

Drug Mix: The Right Drug

Over the last few years, an increasing number of blockbuster brand drugs faced **generic competition** for the first time. Caremark clients benefited significantly from these opportunities by implementing benefit designs that encouraged utilization of generics and promoting member education about these cost-effective alternatives. Our 2005 **generic dispensing rate (GDR) rose to 52.6%**, up from 48.2% in 2004. Major first-time generics in 2005 included the antihistamine fexofenadine (Allegra®) and the antibiotic azithromycin (Zithromax®). In 2006, the market anticipates a number of new generic entries, including cholesterol drugs pravastatin (Pravachol®) and simvastatin (Zocor®). Generic versions of Flonase® (fluticasone propionate), a nasal corticosteroid; Ambien® (zoldipem tartrate), an insomnia agent; and Zoloft® (sertraline), an antidepressant; are also expected to launch in the next 12 to 15 months.

Seniors' drug spend is dramatically higher

Over 65
\$1,269.85
PMPY

Under 65
\$653.53
PMPY

and their utilization of generics is lower.

50.3%
generic utilization

54.5%
generic utilization

Caremark Book of Business data,
generic utilization for December 2005

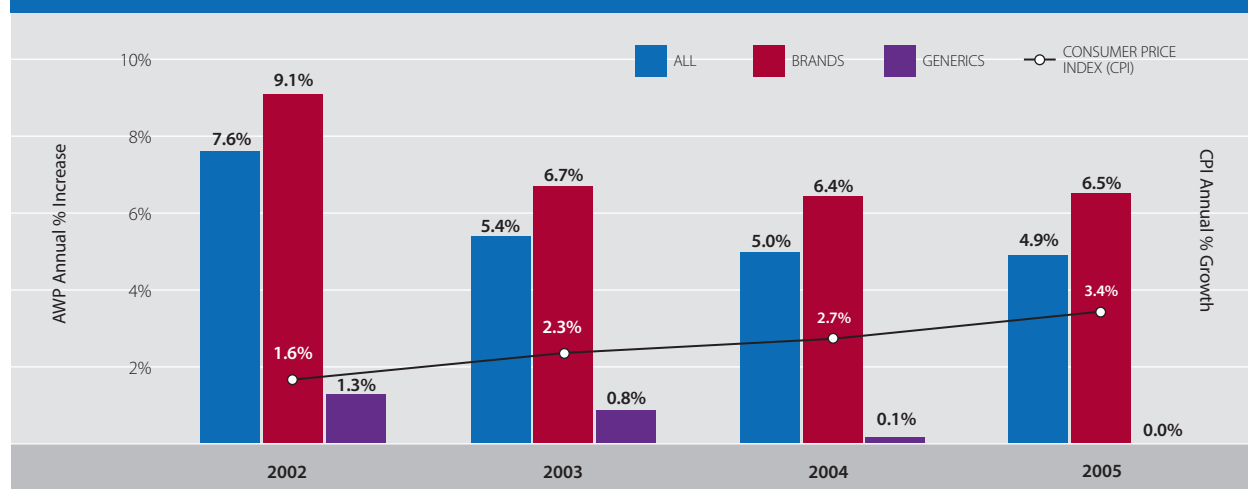
2005 Brand Drug Launches

As has been true for the last few years, the traditional brand drug pipeline produced **few high-performing new products** in 2005, helping to reduce the impact of drug mix as a trend driver. The FDA was cautious, approving only 20 new drugs (includes two biologics) in 2005. This was a decrease of more than 44% from the 36 NME approvals in 2004 (includes five biologics). Notable 2005 launches include the insomnia agents Lunesta® (eszopiclone) and Rozerem™ (ramelteon); Lyrica® (pregabalin), which is indicated for the treatment of pain associated with diabetic peripheral neuropathy; and several products for diabetes including Symlin® (pramlintide) and Byetta® (exenatide).

Price Increases: Generics Hold the Line

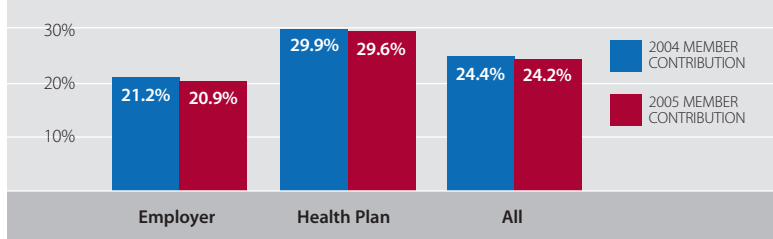
Price inflation was second to utilization as a trend driver in 2005. The rate of brand price increases accelerated, while prices for generics held steady.

Average Wholesale Price Inflation



Caremark Book of Business data

Member Contribution by Market Segment



Member contribution levels remained relatively stable in 2005; plans looked to other strategies help manage trend. Consequently, **BOB net trend at 8.3%** was higher than gross trend.

Caremark 2005 BOB data;
Includes specialty

Top 10 Brand Drugs

| 2005 Rank | 2004 Rank | Drug | Class | % Gross Spend |
|-----------|-----------|---------------|----------------------------------|---------------|
| 1 | 1 | Lipitor | Antihyperlipidemic (cholesterol) | 5.0% |
| 2 | 2 | Nexium | Anti-ulcer Drug | 2.6% |
| 3 | 3 | Prevacid | Anti-ulcer Drug | 2.1% |
| 4 | 6 | Advair Diskus | Antiasthmatic | 1.6% |
| 5 | 5 | Zoloft* | Antidepressant | 1.4% |
| 6 | 10 | Plavix | Anti-platelet medication | 1.4% |
| 7 | 9 | Effexor XR | Antidepressant | 1.3% |
| 8 | 4 | Zocor* | Antihyperlipidemic (cholesterol) | 1.2% |
| 9 | 7 | Pravachol | Antihyperlipidemic (cholesterol) | 1.2% |
| 10 | 12 | Singular | Antiasthmatic | 1.2% |

Drugs ranked by gross cost in the Caremark BOB 2005. Excludes specialty products
*2006 launch of generic equivalent expected

Top 10 Therapeutic Classes by 2005 Gross Cost

| 2005 Rank | 2004 Rank | Class | Gross Trend | Utilization Trend | % Gross Spend |
|-----------|-----------|---|-------------|-------------------|---------------|
| 1 | 1 | Antihyperlipidemics (cholesterol) | 13.7% | 10.5% | 10.6% |
| 2 | 2 | Anti-ulcer drugs | 5.0% | 2.3% | 7.5% |
| 3 | 3 | Antidepressants | -0.3% | 1.3% | 7.0% |
| 4 | 4 | Antihypertensives | 8.6% | 5.3% | 5.3% |
| 5 | 6 | Antidiabetics | 14.1% | 9.7% | 4.9% |
| 6 | 7 | Antiasthmatic and bronchodilator agents | 17.6% | 8.9% | 4.9% |
| 7 | 5 | Analgesics – anti-inflammatory | -13.6% | -23.2% | 4.1% |
| 8 | 10 | Misc. endocrine and metabolic agents | 13.2% | 5.3% | 3.1% |
| 9 | 9 | Analgesics – opioid | 6.0% | 10.6% | 2.9% |
| 10 | 8 | Anticonvulsants | 5.1% | 6.5% | 2.9% |

Asthma drugs, led by Singulair® and Advair Diskus®, posted the highest gross trend of the top therapeutic categories. Anti-inflammatory analgesics, which include the NSAIDs and COX II drugs, had a negative trend as the market con-

tinues to react to the withdrawals of Vioxx® and Bextra® and safety concerns about the class as a whole. The launch of generic antidepressants helped lower trend for the category, which was also affected by safety concerns.

Overview and Projections for Selected Therapeutic Categories

| Therapeutic Category | 2006 Expected Trend | Recent and 2006 Anticipated Brand Launches | Anticipated Generic Launches | Current Generic Opportunity | OTC Opportunity |
|---|---------------------|---|------------------------------|--|----------------------|
| Antihyperlipidemics – Cholesterol-lowering Agents | ▲ | Vytorin, Caduet | Zocor, Pravachol | lovastatin | |
| Anti-ulcer Agents – Proton Pump Inhibitors (PPIs) | ▬ | | | omeprazole | Prilosec OTC |
| Antidepressants | ▼ | | Zoloft | fluoxetine, paroxetine, citalopram, | |
| SSRIs | | | | None available | |
| SNRIs | ▲ | Cymbalta | | | |
| Antihypertensives | | | | | |
| Calcium Channel Blockers (CCBs) | ▬ | | | cartia XT, verapamil SR, nifedical XL | |
| ACE Inhibitors | ▬ | | Aceon | lisinopril, enalapril, quinapril | |
| Angiotensin II Receptor Antagonists (ARBs) | ▲ | | | None available | |
| Antidiabetics | | | | | |
| Insulin Sensitizing Agents | ▲ | ACTOplus met | | None available | |
| Insulin | ▲ | Apidra, Exubera, Levemir | | None available | |
| Antiasthmatic and Bronchodilator Agents | | | | | |
| Sympathomimetics | ▲ | | | albuterol, terbutaline | |
| Anti-inflammatory Analgesics | | | | | |
| NSAIDs | ▬ | Arcoxia, Dynastat | Mobic | ibuprofen, naproxen, nabumetone, oxaprozin | Advil, Motrin, Aleve |
| Misc. Endocrine and Metabolic Agents – Calcium/bone Regulators | | | | | |
| | ▲ | Boniva, Fosamax Plus D, Actonel with Calcium, Oporia, Preos | Miacalcin Nasal | None available | |
| Analgesics, Opioid – Narcotic Agonists | | | | | |
| | ▲ | | | tramadol, fentanyl, morphine sulfate | |
| Anticonvulsants | | | | | |
| | ▲ | Lyrica, Rufinamide | Trileptal, Zonegan | gabapentin, carbamazepine, primidone | |
| MISCELLANEOUS CATEGORIES | | | | | |
| Antihistamines, Non-sedating (NSAs) | ▼ | Allegra-D 24 Hours | Allegra-D | fexofenadine | Claritin, Claritin D |
| Antiplatelet Medication | ▲ | | Plavix | cilostazol, dipyridamole, ticlopidine | |
| Multiple Sclerosis Agents | ▲ | Tysabri (possible relaunch) | | None available | |
| Rheumatoid Arthritis Agents | ▲ | Orencia | | None available | |
| Non-barbiturate Hypnotics | ▲ | Ambien CR, Lunesta, Indiplon | Ambien | temazepam, triazolam, flurazepam | |
| Nasal Steroids | ▼ | | Flonase | flunisolide | |

Arrow Key

- ▲ Estimated greater than 4% increase
- ▬ Estimated flat trend
- ▼ Estimated greater than 4% decrease

Drug Specialty product

These pages contain prescription brand name drugs that are registered or trademarks of pharmaceutical manufacturers that are not affiliated with Caremark.

This information has been compiled from a number of sources and is provided for informational purposes. Due to circumstances beyond our control, information related to prospective drug launches is subject to change without notice. Expected trend is projected based on marketplace events only and does not reflect plan design, clinical programs and other initiatives that may impact trend. This analysis is provided for informational purposes only.

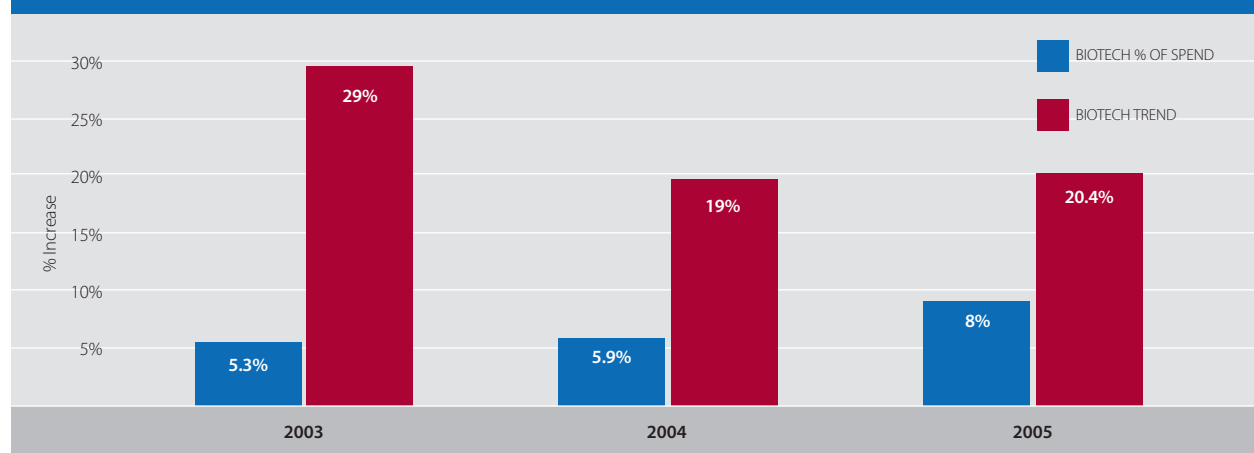
Specialty Trend Driven by Double-Digit Growth in Utilization

Specialty Trend: 20.4%

Specialty Utilization Trend: 11.4%

In 2005, specialty pharmaceuticals grew from less than 6% of our gross BOB spend to 8%. Growth was largely driven by **rapidly increasing utilization**, which grew at more than twice the BOB rate. In addition to the introduction of new specialty products, growth in utilization related to an **ongoing increase in indications** for specialty products already on the market. For example, Humira® (adalimumab) was approved in 2002 for the treatment of symptoms associated with moderate to severe rheumatoid arthritis (RA). Today it has been approved for five additional indications, including the treatment of psoriatic arthritis and plaque psoriasis. Additional indications for Humira are under consideration at the FDA. The **specialty pipeline** continues to be robust. Orencia™ has been approved by the FDA and is expected to launch in 2006, further expanding utilization of specialty products for RA, one of the highest-trending specialty categories. Specialty products for diabetes, macular degeneration and a number of forms of cancer are also pending FDA approval in 2006.

Biotech/Specialty Spend and Trend



2005 Top 10 Biotech/Specialty Products

| 2005 Rank | 2004 Rank | Drug | Common Indication | Drug Utilization Trend |
|-----------|-----------|-----------|---|------------------------|
| 1 | 1 | Enbrel | Rheumatoid arthritis | 26.6% |
| 2 | 2 | Avonex | Multiple sclerosis | -5.7% |
| 3 | 4 | Humira | Rheumatoid arthritis | 38.4% |
| 4 | 3 | Copaxone | Multiple sclerosis | 12.5% |
| 5 | 5 | Procrit | Hematological used for anemia | 6.5% |
| 6 | 6 | Cellcept | Immunosuppressive used with organ transplants | 12.3% |
| 7 | 8 | Gleevec | Antineoplastic used to treat a rare cancer | 16.0% |
| 8 | 9 | Rebif | Multiple sclerosis | 25.0% |
| 9 | 7 | Prograf | Immunosuppressive used with organ transplants | 13.5% |
| 10 | 10 | Betaseron | Multiple sclerosis | 2.2% |

Caremark 2005 Book of Business data

Drugs ranked by gross cost in the Caremark BOB 2005.

Utilization Trend = 2005 Days Supply Per Member Per Month(PMPM) ÷ 2004 Days Supply PMPM

Trend by Industry

| Industry | Gross Cost PMPY Trend |
|--|-----------------------|
| Banking and Financial Services | 8.5% |
| Chemical Manufacturing | 10.5% |
| Colleges and Universities | 8.7% |
| Consumer Products | 11.5% |
| Electrical Equipment and Appliance Manufacturing | 10.7% |
| General Manufacturing | 7.3% |
| Government | 9.4% |
| Healthcare Services | 8.5% |
| Hospitality/Entertainment | 10.4% |
| Hospitals | 6.7% |
| Media | 10.7% |
| Managed Care | 7.4% |
| Pharmaceutical and Medical Device Manufacturing | 10.9% |
| Retail/Wholesale | 8.6% |
| School Systems | 6.3% |
| Technology and Telecommunications | 6.9% |
| Transportation Equipment Manufacturing | 8.4% |
| Transportation Services | 10.2% |
| Union | 8.6% |
| Utilities & Energy | 7.3% |

Caremark Book of Business data for selected categories; trend includes Specialty

Watch for the 2006 TrendsRx® Report for detailed analysis of 2005 and our forecasts for 2006.

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