



## **Comparison of Oral Beta-Blockers**

--Information in chart is from product labeling (see footnote d) unless otherwise noted.--

Agent	Dosing (Adults)	Comments*	Availability
<b>Pharmacokinetics</b>	(also see <b>footnote b</b> )		Cost of 30-day supply <sup>a</sup>
	ts (beta-1 and beta-2 antagonist activity) More likely to worser	n peripheral vasoconstriction or bron	choconstriction, delay
	nia in type 1 diabetes, and impair exercise performance. <sup>4</sup>		
Nadolol	Angina/HTN: Start with 40 mg once daily (Canada:	• US: Reduce dosing interval	<b>US</b> : 20, 40 mg scored
Corgard (US),	Start with 80 mg once daily. Angina patients stable on	for CrCl ≤50 mL/min.	tabs
generics	80 mg once daily can be tried on 40 mg once daily).		~\$53 for 80 mg once
	Usual dose 40 to 80 mg once daily. Max dose:		daily
<ul> <li>Low lipophilicity</li> </ul>	160 to 240 mg once daily for angina, 240 to 320 mg once		
<ul> <li>Kidney excretion</li> </ul>	daily for HTN.		<b>Canada</b> : 40, 80, 160 mg
• Long half-life	<b>A-fib</b> : Usual dose 10 to 240 mg once daily. <sup>1</sup>		scored tabs
	Migraine prevention: Usual dose 80 mg once daily. <sup>3</sup>		~\$11 for 80 mg once
	Max 240 mg once daily. <sup>3</sup>		daily
• High lipophilicity • Extensive first-pass metabolism • Bioavailability variable; increased ~50% by high-protein food • Liver elimination	Angina: 80 to 320 mg/day, divided BID to QID. (Canada: Start with 10 to 20 mg TID to QID. Usual dose 160 mg/day. Max dose 320 to 400 mg/day).  Arrhythmias (Canada): 10 to 30 mg TID to QID  A-Fib: 10 to 40 mg TID to QID¹  Essential tremor (US): Initial 40 mg BID. Usual dose 120 mg/day. Max dose 320 mg/day.  HTN: Start with 40 mg BID. Usual dose 120 to 240 mg/day, divided BID or TID (Canada: 80 to 160 mg BID). Max dose 640 mg/day (US).  Hypertrophic subaortic stenosis: 20 to 40 mg TID to QID Post-MI (US): 40 mg TID, titrated to target dose of 180 to 240 mg/day divided BID to QID.  Migraine prevention: Start with 80 mg/day, divided. Usual dose 160 to 240 mg/day (Canada: 80 to 160 mg/day). Pheochromocytoma (with alpha-blocker): 60 mg/day, divided, for 3 days before surgery, or 30 mg/day, divided, for inoperable tumor, with alpha blockade.	<ul> <li>Post-MI: Reduces CV and total mortality (BHAT)</li> <li>Substrate of CYP2D6 (mainly), CYP1A2, CYP2C19, and P-gp</li> <li>Caution with kidney or liver impairment (Canada). Reduce starting dose in liver impairment (US).</li> <li>Consider stopping one year post-MI without another indication.<sup>7</sup></li> <li>Low concentration in breast milk.<sup>8</sup></li> <li>Dose-related vivid dreams.</li> <li>Risk of fatigue slightly higher than newer agents.<sup>5</sup></li> <li>Wide dosing range may lead to more dosage adjustments than other agents.<sup>4</sup></li> </ul>	US: 10, 20, 40, 60, 80 mg scored tabs; 4.28 mg/mL, 20 mg/5 mL, 40 mg/5 mL oral solution; IV solution ~\$38 for 40 mg TID (tabs)  Canada: 10, 20, 40, 80 mg scored tabs; 4.28 mg/mL; IV solution ~\$13 for 40 mg TID (tabs)

Agent	Dosing (Adults) <sup>b</sup>	Comments*	Availability
Pharmacokinetics**			Cost of 30-day supply <sup>a</sup>
Noncardioselective Age	nts, continued		
Propranolol, extended-release Inderal LA, generics (US); InnoPran XL (US); Lupin- propranolol LA (Canada)  • Once-daily sustained-release formulation • High lipophilicity • Extensive first-pass metabolism • Liver elimination	Inderal LA (US) Angina: Start with 80 mg once daily. Usual dose 160 mg once daily. Max dose 320 mg once daily. HTN: Start with 80 mg once daily. Usual dose 120 to 160 mg once daily. Max dose 640 mg once daily. Hypertrophic subaortic stenosis: Usual dose 80 to 160 mg once daily. Migraine prevention: Start with 80 mg once daily. Usual dose 160 to 240 mg once daily.  InnoPran XL (US) HTN: Start with 80 mg HS. May increase up to 120 mg HS.  Lupin-propranolol LA (Canada) HTN or angina: establish dose with immediate-release tablets, then switch to the equivalent daily dose of Lupin-propranolol LA.	<ul> <li>Substrate of CYP2D6 (mainly), CYP1A2, CYP2C19, and P-gp</li> <li>Caution with kidney or liver impairment.</li> <li>Risk of fatigue slightly higher than newer agents<sup>5</sup></li> </ul>	US: Inderal LA: 60, 80, 120, 160 mg extended-release caps InnoPran XL: 80, 120 mg extended- release caps ~\$43 for 120 mg once daily (generic for Inderal LA)  Canada: Lupin-propranolol LA: 60, 80, 120, 160 mg extended-release caps ~\$50 for 120 mg once daily
Sotalol	See our FAQ, Atrial Fibrillation: Focus on Pharmacother	apy.	1
• Low to moderate lipophilicity <sup>4</sup> • Moderate first-pass metabolism • Liver elimination <sup>4</sup>	HTN: Start with 10 mg BID (Canada: 5 to 10 mg BID with other antihypertensives). Usual dose 20 to 40 mg/day. Max dose 60 mg/day, divided BID.  Angina (Canada): Start with 5 mg BID to TID. Usual dose 35 to 45 mg/day, divided.  Migraine prevention: Start with 10 mg BID. (US: Can give 20 mg once daily as maintenance dose). Max dose 30 mg/day, divided. Some patients may only need 10 mg once daily.  Post-MI: 10 mg BID (Canada: Start with 5 mg BID)	<ul> <li>Post-MI: Reduces cardiovascular and total mortality, including sudden death, and reduces risk of nonfatal reinfarction (NMS)<sup>17</sup></li> <li>Substrate of CYP2D6</li> <li>Caution with kidney or liver impairment</li> <li>Consider stopping one year post-MI without another indication.<sup>7</sup></li> <li>Risk of fatigue slightly higher than newer agents<sup>5</sup></li> </ul>	US: 5, 10, 20 mg scored tabs ~\$140 for 20 mg BID  Canada: 5, 10, 20 mg scored tabs ~\$48 for 20 mg BID

Agent Pharmacokinetics**	Dosing (Adults) <sup>b</sup>	Comments*	Availability Cost of 30-day supply <sup>a</sup>	
Cardioselective Agents (beta-1 antagonist activity only) Can use in asthma or COPD. 20,21 May reduce COPD exacerbations or improve survivial. 20				
Atenolol Tenormin, generics  • Low lipophilicity <sup>4</sup> • Bioavailability about 50% • Kidney elimination	Angina: Start with 50 mg once daily. May increase to 100 mg once daily (Canada: or 50 mg BID). Max dose 200 mg/day.  A-fib: Usual maintenance dose 25 to 100 mg once daily.  HTN: Start with 50 mg once daily. May be increased to 100 mg once daily.  Migraine prevention: Usual dose 100 mg once daily.  Post-MI (US): 50 mg BID or 100 mg once daily (Although it is approved post-MI after IV beta-blockade, due to increased risk of cardiogenic shock in COMMIT/CCS-2 trial of IV metoprolol, IV beta-blockade is used selectively. 16)	<ul> <li>Reduce dose for CrCl ≤35 mL/min/1.73 m².</li> <li>Consider stopping one year post-MI without another indication.<sup>7</sup></li> <li>HTN: may not reduce CV risk.<sup>12</sup> Atenolol not better than placebo for CV outcomes.<sup>18</sup> Losartan had fewer strokes and greater regression of LVH than atenolol in LIFE study.<sup>9</sup> Amlodipine +/- perindopril had lower mortality and stroke than atenolol +/- bendroflumethiazide in ASCOT.<sup>10</sup></li> </ul>	US: 25, 50 (scored), 100 mg tabs <\$5 for 50 mg BID  Canada: 25 (unscored), 50, 100 mg <\$10 for 50 mg BID	
• Moderate lipophilicity <sup>4</sup> • Low first-pass metabolism • Bioavailability ~90% • Mostly liver elimination	HTN: Start 10 mg once daily. May increase to 20 mg once daily. Max dose 40 mg once daily.	• Reduce dose in severe kidney impairment. Start with 5 mg once daily. May increase to 20 mg once daily.	US: 10 (scored), 20 mg tabs ~\$35 for 20 mg once daily	
<ul> <li>Bisoprolol</li> <li>Low lipophilicity<sup>4</sup></li> <li>Low first-pass metabolism</li> <li>Bioavailability 80%</li> <li>50% kidney elimination</li> </ul>	Angina: 5 to 20 mg once daily. <sup>3</sup> A-Fib: Usual maintenance dose 2.5 to 10 mg once daily. <sup>1</sup> HF: Usual starting dose is 1.25 mg once daily, titrated to a target dose of 10 mg once daily. <sup>2,6</sup> HTN: Start 2.5 to 5 mg once daily (Canada: 5 mg once daily). Max 20 mg once daily.	<ul> <li>US: reduce starting dose to 2.5 mg once daily for CrCl &lt;40 mL/min, liver disease, or bronchospastic disease.</li> <li>HF: Reduces mortality (CIBIS-II).<sup>13</sup></li> </ul>	US: 5(scored), 10 mg tabs ~\$18 for 10 mg once daily  Canada: 5 (scored), 10 mg tabs  <\$5 for 10 mg once daily	

Agent	Dosing	Comments*	Availability
Pharmacokinetics** Cardioselective Agents,	(Adults) <sup>b</sup>		Cost of 30-day supply <sup>a</sup>
Metoprolol tartrate, immediate-release Lopressor (US), generics  • Moderate lipophilicity <sup>4</sup> • Bioavailability about 40%-50% due to first-pass metabolism <sup>4</sup> • Liver elimination	A-fib: Usual maintenance dose is 25-100 mg BID.¹ Angina: Start with 50 mg BID. Usual dose 200 mg/day (Canada). Max dose 400 mg/day (Canada: divided BID to TID). HTN: Start with 100 mg once daily or divided (Canada: 50 mg BID. Usual dose 100 to 200 mg/day). Max dose 450 mg/day (Canada: 200 mg BID). If effect does not last 24 h with once-daily dosing, divide dose.  Migraine prevention: Start with 25 mg BID. Max dose 200 mg/day, divided.³  Post-MI: Start with 50 mg every six hours (25 mg if not tolerated), for 48 hours. Thereafter, dose is 100 mg BID. (Approved post-MI after IV beta-blockade. Due to increased risk of cardiogenic shock in COMMIT/CCS-2 trial of IV metoprolol, IV beta-blockade is used selectively.¹6)	<ul> <li>Substrate of CYP2D6</li> <li>Reduce starting dose in liver impairment (US). Canada: Reduce starting dose and maintenance dose in severe liver impairment, and use caution in severe kidney impairment.</li> <li>Consider stopping one year post-MI without another indication.<sup>7</sup></li> <li>HF: Greater reduction in mortality with carvedilol than with immediate-release metoprolol tartrate in COMET.<sup>14</sup></li> <li>Post-MI: Reduces total mortality, sudden death, and reinfarction (Goteborg).<sup>15</sup></li> </ul>	US:50, 100 mg scored tabs; IV formulation <\$5 for 100 mg BID (generic)  Canada: 25, 50, 100 mg scored tablets; IV formulation <\$10 for 100 mg BID
Metoprolol succinate, extended-release (US) Toprol-XL, generics  • Moderate lipophilicity <sup>4</sup> • Sustained-release formulation that maintains therapeutic plasma concentrations for 24 hours  • Bioavailability about 40%-50% due to first-pass metabolism <sup>4</sup> • Liver elimination	Angina: Start with 100 mg once daily. Max dose 400 mg/day.  A-fib: Usual maintenance dose is 50-400 mg once daily.  HF: Start with 12.5 to 25 mg once daily. Target dose 200 mg/day.  HTN: Start with 25-100 mg once daily. Max dose 400 mg/day.	<ul> <li>Substrate of CYP 2D6</li> <li>Reduce starting dose in liver impairment</li> <li>HF: Reduces mortality and cardiovascular hospitalization (MERIT-HF).</li> </ul>	US: 25, 50, 100, 200 mg scored extended-release tabs ~\$22 for 200 mg once daily

Agent	Dosing	Comments*	Availability
Pharmacokinetics**	(Adults) <sup>b</sup>		Cost of 30-day supply <sup>a</sup>
Cardioselective Agents,			
Nebivolol Bystolic, generics (US)  • Low lipophilicity <sup>4</sup> • Bioavailability not determined	HTN: Start with 5 mg once daily. Max dose 40 mg/day (Canada: max dose 20 mg once daily).  HF: Start with 1.25 mg once daily, titrated to target dose of 10 mg once daily.  Migraine prevention: Dose is 5 mg once daily.  Migraine prevention: Dose is 5 mg once daily.	<ul> <li>Substrate of CYP2D6</li> <li>Start with 2.5 mg once daily for CrCl &lt;30 mL/min or moderate hepatic impairment.         <p>Contraindicated in severe liver impairment.     </p></li> <li>Causes peripheral vasodilation by increasing nitric oxide production<sup>4</sup></li> <li>HF: Reduced composite endpoint of mortality and cardiovascular hospitalizations in the elderly (SENIORS)<sup>11</sup></li> </ul>	US: 2.5, 5, 10, 20 mg tabs ~\$24 for 10 mg once daily  Canada: 2.5, 5, 10, 20 mg tabs ~\$48 for 10 mg once daily
Agents with alpha-1 ant			
	ral vasodilation. <sup>4</sup> Alpha-1 antagonist activity poses risk of floppy		1
Carvedilol,	<b>A-Fib:</b> Usual maintenance dose 3.125-25 mg BID. <sup>1</sup>	• Substrate of CYP2D6 and	US: 3.125, 6.25,12.5,
immediate-release	HF: Start with 3.125 mg BID, titrated to target dose of	CYP2C9	25 mg tabs
Coreg (US), generics	25 mg BID (may use 50 mg BID for patients >85 kg [US	• Contraindicated in severe liver	<\$10 for 25 mg BID
NI 4 1: 1 4: 4	labeling: with mild-moderate HF]). <sup>2,6</sup> Reduce dose if HR <55.	impairment. Canada: caution in	Canada: 2 125 6 25 25
• Not cardioselective <sup>4</sup>		milder liver impairment.  • Consider stopping one year	Canada: 3.125, 6.25, 25
• Moderate	HTN (US): Start with 6.25 mg BID. Max dose 25 mg BID.	post-MI without another	mg tabs ~\$13 for 25 mg BID
lipophilicity <sup>4</sup>	LVD after MI (US): Start with	indication. <sup>7</sup>	~\$13 101 23 IIIg BID
• Bioavailability 25%	6.25 mg BID, titrated to target dose of 25 mg BID. (Can	HF: Reduces mortality in	
to 35% due to first-	start with 3.125 mg BID if appropriate [e.g., low blood	NYHA stage 2-4; has the	
pass metabolism	pressure, heart rate, fluid retention].)	strongest evidence for benefit in	
• Liver elimination	pressure, near rate, nata retention].)	severe failure (COPERNICUS).	
	Take with food to slow absorption, thereby reducing risk	Greater reduction in mortality	
	of orthostatic hypotension.	than with immediate-release	
	<b>71</b>	metoprolol tartrate in COMET. <sup>14</sup>	
		• Post-MI with LVD: Reduces	
		mortality and reinfarction in	
		patients taking an ACEI or ARB	
		(CAPRICORN)	

Agent	Dosing	Comments*	Availability
Pharmacokinetics**	(Adults) <sup>b</sup>		Cost of 30-day supply <sup>a</sup>
	agonist activity, continued		
Carvedilol phosphate, extended-release (US) Coreg CR, generics  • Sustained-release formulation that maintains therapeutic plasma concentrations for 24 hours  • Not cardioselective <sup>4</sup> • Moderate lipophilicity <sup>4</sup> • Bioavailability 25% to 35% due to first-pass metabolism  • Liver elimination	<ul> <li>HF: Start with 10 mg once daily, titrated to a target dose of 80 mg once daily (equal to 25 mg BID immediate-release product). Reduce dose if HR &lt;55.</li> <li>HTN: Start 20 mg once daily. Max dose 80 mg once daily.</li> <li>LVD after MI: Start with 20 mg once daily, titrated to target dose of 80 mg once daily. (Can start with 10 mg once daily if appropriate [e.g., low blood pressure, heart rate, fluid retention].)</li> <li>Take with food.</li> </ul>	<ul> <li>Contraindicated in severe hepatic impairment</li> <li>When switching from carvedilol immediate-release 12.5 mg BID or 25 mg BID, consider a starting dose of <i>Coreg CR</i> 20 mg or 40 mg once daily, respectively, especially in patients at increased risk of hypotension, dizziness, or syncope (e.g., the elderly)</li> <li>Consider stopping one year post-MI without another indication.<sup>7</sup></li> </ul>	US: 10, 20, 40, 80 mg extended-release caps ~\$240 for 40 mg QD
Labetalol Trandate (Canada), generics  • Not cardioselective <sup>4</sup> • Low lipophilicity <sup>4</sup> • Bioavailability 20% to 40% due to first- pass metabolism <sup>4</sup> • Bioavailability may be increased by food <sup>4</sup> • Liver elimination <sup>4</sup>	HTN: Start with 100 mg BID. Usual dose 200-400 mg BID. Max dose 2,400 mg/day (Canada: max dose 600 mg BID). If nausea and/or dizziness occur, consider TID dosing.  • Preferably taken after food (Canada).	<ul> <li>Wide dosing range may lead to more dosage adjustments than other agents</li> <li>Caution with liver impairment</li> <li>Rare liver injury</li> <li>A preferred antihypertensive in pregnancy. 19 Low concentrations in breast milk. 8</li> </ul>	US: 100, 200, 300 mg scored tabs; IV formulation ~\$25 for 200 mg BID  Canada: 100, 200 mg scores tabs; IV formulation ~\$23 for 200 mg BID

Agent	Dosing	Comments*	Availability
Pharmacokinetics**	(Adults) <sup>b</sup>		Cost of 30-day supply <sup>a</sup>
	npathomimetic activity (ISA)		
	ate and negative inotropic activity may be less than with other beta		I
• Cardioselective • Mild ISA • Low lipophilicity <sup>4</sup> • Bioavailability 40% due to first-pass metabolism to active metabolite which is excreted in urine	HTN: Start with 400 mg/day once daily or divided BID (Canada: Start with 100 mg BID). Usual dose 400-800 mg/day. Some patients may only need 200 mg/day. Max dose 600 mg BID (Canada: max dose 400 mg BID).  Angina (Canada): Start with 200 mg BID. Usual dose 200 to 600 mg/day, divided BID. Max dose 300 mg BID. Patients stable on 400 mg/day can be tried on 100 mg BID.  Ventricular arrhythmias (US): Start with 200 mg BID. Max dose 300-600 mg BID.	<ul> <li>Agents without ISA are preferred for HTN in patients with angina. 16</li> <li>Reduce dose for CrCl &lt;50 mL/min</li> </ul>	US: 200, 400 mg caps ~\$36 for 200 mg BID Canada: 100, 200, 400 mg caps <\$10 for 200 mg BID
Pindolol Visken (Canada), generics  • Not cardioselective • Low lipophilicity <sup>4</sup>	HTN: Start with 5 mg BID. Max dose 60 mg/day (Canada: max dose 45 mg/day. Doses >30 mg should be divided TID.)  Angina (Canada): Start with 5 mg TID. Usual dose 15 to 40 mg/day  • Take with food (Canada).	<ul> <li>Agents without ISA are preferred for HTN in patients with angina<sup>16</sup></li> <li>Caution with liver or kidney impairment. Dose reduction is not usually needed in mild to moderate impairment.</li> </ul>	US: 5, 10 mg scored tabs ~\$74 for 10 mg BID Canada: 5, 10, 15 mg score tabs ~\$41 for 10 mg BID

**Abbreviations**: ACEI = angiotensin converting-enzyme inhibitor; ACS = acute coronary syndrome; ARB = angiotensin receptor blocker; A-fib = atrial fibrillation; BID = twice daily; CrCl = creatinine clearance; CV = cardiovascular; CYP = cytochrome P450; HF = heart failure; HR = heart rate; HS = at bedtime; HTN = hypertension; ISA = intrinsic sympathomimetic activity; LVD = left ventricular dysfunction; LVH = left ventricular hypertrophy; MI = myocardial infarction; NYHA = New York Heart Association; P-gp = P-glycoprotein; QID = four times daily; TID = three times daily.

- a. Cost is wholesale acquisition cost (WAC) for generic. Medication pricing by Elsevier, accessed November 2023.
- b. Consider reducing starting doses in elderly patients.<sup>11</sup>
- c. For more information on beta-blockers for migraine prophylaxis, see our chart, Migraine Prophylaxis.
- d. US product information used for the above chart (unless otherwise noted): Corgard (August 2021), propranolol immediate-release (Mylan,

March 2022), *Inderal LA* (August 2023), *InnoPran XL* (May 2023), timolol (Athem, May 2021), *Tenormin* (June 2021), betaxolol (KVK Tech, April 2020), bisoprolol (Solco, June 2023), *Lopressor* (July 2023), *Toprol-XL* (April 2023), *Bystolic* (June 2023), *Coreg* (May 2022), *Coreg CR* (November 2020), labetalol (Cadila, September 2022), acebutolol (ANI, August 2022), pindolol (ANI, August 2022).

Canadian product monographs used for the above chart (unless otherwise noted): nadolol (Apotex, August 2020), propranolol immediate-release (Teva, September 2011), propranolol LA (Lupin, January 2022), timolol (AA Pharma, February 2018), *Tenormin* (May 2023), bisoprolol (Sivem, October 2022), metoprolol tartrate (Pro Doc Ltée, August 2023), *Bystolic* (November 2022), carvedilol (Sivem, August 2022), acebutolol (Apotex, October 2018), *Visken* (May 2021).

\* Clinical Trial Acronyms: ASCOT = Anglo-Scandinavian Cardiac Outcomes Trial, BHAT = Beta-Blocker Heart Attack Trial, CAPRICORN = Carvedilol Post Infarction Survival Control in Left Ventricular Dysfunction, CIBIS-II = Cardiac Insufficiency Bisoprolol Study II, COMET = Carvedilol or Metoprolol European Trial, COMMIT/CCS-2 = Clopidogrel and Metoprolol in Myocardial Infarction Trial – Second Chinese Cardiac Study, COPERNICUS = Carvedilol Prospective Randomized Cumulative Survival, LIFE = Losartan Intervention For End point Reduction in Hypertension, MERIT-HF = Metoprolol CR/XL Randomized Intervention Trial in Congestive Heart Failure, NMS = Norwegian Multicenter Study, SENIORS = Study of the Effects of Nebivolol Intervention on Outcomes and Rehospitalisation in Seniors with Heart Failure.

Users of this resource are cautioned to use their own professional judgment and consult any other necessary or appropriate sources prior to making clinical judgments based on the content of this document. Our editors have researched the information with input from experts, government agencies, and national organizations. Information and internet links in this article were current as of the date of publication.

## Levels of Evidence

In accordance with our goal of providing Evidence-Based information, we are citing the **LEVEL OF EVIDENCE** for the clinical recommendations we publish.

Level	Definition	Study Quality	
A	Good-quality	1.	High-quality
	patient-oriented		randomized
	evidence.*		controlled trial (RCT)
		2.	Systematic review
			(SR)/Meta-analysis
			of RCTs with
			consistent findings
		3.	All-or-none study
В	Inconsistent or	1.	Lower-quality RCT
	limited-quality	2.	SR/Meta-analysis
	patient-oriented		with low-quality
	evidence.*		clinical trials or of
			studies with
			inconsistent findings
		3.	Cohort study
		4.	Case control study
C	Consensus; usual practice; expert opinion;		
	disease-oriented evidence (e.g., physiologic or		
	surrogate end points); case series for studies of		
	diagnosis, treatment, prevention, or screening.		

<sup>\*</sup>Outcomes that matter to patients (e.g., morbidity, mortality, symptom improvement, quality of life).

[Adapted from Ebell MH, Siwek J, Weiss BD, et al. Strength of recommendation taxonomy (SORT): a patient-centered approach to grading evidence in the medical literature. Am Fam Physician. 2004 Feb 1;69(3):548-56.

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