

Type 2 diabetes in non-pregnant adult
Initial diagnosis, drug naive

Lifestyle modifications

A1C < 7.5

A1C > 7.5 – 9.0

A1C > 9.0

Monotherapy
METFORMIN

Formulary alternative if contraindicated:
SU (glipizide)
Glinide (repaglitinide)
TZD (pioglitazone)
AGI (acarbose)

If above agents cannot be used due to drug/patient characteristics:
DPP-4i (alogliptin)
SGLT-2i (ertugliflozin)

METFORMIN
May consider 2 agents based on target A1C
+
Formulary agents:
SU (glipizide)
Glinide (repaglitinide)
TZD (pioglitazone)
Basal insulin (insulin glargine)
AGI (acarbose)

May consider dual therapy with
METFORMIN
+
Basal insulin
(insulin glargine)

A1C at target

A1C at target on two agents

A1C at target

Maintain therapy

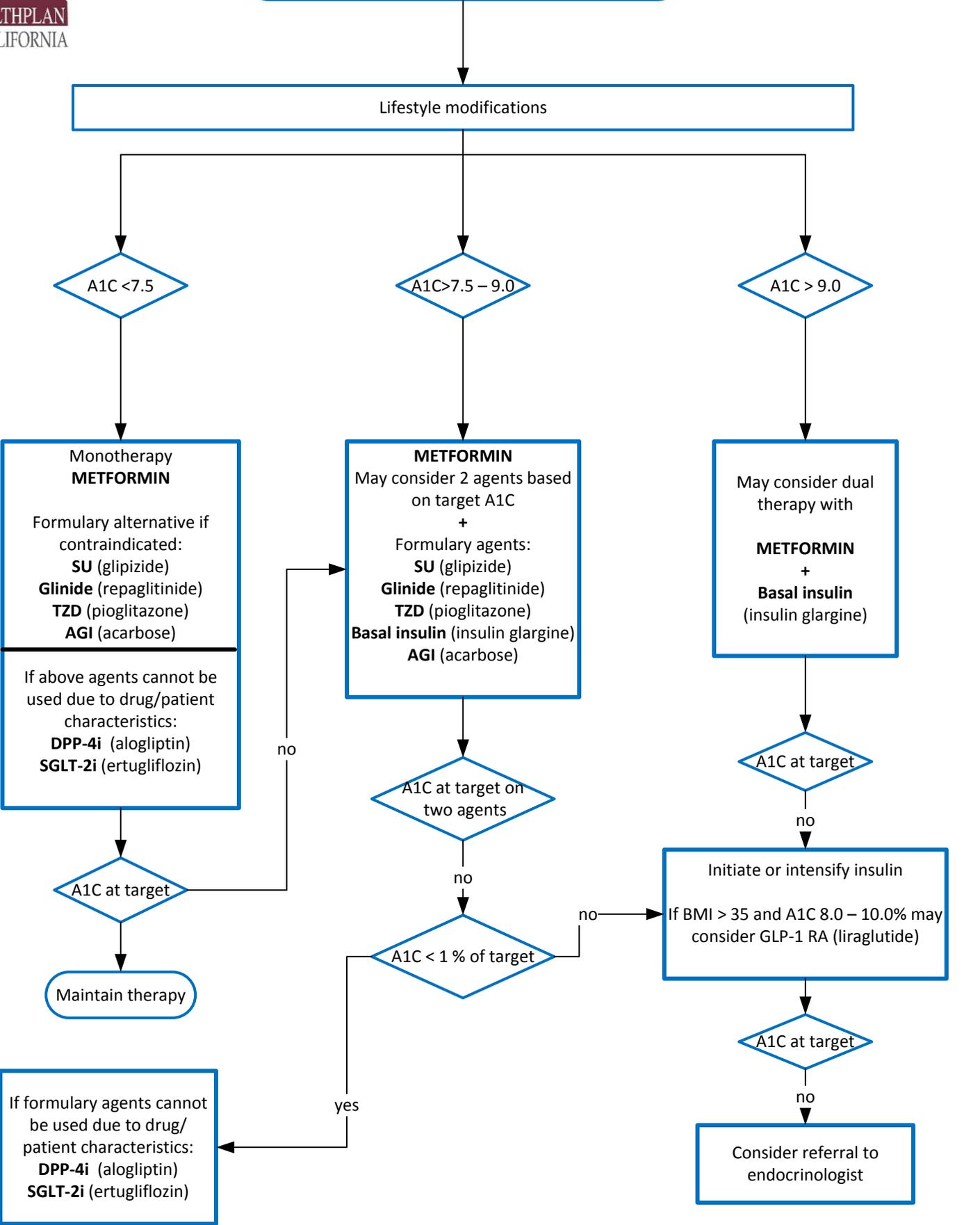
A1C < 1 % of target

Initiate or intensify insulin
If BMI > 35 and A1C 8.0 – 10.0% may consider GLP-1 RA (liraglutide)

A1C at target

Consider referral to endocrinologist

If formulary agents cannot be used due to drug/patient characteristics:
DPP-4i (alogliptin)
SGLT-2i (ertugliflozin)



DRUG CLASS	MEDICATION						
Sulfonylureas (SU)	Glipizide	Glimepiride	Glyburide				
Meglitinides (GLN)	Repaglinide	Nateglinide					
Thiazolidinediones (TZD)	Pioglitazone						
A-glucosidase Inhibitors (AGI)	Acarbose	Miglitol					
SGLT-2 Inhibitors (SGLT-2i) \$\$\$\$	Ertugliflozin <i>Formulary step</i>	Canagliflozin <i>NF</i>	Empagliflozin <i>NF</i>	Dapagliflozin <i>NF</i>			
DPP-4 Inhibitors (DPP-4i) \$\$\$\$	Alogliptin <i>Formulary step</i>	Sitagliptin <i>NF</i>	Linagliptin <i>NF</i>	Saxagliptin <i>NF</i>			
GLP-1 receptor agonists (GLP-1 RA) \$\$\$\$\$\$	Liraglutide <i>Formulary step</i>	Dulaglutide <i>NF</i>	Semaglutide <i>NF</i>	Exenatide ER <i>NF</i>	Exenatide <i>NF</i>	Lixisenatide <i>NF</i>	Albiglutide <i>NF</i>
<p>* Relative cost: each \$ = \$100 * NF: Non-formulary</p>							

- Initiate metformin 500 mg ½ to 1 tablet once a day with meal and slowly titrate up q 5-7 days. If gastrointestinal side effects occur, consider ER formulation (500 mg and 750 mg on formulary).
- Alternatives to SU should be considered if BMI > 35 or eGFR < 30 ml/min.
- Can consider DPP-4i if expected change in A1C is <1% in order to reach patient specific target.
- Can consider GLP-1 RA after maximizing basal insulin, in member with BMI > 35, if A1C 8.0 – 10.0%. If A1C >10.0% consider intensifying insulin.
- Per AACE guidelines, patients taking 2 oral antihyperglycemic agents with an A1C >8.0% and/or long-standing diabetes are unlikely to reach their target A1C with a third oral antihyperglycemic agent. ²
- Per AACE guidelines, prandial insulin should be consider when the total daily dose of basal insulin is greater than 0.5 U/kg. Beyond this dose of basal insulin, the risk of hypoglycemia increases without significant improvement in A1C. ²
- Per ADA guidelines, SU, DPP-4i, and GLP-1 RA are typically stopped once more complex insulin regimens beyond basal insulin are used. ²
- Per ADA guidelines, for patients with suboptimal blood glucose control, especially those requiring increasing insulin doses, adjunctive use of TZD (usually pioglitazone) or SGLT-2i may improve control and reduce the amount of insulin needed. ²

References:

1. American Diabetes Association. Pharmacologic Approaches to Glycemic Treatment. Sec. 8. Standards of Medical Care in Diabetes-2017. *Diabetes Care* 2017;40(Suppl.1):S64-S74
2. Consensus Statement by the American Association of Clinical Endocrinologists and American College of Endocrinology on the Comprehensive Type 2 diabetes Management Algorithm-2018 Executive Summary. *Endocrine Practice* 2018;24 (1):91-120
3. Handelsman Y, Bloomgarden ZT, Grunberger G, et al. American Association of Clinical Endocrinologists and American College of Endocrinology 2015 Clinical Practice Guidelines for Developing a Diabetes Mellitus Comprehensive Care Plan. *Endocrine Practice* 2015;21Suppl 1:1-87