

Caremark Achieves Single-Digit Trend 2 Consecutive Years

Caremark pharmacy benefit management techniques:

- · limited the impact of new drugs and specialty products, and
- · optimized the opportunities presented by the expanding number of generics on the market, and
- kept overall pharmacy trend in the single digits for Caremark clients in 2004.

9.9% Trend

4.6% Price - Inflation

0.6% Product – New Drugs, Drug Mix, Biotech Drugs

4.7% **Utilization** – Prescribers, Politics, People, Philosophy

Utilization, at 4.7%, climbed from an exceptionally low 2003 number, but still remained under levels for the previous three years. **Drug price inflation at 4.6%**, while higher than 2003, was moderate compared to 2002 levels of nearly 6%. As trend drivers, **drug mix** and **new drugs** were essentially flat.

Impact of Price Inflation: 4.6%

Traditional brand manufacturers face a number of challenges and uncertainties in addition to a less-than-robust pipeline: numerous pending patent expirations, the impending Medicare drug benefit, product recalls, concerns about drug safety, and questions about the FDA approval process. All these factors may have contributed to **manufacturer price increases** in 2004.

Impact of Utilization: 4.7%

The aging of our population and increases in the incidence of chronic conditions continue to drive utilization increases across all our top categories. Shifts in prescribing patterns related to the release of new medical data had major impact in 2004. Significantly, utilization of our top category, the antihyperlidemics – which include the **statins** – grew substantially (15.7%) driven by new guidelines on cholesterol levels. Conversely, the withdrawal of Vioxx® late in 2004 and subsequent doubts about other **COX-2s** reduced overall utilization in the anti-inflammatory analgesics. Given the recommendations of the FDA COX-2 advisory panel released early in 2005 it is likely that we will see some rebound in COX-2 utilization in the months ahead. Utilization of **antidepressants** continues to grow despite late-year warnings about use in pediatric patients. However, spend in the category was moderated by the increasing availability of generics.

Ulcer drugs, dominated by the proton pump inhibitiors (**PPIs**), actually decreased in utilization, due to the availability of OTC alternatives; management techniques, such as moving the class to the third tier; and clinical strategies such as dose optimization, step down therapy, and prior authorization.

Increases in the **asthma** and **diabetes** categories are related to increasing incidence, more aggressive treatment, and multi-drug regimens for these diseases. The **miscellaneous endocrine** category is dominated by osteoporosis drugs. Utilization of these products has been impacted by greater awareness of the condition as well as the aging of the population. Moreover, migration from less expensive hormone replacement therapy to more expensive osteoporosis products has driven trend for the category upward. Use of **narcotic pain relievers** has become more accepted and their use has been increasing for several years as pain management guidelines evolve. Growth of **anticonvulsants** continues due to widespread and acceptable off-label use for pain relief.

Top Ten Therapeutic Classes Account for More Than 50% of Book of Business Spend							
Rank	Therapeutic Class	Gross Trend	Utilization trend	% of Total Spend			
1	ANTIHYPERLIPIDEMICS	18.5%	15.7%	9.9%			
2	ANTIDEPRESSANTS	6.3%	5.3%	7.5%			
3	ULCER DRUGS	0.7%	-0.8%	7.3%			
4	ANTIHYPERTENSIVES	12.4%	8.5%	5.3%			
5	ANALGESICS - ANTI-INFLAMMATORY	14.0%	1.4%	4.9%			
6	ANTIDIABETICS	11.9%	7.0%	4.6%			
7	ANTIASTHMATICS	16.2%	6.1%	4.3%			
8	MISC. ENDOCRINE	17.9%	10.0%	3.0%			
9	ANALGESICS - NARCOTIC	13.1%	8.6%	3.0%			
10	ANTICONVULSANTS	19.0%	7.5%	2.9%			
Top 10 T	herapeutic Classes, % of total spend			52.8%			

Impact of Product: 0.6%

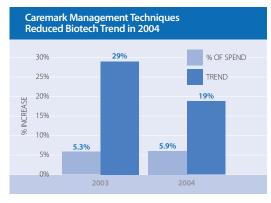
It's especially important to look at 2004 increases in utilization in relationship to our nearly flat numbers for **drug mix** and **new drugs**. There were no immediate blockbuster brand launches in 2004. Several of the significant new launches were brand extensions or products combining existing drugs rather than the breakthrough products that can significantly drive spend. Even innovative new products, such as Spiriva® for chronic obstructive pulmonary disease, and Cymbalta™, a new antidepressant, did not perform as well as expected. Caremark management programs and strategies also helped by limiting utilization of expensive newer drugs.

2004 Launches Did Not Have Significant Impact							
Rank	2004 Top New Drugs	Launch Date	Indication/Comments	% Spend			
1	SPIRIVA	Q2	COPD	0.06%			
2	FOLLISTIM AQ	Q1	Fertility	0.05%			
3	VYTORIN	Q3	Combination product for cholesterol	0.04%			
4	GONAL-F RFF	Q3	Fertility	0.03%			
5	CYMBALTA	Q3	Antidepressant	0.03%			
6	KETEK PAK	Q3	New class of antibiotic	0.02%			
7	CADUET	Q3	Combination Lipitor and Norvasc	0.02%			
8	TRUVADA	Q3	Combination for HIV	0.02%			
9	ZYRTEC	Q2	New dosage form – chewable	0.01%			
10	SENSIPAR	Q2	Specialty for hyperparathyroidism	0.01%			

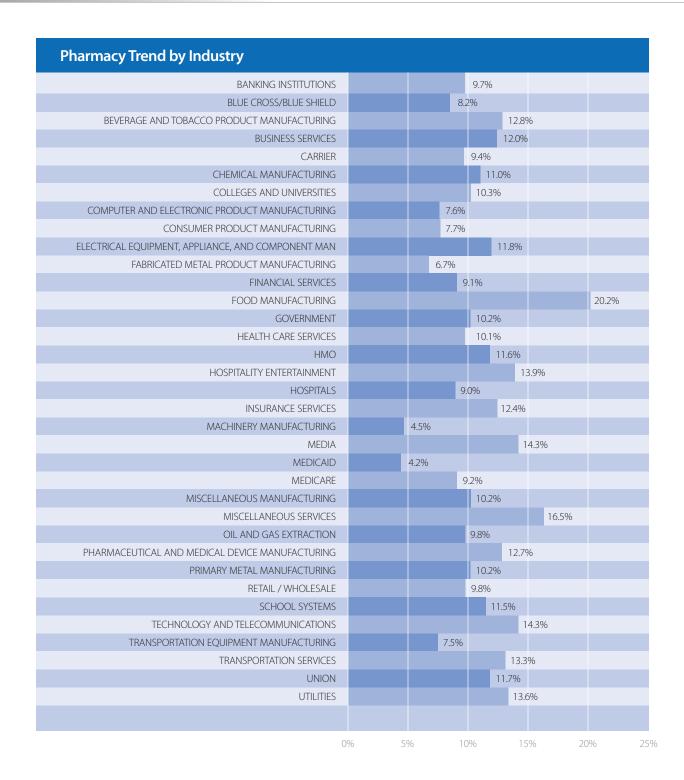
Utilization of **generics** helped keep the drug mix portion of trend low. Our **generic dispensing rate (GDR)** increased by more than five percentage points from 43.7% to 49.5%. **Generic substitution rates (GSR)** increased to 95%. Our Book of Business benefited from the many new generic launches in 2004, including generic forms of Wellbutrin®, Proventil®, Glucovance®, Cipro®, Neurotin®, and Celexa®. Caremark plan participant education as well as targeted retail pharmacist and physician communications are helping to reinforce the message that generics are the good-value choice

Generic Drug Name		Brand Avg Gross Cost/Day Supply*	Generic Avg Gross Cost/Day Supply*	2004 Savings from Generic	
2004 Generic Products					
BUPROPION HCL 150MG	WELLBUTRIN 150 MG	\$3.28	\$2.50	\$331,727,430	
CIPROFLOXACIN SUS, TAB	CIPRO SUS, TAB	\$7.44	\$3.27	\$33,156,452	
OXYCODONE HCL TAB	OXYCONTIN TAB	\$27.45	\$18.73	\$22,169,300	
GLYBURIDE/METFORMIN HCL TAB	GLUCOVANCE TAB	\$2.44	\$1.91	\$20,869,074	
CYCLOSPORINE SOL, CAP	NEORAL SOL, CAP	\$11.59	\$9.01	\$17,575,538	
Sub Total				\$425,497,794	
2003 Generic Products					
SOTALOL HCL TAB	SOTALOL TAB	\$5.93	\$2.58	\$92,939,938	
PAROXETINE HCL TAB	PAXIL TAB	\$2.83	\$2.01	\$73,753,531	
MIRTAZAPINE TAB, ODT	REMERON TAB, ODT	\$2.70	\$1.69	\$25,030,170	
NEFAZODONE HCL TAB	SERZONE TAB	\$3.19	\$1.23	\$19,539,363	
FOSINOPRIL SODIUM TAB	MONOPRIL TAB	\$1.32	\$.88	\$19,323,843	
Sub Total				\$230,586,845	
2002 Generic Products					
LISINOPRIL TAB	ZESTRIL TAB	\$1.20	\$0.45	\$447,610,379	
METFORMIN HCL TAB	GLUCOPHAGE TAB	\$1.95	\$1.14	\$326,001,388	
OMEPRAZOLE CAP	PRILOSEC CAP	\$4.41	\$3.00	\$103,977,235	
TRAMADOL HCL TAB	ULTRAM TAB	\$4.53	\$1.55	\$83,190,181	
OX TR/POTASSIUM CLAVULANATE TAB, CHW, SUS	AUGMENTIN TAB, CHW, SU	IS \$8.06	\$7.33	\$34,225,486	
Sub Total				\$995,004,669	
Total Savings for Top 5 Generics					

^{*} January–December 2004 average



At 5.9 % of our Book of Business overall spend, **biotech** showed only a small increase despite significant new launches in the category. Careful management of all specialty products helped keep specialty spend and trend down for Caremark clients. For example, enhanced guideline management helped ensure that the specialty product was the right drug for a specific patient, limiting inappropriate utilization of these costly products.



Watch for TrendsRx® 2005 Trend Report for detailed analysis of 2004 and our forecasts for 2005.

Trend – the annual increase in the cost of providing prescription benefits to plan participants – is calculated as gross cost per employee per year (PEPY) in the Caremark Book of Business.

Data source: Caremark Book of Business, 2004, Analytics & Outcomes

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