2 Shelf Life: 2 Years				

¹ Solvent Compatibility:

- 1- All solvents, aqueous and organic
- 2- All organic solvents
- 3- Anhydrous aprotic solvents
- 4- Anhydrous aprotic solvents, unstable in DMF

² Prolonged Storage:

- 1- Keep dry
- 2- Keep cool (< $8^{\circ}C$) and dry
- 3- Keep cool (< $8^{\circ}C$), dry, and under inert atmosphere



Potentially Genotoxic Impurities (*PGI*) Scavenger See pages 16-17 or contact us for more information



Synthesis

Chromatography

Sample Preparation

Analysis

R&D Services

SiliaBond Organic Scavengers Selection Table

Silia*Bond* organic scavengers can help you purify your API. The functional group is bound to the silica, that will specifically react with a given product. Use the double-entry chart below to choose the best match for the impurity you are dealing with.

\mathcal{L}	SiliaBond Organic Scavengers Selection Table								
Organic Scavenger	Silia <mark>Bond</mark> Amine (<i>Si-WAX</i>)	Silia <mark>Bond</mark> Carbamate (Si-CAR)	Silia <mark>Bond</mark> Carbonate (Si-CO ₃)	Silia <mark>Bond</mark> Carboxylic Acid (Si-WCX)	Silia MetS DEAM (Si-DEAM)	Silia MetS Diamine (Si-PSA)	Silia <mark>Bond</mark> Diol nec (Si-Diol)		
Product Number	R52030B	R50130B	R66030B	R70030B	R54430B	R49030B	R35030B		
Loading (<i>mmol/g</i>)	≥ 1.20	≥ 1.16	≥ 0.46	≥ 0.92	≥ 0.85	≥ 1.28	≥ 0.97		
Typical Density (g/mL)	0.700	0.741	0.608	0.687	0.691	0.728	0.687		
Solvent Compatibility	All organic solvents	Anhydrous aprotic solvents	Anhydrous aprotic solvents	All aqueous & organic solvents	All aqueous & organic solvents	All organic solvents	All aqueous & organic solvents		

Functionalities to be scavenged

Acids & Acidic Phenols		▼				
Acyl & Sulfonyl Chlorides	•				•	
Alcohols & Alkoxides		▼				
Aldehydes, Anhydrides, Chloroformates, Isocyanates & Ketones	•				•	
Amines & Anilines		▼				
Boronic Acids				•		••
Carboxylic Acids						
Hydrazines		▼				
Organometallics, Thiols & Thiolates		▼				

Potential Genotoxic Impurities (PGI) Scavengers Selection Table

Eleven compounds containing structurally alerting functional groups were studied and scavenged by at least one of our grafted silica.

Dest scavenger.	Guou scaverige	ı. •								
	Scav	Scavenging Affinity for Various Potentially Genotoxic Impurities								
Alerting Functional Groups	Acetamide	Pyridine N-oxide	Aniline	Phenyl- hydroxylamine	Benzaldehyde	Octaldehyde				
Silia <i>MetS</i> DMT				-						
SiliaMetS Thiourea										
SiliaMetS Triamine										
SiliaMetS TAAcOH										
SiliaMetS TAAcONa										
SiliaBond Tosic Acid										
Silia <mark>Bond</mark> Tosyl Chloride										





Electrophile scavenger (*covalent bonding*): •

Nucleophile scavenger (covalent bonding):

Catch and release:

Ionic bonding: 🔵

	SiliaBond Organic Scavengers Selection Table								
Silia <mark>Bond</mark> DMAP (Si-DMAP)	Silia <mark>Bond</mark> Guanidine (Si-GUA)	Silia <mark>Bond</mark> Piperazine (Si-PPZ)	Silia Bond Propylsulfonic Acid (<i>Si-SCX-2</i>)	Silia <mark>Bond</mark> Tosic Acid (Si-SCX)	Silia <mark>Bond</mark> TMA Acetate nec (Si-SAX ₂)	Silia <mark>Bond</mark> Tosyl Chloride <i>nec (Si-TsCl</i>)	Silia <i>MetS</i> Triamine (<i>Si-TRI</i>)		
R75630B	R68230B	R60030B	R51230B	R60530B	R66430B	R44030B	R48030B		
≥ 0.53	≥ 0.80	≥ 0.83	≥ 0.63 meq/g	≥ 0.54 meq/g	≥ 0.71	≥ 0.63	≥ 1.11		
0.674	0.732	0.671	0.728	0.698	0.665	0.761	0.736		
All aqueous & organic solvents	All aqueous & organic solvents	All organic solvents	All organic solvents	All organic solvents	All aqueous & organic solvents	Anhydrous aprotic solvents, unstable in DMF	All organic solvents		

•	•			•
	•			•
			▼	
			▼	
			▼	

Scavenging Affinity for Various Potentially Genotoxic Impurities									
Methylmethane sulfonate	Propiolactone	Benzyl bromide	Allyl bromide	1,2-epoxyoctane	Alerting Functional				
0, 0 ~S~0~		Br	<i>₽</i> ►► Br		Groups				
					Silia <i>MetS</i> DMT				
					SiliaMetS Thiourea				
					SiliaMetS Triamine				
					SiliaMetS TAAcOH				
					Silia <i>MetS</i> TAAcONa				
					SiliaBond Tosic Acid				
					SiliaBond Tosyl Chloride				

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Typical Experimental Procedures

In Batch Reactor Mode (Bulk)

Ya.	Typical Experimental Procedures
STEP	For initial screening, start with 4 - 8 molar equivalents* of Silia <i>MetS /</i> Silia <i>Bond</i> in respect to the residual metal or organic impurity concentration.
1	Dissolve the crude product to be treated in a suitable solvent (or directly use the crude reaction <i>mixture</i>) and prepare vials containing the same solution volume.
T	Directly add your chosen Silia <i>MetS /</i> Silia <i>Bond</i> to these vials.
	Note: no pre-wetting / pre-activation is required.
2	For initial tests, stir the solution for at least one hour at room temperature.
	Scavenging progress can be followed by normal analytical techniques.
	The scavenging progress can also be estimated by looking at the color of the solution, as depicted herein:
3	
	When the scavenging is almost complete, the solution is less colored and Silia <i>MetS /</i> Silia <i>Bond</i> becomes colored.
	In some occasional cases, if all the samples are still colored, try one or all of the following: let them react for a longer period of time; add more equivalents of the Silia <i>MetS /</i> Silia <i>Bond</i> ; increase the temperature of the reaction.
	Keep in mind that colors can come from other factors unrelated to the impurities.
4	At the end of the scavenging, filter off the scavenger using a fritted funnel or filtration device.
5	Wash the Silia <i>MetS</i> / Silia <i>Bond</i> with additional solvent for total recovery of the API (or <i>compound of interest</i>) and concentrate the solution under vacuum.
6	Analyze the residual metal or organic impurity concentration of each vial to identify the most efficient scavenger.
7	Direct scale-up is now possible. Otherwise, scavenging optimization can be examined.



Watch the step-by-step experimental procedure video



Take a look at our FAQ about how to calculate the amount of scavenger needed

* Please see page 20 for calculation exemple. Please keep in mind that the above procedures are standard and introductory, but optimization of conditions is key to optimal scavenging efficiency.

In SPE or Flash Cartridges

Fixed-bed formats are a great alternative for metal or organic removal and are directly scalable. We suggest initial screening investigations to be done using Silia*Prep* 2 g / 6 mL SPE cartridges.

	Typical Experimental Procedures
STEP	Description
1	Condition the cartridge with 1 column volume using the same solvent as the solution to be treated.
2	Add the solution containing the API and the metal or organic impurity to the top of the cartridge and let it pass through the cartridge under gravity.
2	Note : if needed, a slight positive pressure on the top of the cartridge or a light vacuum at the bottom can be applied to speed up the flow rate.
2	For most Silia <i>MetS</i> metal scavengers, a dark colored band will be observed on the top of the silica bed.
3	Note : if the residual solution is still colored, multiple passes through the same cartridge can be done.
4	Once the scavenging is completed, wash the cartridge using at least 1 column volume of solvent to ensure total API recovery.



See the 5 steps of SPE here



In E-PAK Cartridges

Y)	Typical Experimental Procedures						
STEP	Description						
1	Ensuring that the selected E-PAK cartridge has been installed, it is recommended to flush the system with at least 3 tank volumes using a compatible solvent. This will remove loose silica / carbon fines and other particulates from the system that could plug the back-pressure control valve. This is best accomplished using solvent delivered at \geq 10 mL/min.						
2	Use bleed valve to remove trapped air from the housing then close the valve and pump at least 3 tank volumes of solvent through the system. The solvent should be collected and discarded. Typical tank volume per cartridge size can be found in the table page 7.						
3	Once the system flushing is completed, set the pump to a low flow rate.						
4	Fully open the air bleed valve on top of the head of the housing.						
	Turn the pump on and watch for test fluid exiting from the bleed valve.						
5	Note : over the course of the test run, additional gas produced due to degassing from the pump action can accumulate in the housing. Periodically this gas should be removed through the bleed valve to prevent interference with even flow distribution through the adsorbent cartridge.						
6	When the bleed valve begins discharging liquid, indicating that all air has been bled from the system, close the bleed valve.						
7	Observe the pressure gauge after closing the bleed valve. Pressure should increase slowly. If pressure increases rapidly, immediately shut the pump off and disassemble the cartridge housing and inspect for blockage. If no blockages are identified, consider feed material characteristics; high viscosity or high solids can create pressure difficulties.						
	Note : we recommend that users pre-filter test solutions containing a large amount of insoluble solids.						
0	Begin sample loading on the cartridge and collection at intervals according to desired data needs or circulate the solution for the number of desired runs.						
ŏ	Note : during sample collection, periodically monitor flow and pressure, adjusting metering valve and pump setting as necessary to maintain target range.						
9	Once the process is completed, flush and collect the system with 2 or 3 tank volumes of fresh clean compatible solvent.						
10	When no more liquid is collected from the outlet valve, turn off the pump.						



See how easy it is working with E-PAK!

Scavenging

Synthesis

Chromatography

E-PAK Housings Installation and Operation Manuals

To get your copies of our Installation and Operation Manuals, contact us at scavengers@silicycle.com







Experimental Procedure Optimization in Bulk

Following a screening test whereby the most performant scavenger has been selected, optimization steps can then be undertaken. There are nine main parameters that can be adjusted either independently one at a time or simultaneously, to improve overall scavenging efficiency. By experience, some are known to influence scavenging more than others and have therefore been grouped in three main steps, classified in order of efficiency.

1. First Bulk Optimization Step

1.1 Reaction Time

Increasing the contact time of the scavenging process can lead to improved efficiency. In fact, scavenging should be pursued until a plateau is reached.

See the example on the right and you can take a look at the <u>Application Note Appn_EP008-0</u> where the scavenging was done using the E-PAK technology, but nevertheless the same concept is applicable to the bulk media.

Ruthenium scavenging yield using E-PAK DMT at 65°C in toluene



1.2 Amount of Scavenger

The amount of scavenger can play a role and should be fine tuned once the initial screening is completed.

Two different methods are suggested to calculate the amount of scavenger required: the molar equivalent method and the weight / weight method. The first makes it possible to calculate the exact number of equivalents to add, however it also requires having previously determined the level of metal contamination. On the other hand, the weight / weight method will allow to skip the pre-analysis step at the expense of risking an underestimate / overestimate of the quantity really needed.

Molar Equivalent Method

For initial screening experiments, we suggest using 4 - 8 molar equivalents of scavengers relative to the concentration of the residual impurity. Once the optimized number of equivalents is selected, redo the calculation for the complete batch to purify.

Here is how to calculate the amount of scavenger to use:



For example, if you need to scavenge 800 g of material containing 500 ppm (*mg/kg*) of Pd (*MW of Pd: 106.42 g/mol*) with Silia*MetS* Thiol with a loading of 1.2 mmol/g and you want to add 4 molar equivalents:

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500 mg				1 mol		1 g Silia <i>MetS</i> or Silia <mark>Bond</mark>		
kg	×	0.800 kg	×	106.42 g	×	1.2 mmol	×	4 equiv. = 12.5 g of Silia <i>MetS</i> or Silia <i>Bond</i>



The Weight / Weight Method

For initial screening experiments we suggest starting your tests with a 20 % w / w ratio of scavenger vs crude (a higher quantity might be required for high levels of impurities). Once the optimized number of equivalents is selected, redo the calculation for the complete batch to purify.



For example, if you need to scavenge 800 g of material containing an unknown quantity of Pd.

800 g × 20 % = 160 g of SiliaMetS or SiliaBond

1.3 Temperature

Generally, it is recommended to perform initial screenings at room temperature, and scavenging is usually completed within a few hours. When shorter scavenging times are required, higher scavenging rates can be achieved by increasing the temperature as shown below. Increasing temperature not only improves the kinetic of the reaction, but it can also improve the scavenging efficiency. The SiliaMetS and SiliaBond products can be safely used without degradation at temperatures up to 150°C.





Residual concentration (%) of Pd(OAc), with SiliaMetS Thiol in DMF

Scavenging



Residual Palladium Concentration (%)

2. Second Bulk Optimization Step

2.1 Combination of Scavengers

A strategy that may be worthwhile is to mix multiple scavengers together, as some have shown to exhibit a synergistic effect that can produce a higher scavenging yield than either one independently. For example, in the <u>Application Note Appn_SM006-0</u>, mixing scavengers achieved greater scavenging yields for Rhodium than using the same amount individually.

2.2 Subsequent Treatments

In some cases (*equilibrium process or the presence of multiple species*), multiple treatments in a row are preferable over a single larger treatment. It should be noted that for optimal results, filtration between each treatment must be carried out. An example of this method is shared in the Case Study CS_SM010-0. For their experiment, the researchers performed two subsequent treatments with Silia*MetS* Thiol in order to reduce the Palladium concentration to the desired amount instead of doing one larger batch treatment.

2.3 Pre-treatment with Activated Carbon

For large quantities of metal removal, a good strategy could be to pre-treat the solution with Silia*Carb* activated carbon prior to the scavenger treatment. This is an approach sometimes used by SiliCycle's researchers.

For example, in the <u>Application Note Appn_EP002-0</u>, a pre-treatment with activated carbon greatly reduced the Palladium concentration prior to the metal scavenger treatment.

3. Third Bulk Optimization Step

Finally, if the two first sets of parameters do not provide a satisfying level of impurities removal, a final third set can be attempted. While these tend to be less effective than the previous ones described above, their effect on scavenging has been helpful in some cases. Hence why they are still recommended.

3.1 Solvent

Our scavengers can be safely used in a wide range of organic and aqueous solvents commonly used in laboratory and in process, such as DMF, DMSO, THF, 2-butanone, alcohols, ethers, chlorinated solvent, etc.

Some scavengers should not be used under aqueous conditions, they are stable only in organic solvents. Others are solely compatible with aprotic solvents. One is also unstable in DMF.

As demonstrated in the graph below, the nature of the solvent can sometimes influence the scavenging efficiency. If scavenging or kinetics are too slow, changing the solvent or adding a co-solvent should be considered.



Pd(PPh₃)₄ (500 ppm) removal in various solvents

3.2 Mixing Rate

SiliCycle's scavengers are mechanically stable and offer excellent scavenging efficiency in batch processes agitated by overhead stirrers as well as orbital shaking under low to moderate agitation rates. If required, mixing rates can be increased to get better scavenging results. With faster stirring, scavenger dispersion in solution is improved. Since silica-based products have an excellent mechanical stability, no worries grinding nor crushing the Silia*MetS* and Silia*Bond* products by applying a more vigorous stirring.

3.3 pH of the Aqueous Solution

When the scavenging is carried out in aqueous conditions, our recommended pH range is between 2 and 10. Depending on the nature of the scavenging agent, the pH can modify the functional groups present on the scavengers by charging them, thereby possibly affecting scavenging (*e.g.: amine groups in acidic conditions*). The various species in solution could also be affected by the pH variation modifying electronic effects, such as H-bonding for example, which could also have an influence on the scavenging efficiency.

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Experimental Procedure Optimization in E-PAK

Following the screening tests described for bulk scavengers where the best Silia*Carb* and / or Silia*MetS* – Silia*Bond* have been identified, the optimization of the experimental conditions can start. There are 5 parameters that can be optimized to improve the scavenging efficiency with E-PAK.

1. First E-PAK Optimization Step

1.1 Amount of Scavenger

Both the molar equivalent and the weight / weight methods (*pages 20 - 21*) are suitable to calculate the amount of scavenger needed for the reaction. Once this has been determined, the corresponding E-PAK cartridge needs to be selected using the table below.

	E-PAK Cartridge Selection	
Weight of Silia <i>Flash</i> , Silia <i>MetS</i> and Silia <mark>Bond</mark> (g)	Weight of Silia <i>Carb</i> (g)	Cartridge Size Required (cm)
0 to 8	0 to 5	5 × 1
> 8 to 75	> 5 to 50	5 × 10
> 75 to 200	> 50 to 125	5 × 25
> 200 to 875	> 125 to 550	16.5 × 12.5
> 875 to 1,750	> 550 to 1,100	16.5 × 25
> 1,750 to 3,500	> 1,100 to 2,100	16.5 × 50
> 3,500 to 7,000	> 2,100 to 4,100	16.5 × 100

1.2 Residence Time

The residence time or contact time refers to the time the solution spend in the E-PAK cartridge. We suggest to start your experiment with a residence time of 2.5 minutes, as it was experimentally determined that this gave the best results. However, depending on the conditions and the type of impurities the results can vary. The residence time is correlated to the flow rate of the experiment: a higher flow rate will give a shorter residence time, while a lower flow rate will give a longer residence time.

2. Second E-PAK Optimization Step

2.1 Selecting the Method

When using E-PAK, four methods are possible to maximize your scavenging yield (see figure on the right).

- **Single Pass**: consists on passing the solution through the E-PAK housing once, yielding the desired purity.
- **Recirculation:** which is the most common, consists on recirculating the solution until getting the desired scavenging yield.
- Multiple Sequential Passes: the solution is passing through like a single pass, but several times.
- Series Pass: consists on having 2 or more housings connected in series where you can recirculate the solution. In this case you can have different adsorbants or the same one.

2.2 Pre-treatment with Activated Carbon

Similar to the Silia*MetS*, if a large amount of contaminant needs to be removed, a pre-treatment with an E-PAK Silia*Carb* activated carbon cartridge could reduce it significantly, allowing the next scavenging step to get to the desired purity level (*for more infomation, take a look at the <u>Application Note Appn EP002-0</u>).*

2.3 Temperature

Keeping in mind that the housings do not have any heating element and that the E-PAK cartridges are made with some polymers, it is possible to increase the temperature up to a maximum of 80°C to see if any significant increase in scavenging yield is noticed. When using the lab scale housing, the heat dissipation can be negligeable, therefore getting your solution in the housing at the right temperature could be done. However, when using pilot and commercial scale housings, an alternative will have to be found to minimize the heat dissipation (*heating up the pipes, etc.*).

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Analysis

Ordering Information

Batch Reactor Mode (Bulk)

All scavengers particle size and pore size are respectively 40 - 63 μm and 60 Å, and are available from 10 g to 25 kg formats (*up to multi-ton scale!*). To build your own product number, just add the Format Code to the Phase PN: [Phase PN]-[Format Code] Example: 100 g of Thiol silica gel, 40 - 63 μm, 60 Å: R51030B-100G

Silia <i>MetS</i> Metal Scavengers							
Scavenger Name	Scavenger PN						
SiliaMetS AMPA	R85130B						
SiliaBond Amine (WAX)	R52030B						
SiliaMetS Cysteine	R80530B						
SiliaMetS DEAM	R54430B						
SiliaMetS DOTA	R91030B						
SiliaMetS DMT	R79030B						
SiliaMetS Diamine	R49030B						
SiliaMetS Imidazole	R79230B						
SiliaMetS TAAcOH nec	R69030B						
SiliaMetS TAAcONa nec	R69230B						
SiliaMetS Thiol	R51030B						
SiliaMetS Thiourea	R69530B						
SiliaBond Tosic Acid (SCX)	R60530B						
SiliaMetS Triamine	R48030B						

SiliaBond Organic Scavengers			
Scavenger Name	Scavenger PN		
SiliaBond Amine (WAX)	R52030B		
SiliaBond Carbonate	R66030B		
SiliaBond Carboxylic Acid (WCX)	R70030B		
SiliaMetS DEAM	R54430B		
SiliaBond DMAP	R75630B		
SiliaMetS Diamine	R49030B		
SiliaBond Diol	R35030B		
SiliaBond Guanidine	R68230B		
SiliaBond Carbamate	R50130B		
SiliaBond Piperazine	R60030B		
SiliaBond Propylsulfonic Acid (SCX-2)	R51230B		
SiliaBond TMA Acetate nec	R66430B		
SiliaBond Tosic Acid (SCX)	R60530B		
SiliaBond Tosyl Chloride	R44030B		
SiliaMetS Triamine	R48030B		

All activated carbons are available in 100 g, 1 kg, and 20 kg packagings.

Silia <i>Carb</i> Activated Carbon			
Carbon Name	Carbon PN		
SiliaCarb CA	C-CA		
SiliaCarb HA	C-HA		
SiliaCarb VA	C-VA		
SiliaCarb VW	C-VW		

Y.	Bulk Formats				
Quantity	Code	Quantity	Code		
5 g	5G	500 g	500G		
10 g	10G	1 kg	1KG		
25 g	25G	5 kg	5KG		
50 g	50G	10 kg	10KG		
100 g	100G	20 kg	20KG		
250 g	250G	25 kg	25KG		

Available Kit

Because all matrices are unique, and that small differences can influence scavenging efficiency, we recommend trying first our scavenger kit for screening purposes, especially if you are new to this technology. Steric hindrance of the catalyst, electronic effects, solubility in solvents, all are factors that can influence the removal of your impurity.

<u>How to order</u>: simply note the kit number which starts with "K", add a dash mark and your choice of format. Example: K30730B-10G to obtain 10 g of each one of the scavengers listed in the kit.

SiliaMetS Metal Scavenger Kit				
Kit Name	Kit Number	Composition		
SiliaMetS Universal Metal Scavenger Kit	K30730B	Cysteine, DMT, Imidazole, TAAcOH, TAAcONa, Thiol, Thiourea & Triamine		

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This kit is available in 5 g, 10 g, 25 g, 50 g, and 100 g formats.



E-PAK Cartridges

Y.	E-PAK Cartridges						
Type	Labora	tory Scale - 5 cm D	/ Scale - 5 cm Diameter Pilot Scale - 16.5 cm Diameter (Commercial Scale	- 16.5 cm Diameter	
туре	1 cm	10 cm	25 cm	12.5 cm	25 cm	50 cm	100 cm
SiliaFlash Irr	egular Silica Gel						
40 - 63 µm	LS-R10030B-51	LS-R10030B-510	LS-R10030B-525	PS-R10030B-1612	PS-R10030B-1625	CS-R10030B-1650	CS-R10030B-16100
60 - 200 µm	LS-R10040B-51	LS-R10040B-510	LS-R10040B-525	PS-R10040B-1612	PS-R10040B-1625	CS-R10040B-1650	CS-R10040B-16100
SiliaMetS Me	tal Scavengers						
Thiol	LS-R51030B-51	LS-R51030B-510	LS-R51030B-525	PS-R51030B-1612	PS-R51030B-1625	CS-R51030B-1650	CS-R51030B-16100
DMT	LS-R79030B-51	LS-R79030B-510	LS-R79030B-525	PS-R79030B-1612	PS-R79030B-1625	CS-R79030B-1650	CS-R79030B-16100
Imidazole	LS-R79230B-51	LS-R79230B-510	LS-R79230B-525	PS-R79230B-1612	PS-R79230B-1625	CS-R79230B-1650	CS-R79230B-16100
AMPA	LS-R85130B-51	LS-R85130B-510	LS-R85130B-525	PS-R85130B-1612	PS-R85130B-1625	CS-R85130B-1650	CS-R85130B-16100
Amine	LS-R52030B-51	LS-R52030B-510	LS-R52030B-525	PS-R52030B-1612	PS-R52030B-1625	CS-R52030B-1650	CS-R52030B-16100
Diamine	LS-R49030B-51	LS-R49030B-510	LS-R49030B-525	PS-R49030B-1612	PS-R49030B-1625	CS-R49030B-1650	CS-R49030B-16100
Triamine	LS-R48030B-51	LS-R48030B-510	LS-R48030B-525	PS-R48030B-1612	PS-R48030B-1625	CS-R48030B-1650	CS-R48030B-16100
TAAcONa	LS-R69230B-51	LS-R69230B-510	LS-R69230B-525	PS-R69230B-1612	PS-R69230B-1625	CS-R69230B-1650	CS-R69230B-16100
Cyano	LS-R38030B-51	LS-R38030B-510	LS-R38030B-525	PS-R38030B-1612	PS-R38030B-1625	CS-R38030B-1650	CS-R38030B-16100
SCX-2	LS-R51230B-51	LS-R51230B-510	LS-R51230B-525	PS-R51230B-1612	PS-R51230B-1625	CS-R51230B-1650	CS-R51230B-16100
SiliaCarb Act	tivated Carbon			-		^	
C-CA	LS-CCA-51	LS-CCA-510	LS-CCA-525	PS-CCA-1612	PS-CCA-1625	CS-CCA-1650	CS-CCA-16100
C-HA	LS-CHA-51	LS-CHA-510	LS-CHA-525	PS-CHA-1612	PS-CHA-1625	CS-CHA-1650	CS-CHA-16100
C-VA	LS-CVA-51	LS-CVA-510	LS-CVA-525	PS-CVA-1612	PS-CVA-1625	CS-CVA-1650	CS-CVA-16100
C-VW	LS-CVW-51	LS-CVW-510	LS-CVW-525	PS-CVW-1612	PS-CVW-1625	CS-CVW-1650	CS-CVW-16100

E-PAK Housings

Lab Scale

Housing Kits for Lab Scale E-PAK				
Housing Size	316L	Hastelloy		
5 × 1 cm	LSHK-1-S	LSHK-1-H		
5 × 10 cm	LSHK-10-S	LSHK-10-H		
5 × 25 cm	LSHK-25-S	LSHK-25-H		
5 × 1 cm + 5 × 10 cm	LSHK-110-S	LSHK-110-H		
5 × 1 cm + 5 × 25 cm	LSHK-125-S	LSHK-125-H		
5 × 1 cm + 5 × 10 cm + 5 × 25 cm	LSHK-11025-S	LSHK-11025-H		

Bowls for Lab Scale E-PAK Housing				
Bowl Length 316L Hastelloy				
1 cm	LS-BOWL1-S	LS-BOWL1-H		
5 cm	LS-BOWL10-S	LS-BOWL10-H		
25 cm	LS-BOWL25-S	LS-BOWL25-H		

Pilot Scale*

	Housings for Pilot Scale E-PAK				
Housing Size	316L Hastelloy				
16.5 × 12.5 cm	PSHSR-12-S	PSHSR-12-H			
16.5 × 25 cm	PSHSR-25-S	PSHSR-25-H			

* For PED certified housing, simply add "-PED" at the end of the product number.

Commercial Scale*

Housings for Commercial Scale E-PAK				
Housing Type 316L		Hastelloy		
Single Round, 16.5 × 50 cm Single Round, 16.5 × 100 cm	CSHSR-50100-S	CSHSR-50100-H		
3-Round, 16.5 × 50 cm 3-Round, 16.5 × 100 cm	CSH3R-50100-S	CSH3R-50100-H		
7-Round, 16.5 × 50 cm 7-Round, 16.5 × 100 cm	CSH7R-50100-S	CSH7R-50100-H		
12-Round, 16.5 × 50 cm 12-Round, 16.5 × 100 cm	CSH12R-50100-S	CSH12R-50100-H		



Fixed-Bed Mode (SPE or Flash Cartridges)



SiliaPrep SPE Cartridges and SiliaSep Flash Cartridges

To build your SPE or flash cartridge Product Number, simply start with the **Prefix SPE** or **FLH**, followed by the Phase Number you wish your cartridge to be packed with (*see page 24*), followed by the Format Code.

- Examples: SiliaPrep Thiourea, 6 mL, 500 mg = SPE-R69530B-06P
 - SiliaSep Open-Top TAAcONa, 70 mL, 10 g = FLH-R69230B-70Y
 - SiliaSep TAAcONa, 4 g = FLH-R69230B-ISO04

SiliaPrep SPE and SiliaSep OT Cartridges					
Formats Available	Prefix	Code	Units / Box		
3 mL / 200 mg	SPE	03G	50		
3 mL / 500 mg	SPE	03P	50		
6 mL / 500 mg	SPE	06P	50		
6 mL / 1 g	SPE	06S	50		
6 mL / 2 g	SPE	06U	50		
12 mL / 2 g	SPE	12U	20		
25 mL / 5 g	FLH	20X	20		
70 mL / 10 g	FLH	70Y	16		
70 mL / 15 g	FLH	70i	16		
70 mL / 20 g	FLH	70Z	16		
150 mL / 25 g	FLH	95K	10		
150 mL / 50 g	FLH	95M	10		
150 mL / 70 g	FLH	95N	10		

SiliaSep Flash Cartridges					
Formats Available	Prefix	Code	Units / Box		
4 g	FLH	ISO04	2		
12 g	FLH	ISO12	1		
25 g	FLH	ISO25	1		
40 g	FLH	ISO40	1		
80 g	FLH	ISO80	1		
120 g	FLH	IS120	1		
220 g	FLH	IS220	1		
330 g	FLH	IS330	1		
800 g	FLH	IS750	1		
1,600 g	FLH	11500	1		



SiliaChrom Guard Cartridges

Silia*Chrom* guard cartridges are available in lengths of 10, 20 and 30 mm and four internal diameters (*ID: 4.0, 10, 21.2 and 30 mm*).

The guard cartridges internal diameter should be the same as the HPLC column or one size smaller. Never use a guard cartridge with a larger ID than the HPLC column (*risk of efficiency loss*).

<u>Best Scavenger for</u>: Pd, Ag, Au, Bi, Hg, Ir, Ni, Os, Pt, Rh, Ru, Se and U. <u>Good Scavenger for</u>: Cd, Co, Cu, Fe, Pb, Sc, Sn and Zn.

SiliaChrom Guard Cartridges						
Formats Available (Internal Diameter × Length in mm)	5 μm PN	10 µm PN	Units / Box	Guard Holder PN		
4.0 × 10	HPLG-K34605E-A-N010	HPLG-K34607E-A-N010	4	HPH-N010		
4.0 × 20	HPLG-K34605E-A-N020	HPLG-K34607E-A-N020	4	HPH-N020		
10 × 10	HPLG-K34605E-A-Q010	HPLG-K34607E-A-Q010	2	HPH-Q010		
21.2 × 10	HPLG-K34605E-A-T010	HPLG-K34607E-A-T010	1	HPH-T010		
30 × 10	HPLG-K34605E-A-V010	HPLG-K34607E-A-V010	1	HPH-V010		
30 × 20	HPLG-K34605E-A-V020	HPLG-K34607E-A-V020	1	HPH-V020		
30 × 30	HPLG-K34605E-A-V030	HPLG-K34607E-A-V030	1	HPH-V030		



Resource Center

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Choosing your metal & organic scavenging solution

Take a Look at some of our Multimedia Contents



Introduction to metal and organic Scavengers



E-PAK flow purification cartridges



SiliCycle's most popular functionalized silicas

SILICYCLE CO

Metal scavenging using bulk

SiliaMetS functionalized silica

SILICYCLE

Scale-up of impurity scavenging with E-PAI

Scale-up impurity

scavenging with E-PAK



SiliCycle packing formats in a nutshell



Scavenging

Synthesis

Chromatography

Sample Preparation



How to calculate the amount of scavenger needed



E-PAK cartridge housings, from lab to commercial scale

SILICYCLE 🦚

Understanding Column

Volume

SILICYCLE 🧑





What are the parameters that influence scavenging efficiency?



See how easy it is working with E-PAK



Flash separation of dye mixture with SiliaSep Premium



The 5 steps of a solid phase extraction (SPE)



What pH range is suitable for functionalized silica?



How does flash chromatography work?



Understanding particle size distribution - D50, D90 and D10



What is the sample mass loading capacity of preparative TLC plates?

What is the relationship between retention factor and column volume



A collection of various case studies and application notes using scavengers

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