ORAL ANTIRETROVIRAL/HCV DAA ADMINISTRATION: INFORMATION ON CRUSHING AND LIQUID DRUG FORMULATIONS

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
Combination Products:					
Atripla® (efavirenz/ emtricitabine/ tenofovir DF)	no	Consider use of Truvada® tabs and efavirenz caps as alternate formulations (see separate entries)		Atripla® tablet was crushed, dissolved in 5 mL of water and diluted to 20 mL with Ora-Sweet oral vehicle. The solution was prepared within 24 hours of administration to ensure drug stability in solution. Bioequivalence of Atripla® tablet and compounded oral liquid formulation (above) in HIV-negative volunteers was **not** demonstrated. The 90% CI for FTC Cmax and AUC fell within the range of 0.8-1.25 thus, bioequivalence was met, but the 90% CI for efavirenz Cmax fell below the range of bioequivalence while efavirenz AUC∞ fell slightly above the range and tenofovir Cmax and AUC∞ fell above the range. Tenofovir Cmax and AUC∞ were approximately 40% and 20% higher, respectively with the liquid formulation. The clinical implications of these data are unknown, however the authors state that crushed Atripla® may be a viable option in certain patients and risks vs. benefits should be carefully	See information on crushing Atripla® in the Case Reports section. Although Truvada® tablets may be split, splitting Atripla® tablets has not been studied. There are no studies evaluating the pharmacokinetics of a split tablet vs. a whole tablet. Efavirenz is not water soluble.

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				considered. (King et al. JAIDS 2011;56(5):e131-2.)	
Biktarvy® (bictegravir/ emtricitabine/ tenofovir alafenamide)	no				Note that Biktarvy tablets should be administered whole. Crushing BIC/FTC/TAF tablets into a liquid medium has not been studied and is not recommended. While TAF is soluble in water, it has a bitter and burnt aromatic taste flavor profile. While FTC is soluble in water, BIC is practically insoluble (solubility of 0.1 mg/mL in water at 20 °C). Currently, there are no studies evaluating the pharmacokinetics (e.g., oral bioavailability) of a crushed BIC/FTC/TAF tablet dispersed into a liquid medium (e.g., milk, water, juice) compared to a whole tablet. (Data on File, Gilead US, May 2018) Similarly, splitting BIC/FTC/TAF tablets has not been studied and it is not recommended. Currently, there is no study evaluating the pharmacokinetics of a split tablet versus a whole tablet. (Data on File, Gilead US, May 2018).
Combivir® (lamivudine/ zidovudine)	yes (individual components)	Use lamivudine & zidovudine liquid products			No data, but likely OK to crush tablets (film-coated); crush immediately before ingestion.

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					May have bitter taste. Tablets can be crushed and added to small amount of liquid or semi-solid food; consume immediately. (Duggan et al. Am J Health-Syst Pharm. 2015; 72:1555-65.)
Complera® (tenofovir DF/ emtricitabine/ rilpivirine)	no				Splitting or crushing Complera® tablets into a liquid medium has not been studied and is not recommended. Rilpivirine hydrochloride is insoluble in water over a wide pH range. (Email communication, Gilead July 2012).
Descovy® (tenofovir AF emtricitabine)	no			Case report of successful administration of crushed tenofovir AF/emtricitabine and dolutegravir tablets, mixed with water and injection via catheter syringe into percutaneous endoscopic gastrotomy (PEG) tube, immediately followed by a can of enteral nutrition (Ensure). (Fulco & Higginson, AJHP 2018;75:594-5.)	Crushing or splitting Descovy® tablets has not been studied and is not recommended. TAF is soluble in water. However, it has a bitter and burnt aromatic flavor profile. Emtricitabine is soluble in water. (Email communication, Gilead January 2017).
Delstrigo® (doravirine/ lamivudine/ tenofovir DF)	yes	Only lamivudine liquid available			Crushing or splitting Delstrigo® has not been studied and is not recommended. The product should be swallowed whole. The tablets are film-coated (e-mail communication Merck Canada Inc, September 2018).

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Dovato® (dolutegravir/ lamivudine)	yes	Only lamivudine liquid available			The manufacturer recommends swallowing the tablet whole. The tablets are film-coated. For patients who cannot swallow the tablet who, in theory, the tablet may be split in halves or crushed and added to a small amount of semi-solid food or liquid and the full contents consumed immediately (e-mail communication, ViiV Healthcare, Med US, March 2019).
Evotaz® (atazanavir/ cobicistat)	no				The manufacturer recommends swallowing the tablet whole; do not crush or chew tablets (Evotaz Product Monograph, Sept 2015).
Genvoya® (elvitegravir/ cobicistat/ tenofovir AF/ emtricitabine)	no			A cancer patient received Genvoya® via a percutaneous gastrostomy tube. The tablet was crushed (pulverized), mixed with 30 mL of tap water, and administered via a syringe followed by 2 x 250 mL cans of enteral feeds (Diabetisource AC). At week 14, the viral load was < 20 copies/mL. The patient died of metastatic cancer within 15 weeks of starting crushed Genvoya®. (Kaplun et al. Am J Health-Syst Pharm 2019;76(16):1180-81.)	Splitting or crushing Genvoya® tablets into a liquid medium has not been studied and is not recommended. TAF is soluble in water but has a bitter and burnt aromatic taste. Emtricitabine is soluble in water, but cobicistat and elvitegravir are practically insoluble (email communication, Gilead, March 2016). See Stribild® for more information.

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				See case reports for Stribild®.	
Juluca® (dolutegravir/ rilpivirine)	no				The efficacy, safety and pharmacokinetics of crushing dolutegravir/rilpivirine have not been evaluated. The tablet should be swallowed whole and taken with a meal to ensure administration of the entire dose. The tablets are film-coated. Based on clinical judgment if the Juluca® tablet requires splitting, it should be split in half and both halves ingested immediately with a meal. If the Juluca® tablet requires crushing, it should be crushed and added to a small amount of liquid or semi-solid food and the full tablet content consumed immediately with a meal. (Data on File, ViiV Healthcare, May 2018)
Kivexa® (abacavir/ lamivudine)	yes (individual components)	Use abacavir & lamivudine liquid products			Film-coated immediate release tablet; however, no studies regarding stability of split or crushed tablets. (Email communication, GlaxoSmithKline, May 2008) Tablet may be split or crushed and added to a small amount of food or water. (European

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					Medicines Agency, EPAR summary for the public, Ziagen updated 05-2010)
Odefsey® (tenofovir AF/ emtricitabine/ rilpivirine)	no				Crushing or splitting Odefsey® tablets has not been studied and is not recommended. TAF is soluble in water. However, it has a bitter and burnt aromatic flavour profile. Rilpivirine hydrochloride is insoluble in water over a wide pH range. (Email communication, Gilead January 2017).
Prezcobix® (darunavir/ cobicistat)	no	See darunavir and ritonavir for other liquid options (substitution of cobicistat with ritonavir may be required)			Splitting Prezcobix® film-coated tablets has not been studied. Tablets should be swallowed whole without breaking or crushing to ensure administration of the entire dose. (Prezcobix® Product Monograph, 2014) Tablets are immediate-release formulation; no anticipated absorption issues if the tablets are chewed, split or crushed. (Huesgen et al. Pharmacother 2016;36(11):1145-65.)
Stribild® (elvitegravir/ cobicistat/ emtricitabine/ tenofovir)	no			Case report describing successful virological suppression with crushed Stribild® in juice (Fulco et al. AJHP 2014;71(10);784-6.)	Pharmacokinetics of crushed Stribild® tablets were studied in healthy volunteers. Whole tablets with breakfast were compared to: I. Crushed and suspended with breakfast

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Symtuza® (darunavir/ cobicistat/ emtricitabine/ tenofovir alafenamide)					II. Crushed and suspended with enteral nutrition (Nutrison®). The groups were shown to be bioequivalent for elvitegravir, tenofovir and emtricitabine. Elvitegravir Cmax failed to fall within bioequivalence range (100-120%), but this difference is unlikely to be clinically significant. Cobicistat AUC was reduced by 10% for intervention I only. (Jongbloed-de Hoon et al. JAIDS. 2017;74(5):571-574.) Symtuza® should be swallowed whole. The manufacturer does not recommend breaking or crushing Symtuza® to ensure administration of the entire dose. Film-coated tablets. The relative bioavailability of darunavir/cobicistat/emtricitabine/tenofovir alafenamide (D/C/F/TAF) single tablet regimen was compared with the tablet administered whole, split or crushed. In the split group there was an 11% decrease in TAF Cmax only (not clinically relevant). In the crushed group there was a 17% decrease in the emtricitabine Cmax and TAF Cmax and AUC were decreased by 29% and 18%, respectively (clinical relevance

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Triumeq® (abacavir/ lamivudine/ dolutegravir (DTG))	no	Use abacavir & lamivudine liquid products. Dolutegravir tablets may be crushed (see dolutegravir). See Crushing &Splitting section also.		Case report of successful administration of crushed dolutegravir/abacavir/lamivudine tablet with administration via NG tube; adequate antiretroviral concentrations confirmed via TDM and patient achieved rapid viral suppression. (Chrdle et al. Int J STD AIDS 2019;30(1):94-8.)	not assessed, but impact expected to be minimal based on wide therapeutic window for TAF). (Brown K et al. Clin Pharmacol Drug Dev 2019;8(4):541-48.) Triumeq® is film-coated, non-scored, and non-sustained released formulation. Although not studied, splitting or crushing tablets is not expected to affect the dissolution or absorption. Tablets may be crushed and added to a small amount of semi-solid food or liquid, and consumed immediately. (Data on File, ViiV Healthcare, Oct 2014) **Nasogastric or gastric feeding tubes:** No clinical or pharmacokinetic studies done to evaluate. The administration of crushed Triumeq® tablets should not have an effect on the absorption of the components of Triumeq®. The absorption of Triumeq® is thought to occur in the proximal small intesting (duodenum/jejunum). (Data on File, ViiV Healthcare, March 2017) Pharmacokinetics of crushed Triumeq® tablets were studied in healthy volunteers. Whole

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					tablets in fasting state were compared to: I. Crushed and suspended in fasting state II. Crushed and suspended with enteral nutrition taken PO (Nutrison®-250 mL contains elemental calcium 200 mg, iron 4 mg, protein 10 g, carbohydrates 30.8 g, fat 9.8 g). Intervention I showed 26% and 30% increase in DTG AUC and Cmax. Intervention II showed an 18 % and 21 % increase in DTG AUC and Cmax, respectively. Although bio-equivalence was not demonstrated, the increase in DTG exposure was not considered to be clinically relevant. However, caution is warranted if crushed DTG is given once daily or BID with food, as DTG exposure will likely be higher. (Roskam-Kwint et al. JAC 2018;73:2430-4.)
Trizivir® (abacavir/ lamivudine/ zidovudine)	yes (individual components)	Use abacavir, lamivudine & zidovudine liquid products.			Film coated immediate release tablet however no studies regarding stability of split or crushed tablets.
Truvada® (tenofovir/ emtricitabine)	yes (individual components-US only)	See tenofovir & emtricitabine.		The absorption of raltegravir, etravirine, emtricitabine, and tenofovir was not compromised when the drugs were crushed,	May split tablets. May crush and stir into water, grape juice or orange juice. The stability of the mixture is unknown. (Email

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				dissolved in 60 mL warm water, and administered by gastrostomy tube to a 52 year old HIV-positive male with ulcerative esophagitis. (Sandkovsky et al. Pharmacother 2012;32(2):142-7.) Case report of complex HIV patient with MAC with intractable nausea/vomiting requiring ARVs (tenofovir DF 300 mg/emtricitabine 200 mg as Truvada® and dolutegravir 50 mg daily) via jejunostomy (J)-tube. ARVs were crushed, mixed with 3-5 mL or water, administered, and flushed with 10 mL of water. Concentrations of oral and J-tube administration of ARVs were assessed. DTG and TDF exposures were similar between J-tube and oral administration. FTC AUC was 38% lower for J-tube vs. oral. Compared to a reference population, overall AUC was lower for both routes- DTG 75-76% lower and TDF 55-61% lower. However, FTC via J-tube AUC was similar to the reference population and 71% higher when given orally. Reduced drug absorption was the primary cause for decreased drug exposure. TDM	communication, Gilead, July 2012). Case where patient chewed TDF/FTC tablets (for PrEP) due to swallowing difficulties. Monthly urine samples indicated tenofovir concentrations >1000 ng/mL, indicative of protection from HIV acquisition. (Lalley-Chareczko et al. Antivir Ther 2017;22:639-41.)

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				is recommended to assess drug concentrations in patients with the potential for impaired absorption (Brooks et al. Pharmacother 2017;37:e82-9.) Case report of an HIV patient with difficulty swallowing pills who preferred ARV formulations that he could crush. Tenofovir DF-emtricitabine (Truvada®) 1 tab daily and dolutegravir 50 mg daily were crushed using a pill crusher, added to applesauce, and consumed immediately. The 4- week viral load decreased from 10,800 to < 20 copies/mL (Buscemi L. Am J Health-Syst Pharm 2016;73(15):1125-26.) Case report of a 22 year-old male with difficulty swallowing tenofovir DF-emtricitabine (Truvada®) for pre-exposure prophylaxis (PrEP) chewed the whole tablets. Monthly urine samples showed tenofovir concentrations > 1000 ng/mL (protective level). Plasma samples collected at weeks 24 and 48 showed tenofovir concentrations > 10 ng/mL (also protective). (Lalley-Chareczko et al. Antivir	

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				Ther 2017;22(7):639-641.) Case report of a 20 year-old male with acute-chronic inflammatory demyelinating polyneuropathy (IDP) who was successfully treated for HIV with crushed ARVs (Truvada (tenofovir DF-emtricitabine + raltegravir) administered via gastrojejunostomy tube. The viral load decreased from 100,123 copies/mL to < 20 after two weeks of ARVs. The patient was also undergoing plasmapheresis and ARVs were administered after the plasmapheresis sessions. (Lindholm et al. J AIDS Clin Res 2013;4(12):1-7.)	
INDIVIDUAL ANTIRETROVIRAL AGENTS:					
abacavir	yes	20 mg/mL oral solution; 240 mL bottle. Yellow, strawberry-banana flavoured liquid. Contains sorbitol 340 mg/mL (E-mail communication, ViiV, April 2017)	Store oral solution at room temperature.		Tablet is film-coated. Tablet can be crushed and added to a small amount of liquid or semi-solid food; consume immediately (Duggan et al. Am J Health-Syst Pharm. 2015; 72:1555-65.)
amprenavir	no-product	See fosamprenavir for			

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	discontinued	liquid formulation.			
atazanavir	yes (US only)	50 mg/1.5 g dispersible oral powder packet	**Powder:** mix with food such as applesauce or yogurt (1 TBSP minimum). Mixing with a beverage (milk, formula, water- 30 mL + additional 15 mL after to consume residual drug) can be used if infant is able to drink from a cup. For younger infants who cannot eat solid food, mix with infant formula (10 mL + additional 10 mL after to consume residual drug) and administer via oral syringe. Stable for 1 hour at room temperature once mixed in food or beverage. (Refer to Reyataz® US Product	**Capsules**: In an open label, multicentre study of atazanavir and atazanavir/ritonavir in children 91 days-21 years, the pharmacokinetics of atazanavir capsules and atazanavir orange-vanilla flavoured powder were studied. Day 7 atazanavir kinetics were compared in children of similar age receiving powder vs. capsules; the powder was found to be 40% less bioavailable at the same BSA-based dose. Therefore, suggest converting from powder to capsule by multiplying the powder dose by 0.6 and rounding up to the nearest 50 mg. (Kiser J et al. AIDS 2011;25:1489-96.)	**Capsules:** May be opened and the contents mixed with applesauce for immediate ingestion with a light meal. In-house study showed that the bioavailability of the contents of two 200-mg atazanavir capsules mixed with applesauce was 91.7% relative to atazanavir capsules taken intact. In addition, administration of the contents of two 200-mg capsules was well tolerated (Bristol Myers Squibb, Personal Communication, November 20, 2015).

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			Monograph for additional information on mixing/administration).		
darunavir	yes- compassionate access through Janssen Canada (call Janssen Medical Information at 1-800-567-3331 or submit Compassionate Use form posted on website https://www.janssenmedicalinformation.ca/login?destination=compassionate-use; updated Feb 2020) Available in US	100 mg/mL oral suspension	Store oral suspension at room temperature. Shake well before use.	In two patients, one with dysphagia and Candida esophagitis and one with a stomach tube, who received darunavir tablets crushed and dissolved and administered with ritonavir oral solution, adequate plasma darunavir levels were achieved along with good virologic response.(Scholten et al. J Int AIDS Soc 2010;13(Suppl 4):P114.) A case report describes an intubated 44 year-old man on tenofovir/emtricitabine, darunavir, and ritonavir in ICU who was given darunavir tablets via orogastric tube crushed and dissolved in 15-20mls of water. Viral load did not change significantly and adequate darunavir trough levels were achieved. (Kim et al. CJHP 2014;67(1):39-42.) An HIV-positive patient on continuous venovenous hemodiafiltration (CVVHDF) received raltegravir 400 mg	No pharmacokinetic data are available on chewing or crushing of Prezista® film-coated tablets. However, since the tablets are not formulated as an extended release formulation, no potential problem is anticipated if the tablets are chewed or crushed for administration through a nasogastric (NG) tube. It is unlikely that chewing or crushing Prezista® tablets would have a significant impact on pharmacokinetics (Data on File, Tibotec, November 2006).

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				BID, darunavir 600/100 mg BID, zidovudine 300 mg BID and 3TC 50 mg q24h in suspension via gastric port and simultaneous enteral feeding via the duodenal port of a double-lumen nasogastroduodenal tube. Pharmacokinetic sampling and analysis indicated that darunavir and raltegravir were removed by CVVHDF with approximately the same clearance as provided by a normally functioning kidney. **Absorption of both darunavir and raltegravir after suspension and application via the gastric port with continued administration of feed via the duodenal port of the double-lumen tube was good.** As such, dose adjustments are not required for patients receiving darunavir and/or raltegravir while undergoing CVVHDF and that absorption of darunavir and raltegravir is not significantly affected by postpyloric enteral feeding. (Taegtmeyer et al. AIDS 2011;25(10):1339-41.)	
delavirdine	no				Can dissolve 100 mg tablets in water to make slurry (20% ↑ bioavailability). Disperse tablets

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					in at least 90 mL of water, allow to stand for a few minutes, stir and consume.
didanosine (ddI)	no Note: Canada-product was discontinued May 2019 by manufacturer and is no longer available via SAP; no longer available in US	Note: Product is discontinued. 4 g oral powder (pediatric solution); 10 mg/mL final concentration. Take on an empty stomach. Do not give with fruit juices or acidic drinks, feeds or milk. Chew Tablets discontinued (Canada and US)	30 days	Note: Product is discontinued. Reconstitute with commercially available antacid that contains as active ingredients aluminum hydroxide (400 mg per 5 mL), magnesium hydroxide (400 mg per 5 mL), and simethicone (40 mg per 5 mL)) If above strength not available, reconstitute with similar antacid of ½ strength using these alternative instructions: Add 400 mL of antacid in two, 200 mL portions, shaking the contents after each addition of 200 mL. The admixture may be dispensed in flint-glass or plastic bottles. Shake well before using. Stable for 30 days in fridge.	
dolutegravir	no	Pediatric film-coated tablets (10, 25 mg) available via compassionate access directly through ViiV Canada Healthcare Supports (1-855-525-5300). (Updated Feb		In comparison to the commercially available tablet, dolutegravir exposures following administration of the granule formulation alone, with different types of water and with formula exceeded that of the tablet, demonstrating the dolutegravir oral granule can be given without restriction on the type of liquid, or can be administered directly to mouth (e.g., when	10, 25 and 50 mg tablets should ideally be swallowed whole. All tablet strengths may also be split into halves followed by immediate ingestion of both halves or crushed and added to a small amount of semi-solid food or liquid, all of which should be consumed immediately. (ViiV Healthcare communication, February 2017)

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		2020) 5 mg dispersible tablet under investigation (US and Canada). Pediatric granules no longer under development.		potable water is not available). (Patel et al. Antivir Ther 2014;19(3):229-33.) Case report of a critically ill patient with lymphoma requiring enteral administration of ARVs. Both abacavir and 3TC solutions were administered enterally. Crushed **dolutegravir 50 mg BID** (separated from enteral nutrition by 2 hours) and rilpivirine 25 mg daily (given with a 240-mL bolus of an enteral formula (2 kcal/mL)) were administered via orogastric tube. Crushed tablets were each mixed with 10 mL of water and flushed down the tube at separate administration times. Trough concentrations were: day 8, rilpivirine 30 ng/mL (reference range 40-120 ng/mL), and day 9, dolutegravir 820 ng/mL (reference range 830 ng/mL steady-state trough concentration for 50 mg once daily dose). Virologic suppression was maintained after ARV enteral administration (hospital day 29). Given somewhat decreased levels of these ARVs, the authors recommended consideration to increase dolutegravir to 150-200 mg total daily dose,	See Triumeq® for more information.

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				particularly in integrase-experienced patients, and rilpivirine 50 mg daily (similar to dosing with an inducer such as rifabutin). (Turley et al. JIAPAC 2017;16(2):117-119.) Case report of complex HIV patient with MAC with intractable nausea/vomiting requiring ARVs (tenofovir DF 300 mg/emtricitabine 200 mg as Truvada® and **dolutegravir 50 mg daily)** via jejunostomy (J)-tube. ARVs were crushed, mixed with 3-5 mL or water, administered, and flushed with 10 mL of water. Concentrations of oral and J-tube administration of ARVs were assessed. DTG and TDF exposures were similar between J-tube and oral administration. FTC AUC was 38% lower for J-tube vs. oral. Compared to a reference population, overall AUC was lower for both routes- DTG 75-76% lower and TDF 55-61% lower. However, FTC via J-tube AUC was similar to the reference population and 71% higher when given orally. Reduced drug absorption was the primary cause for decreased drug exposure. TDM	

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				is recommended to assess drug concentrations in patients with the potential for impaired absorption (Brooks et al. Pharmacother 2017;37:e82-9.) Case report of an HIV patient with difficulty swallowing pills who preferred ARV formulations that he could crush. Tenofovir DF-emtricitabine (Truvada®) 1 tab daily and **dolutegravir 50 mg daily** were crushed using a pill crusher, added to applesauce, and consumed immediately. The 4- week viral load decreased from 10,800 to < 20 copies/mL (Buscemi L. Am J Health-Syst Pharm 2016;73(15):1125-26.)	
doravirine	no				Crushing or splitting Pifeltro® has not been studied and is not recommended. The product should be swallowed whole. The tablets are film coated (e-mail communication Merck Canada, Inc, September 2018).
efavirenz	no Note: pediatric suspension is no longer available internationally (2014)	30 mg/mL; 180 mL bottle- not available Consider use of capsule formulation as		**Tablets**: A pediatric pharmacokinetic intensive study that utilized weight band dosing and a combination of capsules or half of a 600 mg tablet reported low overall plasma efavirenz concentrations in both	Splitting efavirenz tablets has not been well studied. With the exception of the study by (Fillekes et al. JAIDS 2011;58(4):392-8.), there are no well controlled pharmacokinetic studies evaluating a split tablet

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		described in Clinical Compounding		groups (higher doses need to be investigated). They found no significant differences across weight bands, suggesting no discernible effect of using half tablets. (Fillekes et al. JAIDS 2011;58(4):392-8.) **Capsules:** may be opened and added to 1-2 tsp of liquids or foods (e.g. applesauce, grape jelly, yogurt, reconstituted infant formula at room temperature) but may result in peppery taste. Grape jelly may mask taste. Specific instructions: (Kaul et al. AJHP 2010;67(3):217-22.) 1. Hold the capsule horizontally over a small container and twist open to avoid spillage. 2. Pull the capsule away from the body of the capsule carefully, sprinkle and mix the contents with 1-2 tsp of food or formula. 3. Administer the mixture with a spoon as soon as possible but no more than 30 minutes after mixing. 4. After administration of the efavirenz–food mixture, an additional 2 tsp of food or infant formula must be added to the container, stirred, and given to the patient. For nasogastric administration,	vs. a whole tablet. Efavirenz is not water soluble. The use of the capsule formulation is preferred when possible. (see Case Reports/Clinical Compounding)

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				may open **capsules** and mix with either 5 mL MCT oil or 15 mL Ora-Sweet/any aqueous vehicle (grind powder first to enhance dissolution). Do NOT mix with polyethylene glycol (will ↓ bioavailability). Splitting **tablets** is not recommended (Email communication, Bristol-Myers Squibb, June 1, 2011).	
elvitegravir	no			Case report describing successful virological suppression with crushed Stribild® in juice (Fulco et al. AJHP 2014 71(10);784-6.)	Crushing or splitting Genvoya® tablets has not been studied and is not recommended. While emtricitabine and TAF are soluble in water, cobicistat and elvitegravir are practically insoluble in water. See Genvoya® for more information. (Communication from Gilead Canada, March 2016). See Stribild® for more information.
emtricitabine (FTC)	no (US only)	10 mg/mL oral solution	Store oral solution refrigerated; stable for 3 mos at room temperature.		200 mg capsules may be opened and mixed with water.
etravirine	no	See Crushing and Splitting section for dispersion information.	Consume immediately.	The absorption of raltegravir, etravirine, emtricitabine, and tenofovir was not compromised when the drugs were crushed, dissolved in 60 mL warm water, and administered by gastrostomy tube to a 52 year	Patients who are unable to swallow etravirine tablets whole may disperse the tablets in a glass of water. A bioavailability study has shown that the PK of etravirine tablets when swallowed whole and when

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				old HIV-positive male with ulcerative esophagitis. (Sandkovsky et al. Pharmacother 2012;32(2):142-7.)	taken after dispersion in a glass of water are comparable. Both the 100 mg and 200 mg tablet formulations of etravirine may be dispersed in water (Kakuda et al. Int J Clin Pharmacol Ther 2013;51(9):725-37.) Place the tablet in 5 mL of cold water or at least enough liquid to cover the medication. Stir until a homogenous, white, cloudy, suspension is obtained. If desired, add more water or alternatively orange juice or milk. Once dispersed, patients should stir the dispersion well and drink it immediately. The glass should be rinsed with water, orange juice or milk several times and each rinse completely swallowed to ensure the entire dose is consumed. Avoid the use of grapefruit juice, warm liquids (> 40°C) or carbonated beverages. (Intelence® Product Monograph, 2014).
fosamprenavir	yes	50 mg/mL oral suspension, 225 mL bottle. 0.6% propylene glycol Grape bubblegum and peppermint flavour.	Store oral suspension between 2-30°C. Do not freeze. Discard the suspension 28 days after first		No information on crushing or dissolution of 700 mg tablets. Fosamprenavir calcium tablets and suspension are equivalent on a mg per mg basis.

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		In adults, oral suspension should be taken on an empty stomach (1 hr before or 2 hours after food). In pediatric patients, oral suspension should be given with food.	opening.		
indinavir	no			10 mg/mL indinavir syrup complex compounding formulation. Stable for 14 days in refrigerator, store in glass bottle. (Hugen et al. AJHP 2000; 57(14):1332-9.)	Do NOT open capsules (bitter taste; stability uncertain).
lamivudine (3TC)	yes	10 mg/mL oral solution; 240 mL bottle. Pale yellow, strawberry-banana flavoured solution); (NB: contains 6% v/v ETOH & 3g sugar). Does not contain sorbitol.	Store at room temperature.		Can also crush or split tablets. Pharmacokinetic study in adults on co-administration of 3TC 300 mg and sorbitol solution (low (3.2 g), medium (10.2 g) and high (13.4 g) sorbitol doses) given with 240 mL water in the fasting state. A dose-dependent decrease in 3TC exposure was seen and is likely due to decreased absorption and bioavailability of 3TC (accelerated small intestinal transit time mediated by sorbitol). Higher doses of sorbitol resulted in lower 3TC concentrations (decreased $AUC_{0-\infty}$ by 14%, 32%, and 36%,

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
					respectively). Caution is warranted with chronic administration of 3TC solution and other liquid drugs containing sorbitol (e.g. abacavir, nevirapine, cotrimoxazole). (Adkison et al. Clin Pharmacol Ther 2018;103:402-8.) In addition, in pediatric patients, ensure lamivudine dose is optimized based on weight.(Choi et al. Clin Pharmacol Ther 2018;104:785-7.)
lopinavir/ ritonavir	yes	80 mg/20 mg per mL; 160 mL bottle. Cotton-candy flavoured yellow-orange oral solution. Oral solution contains the excipients alcohol (42.4% v/v) and propylene glycol (15.3% w/v). Increased risk of toxicity in preterm infants.	Stable in refrigerator until expiry date; stable at room temperature for 42 days.	Administration of crushed 200/50 mg lopinavir/ritonavir tablets to children significantly reduced lopinavir and ritonavir exposure with a decrease in AUC by 45% and 47%, respectively. Therefore, the use of crushed lopinavir/ritonavir tablets should be avoided, if possible. (Best et al. JAIDS 2011;58:385-91.)	Adult and pediatric Kaletra® tablets should be swallowed whole and not chewed, broken, or crushed. Risk of tablets shattering if broken/crushed. Tablets are film-coated and formulated using Meltrex (Melt Extrusion Technology) which improves the poor solubility of lopinavir/ritonavir by dissolving drug in a polymer and allowing drug to remain in dispersion as the polymer hardens. The extruded material can then be processed into tablets. (Klein et al. JAIDS 2007;44:401-410.)
maraviroc	yes (US only); not available via SAP in Canada	20 mg/ml clear, colourless, strawberry flavoured oral solution		Successful administration of abacavir, lamivudine, raltegravir and maraviroc given via a percutaneous endoscopic gastrostomy (PEG) tube in a 40 year old male with HIV and	Film coated immediate release tablet. No pharmacokinetic data available for crushing/chewing tablet. (Data on File, Pfizer). While the company does not have any specific kinetic

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
				PML. Abacavir and lamivudine oral liquid formulations were used. Maraviroc and raltegravir tablets were each crushed and separately mixed with 60 mL sterile water. Maraviroc trough concentrations were therapeutic. (Pecora Fulco P et al. AJHP 2019;76:265-7.)	information, crushing or cutting the tablets is not expected to negatively affect bioavailability.
nelfinavir	no; discontinued in Canada & US	Discontinued in Canada & US; 50 mg/g oral powder; 144 g bottle. (1g = 1 level scoop)	**Oral Powder:** mix with small amount of water, milk, formula, or dietary supplements (acidic food or juice such as apple juice, orange juice, apple sauce not recommended-bitter taste); consume immediately; may be stored in fridge for up to 6 hours.		For infants, can also dissolve tablets (i.e. 250 mg tablet) in 5 mL sterile water to yield a 50 mg/mL liquid. Use syringe with 1 mL increments to measure. Round dose to nearest 50 mg and consume immediately. Tablets also readily dissolve in water and produce a dispersion that can be mixed with milk/chocolate milk. Tablets can be crushed and given with pudding. Tablet may be mixed with food or liquid and taken immediately. Do not mix with acidic food/juice (orange or apple juice) due to bitter taste.
nevirapine	Yes (SAP)	10 mg/mL;240 mL bottle suspension Contains sorbitol 162 mg/mL (Personal communication, Boehringer Ingelheim Canada, April	Stable at room temperature. Shake well before use.	Recipe for nevirapine 5 mg/mL oral suspension: 1) Crush nevirapine 200 mg tablet; 2) Measure out 40 mL of simple syrup in a bottle; 3) Triturate crushed tablet with 10 mL of simple syrup; 4) Add nevirapine mixture back into the bottle of remaining syrup and shake well.	Can crush immediate release (200 mg) nevirapine tablets in water. NB: Extended-release (400 mg XR) tablets must be swallowed whole; they must not be chewed, crushed or divided. (Viramune Product Monograph, Burlington, ON, Nov 18, 2013).

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
		2017).		Refrigerate. Stable for 24 hours. (Data on file, Foothills Hospital Pharmacy Department, Alberta Health Services, Calgary, Alberta, Canada)	
raltegravir	Yes-SAP in Canada; available in US	20 mg/mL oral banana flavoured granular powder (single-use packet of 100 mg raltegravir) 25 mg & 100 mg pediatric chewable tablets (Canada & US)	The oral suspension should be administered orally within 30 minutes of mixing	An HIV-positive patient on continuous venovenous hemodiafiltration (CVVHDF) received raltegravir 400 mg BID, darunavir 600/100 mg BID, zidovudine 300 mg BID and 3TC 50 mg q24h in suspension via gastric port and simultaneous enteral feeding via the duodenal port of a double-lumen nasogastroduodenal tube. Pharmacokinetic sampling and analysis indicated that darunavir and raltegravir were removed by CVVHDF with approximately the same clearance as provided by a normally functioning kidney. **Absorption of both darunavir and raltegravir after suspension and application via the gastric port with continued administration of feed via the duodenal port of the double-lumen tube was good.** As such, dose adjustments are not required for patients receiving darunavir and/or raltegravir while undergoing CVVHDF and that	Crushing 400 mg and 600 mg HD film coated tablets is not recommended. (Data on file, Merck US, November 29, 2017). Granules (sub-units of the tablet) dissolve faster than intact tablets and may result in faster release of drug which could affect in-vivo performance. (Data on file, Merck Frosst, May 2008). Drug has a bitter taste which is masked by the film coating. Chewable tablets may be chewed or swallowed whole. Oral suspension, chewable tablets and film-coated tablets are NOT interchangeable. The maximum dose of the chewable tablets is 300 mg BID and the maximum dose of the oral suspension is 100 mg BID.

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
				absorption of darunavir and raltegravir is not significantly affected by postpyloric enteral feeding. (Taegtmeyer et al. AIDS 2011;25(10):1339-41.)	
rilpivirine	Dispersible tablet (2.5 mg) and granule (2.5 mg/g) formulations under investigation- Janssen Ireland (https://clinicaltrials.gov/ct2/show/NCT02561936) (not available in Canada)			Case report of a critically ill patient with lymphoma requiring enteral administration of ARVs. Both abacavir and 3TC solutions were administered enterally. Crushed dolutegravir 50 mg BID (separated from enteral nutrition by 2 hours) and rilpivirine 25 mg daily (given with a 240-mL bolus of an enteral formula (2 kcal/mL)) were administered via orogastric tube. Crushed tablets were each mixed with 10 mL of water and flushed down the tube at separate administration times. Trough concentrations were: day 8, rilpivirine 30 ng/mL (reference range 40-120 ng/mL), and day 9, dolutegravir 820 ng/mL (reference range 830 ng/mL steady-state trough concentration for 50 mg once daily dose). Virologic suppression was maintained after ARV enteral administration (hospital day 29). Given somewhat decreased levels of	Film coated tablet. No data available on stability of splitting or crushing rilpivirine tablets. Rilpivirine is insoluble in water over wide pH range. (Email communication, Janssen, July 2012). Crushed tablets added to a small amount of semisolid food or liquid is not expected to have an adverse effect if consumed immediately. Since tablets are small, ensure the whole dose is consumed. (Huesgen et al. Pharmacother 2016;36(11):1145-65.)

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
				these ARVs, the authors recommended consideration to increase dolutegravir to 150-200 mg total daily dose, particularly in integrase-experienced patients, and rilpivirine 50 mg daily (similar to dosing with an inducer such as rifabutin). (Turley et al. JIAPAC 2017;16(2):117-119.)	
ritonavir	No- solution discontinued in Canada (May 2019) Ritonavir solution and powder not available via SAP. (AbbVie Canada, Data on file, May 2019); ritonavir liquid and powder available in US	Not available in Canada. Available in US. 80 mg/mL oral liquid; 240 mL bottle. Orange-coloured oral solution, peppermint & caramel-flavoured. 43.2% v/v alcohol, propylene glycol 26.57% w/v. Shake well before each use. 100 mg Oral Powder (100 mg/packet)- Available in the US only.	Stable at room temperature; do not refrigerate. See tips for taking liquid in Crushing & Splitting section.		Tablets should be swallowed whole and not chewed, broken or crushed. Tablets are film-coated. (Norvir® Product Monograph, AbbVie Corp, St-Laurent, QC, Sept 25, 2018). Norvir® tablets are formulated using Meltrex (Melt Extrusion Technology) which improves the stability in heat and poor solubility of ritonavir by dissolving drug in a polymer and allowing drug to remain in dispersion as the polymer hardens. The extruded material can then be processed into tablets. (Klein et al. JAIDS 2007;44:401-410); (FDA Norvir Tablets Drug Approval Package, October 21, 2013). A manufacturer study demonstrated that a ritonavir 100 mg tablet produced an aqueous suspension when

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
					placed whole in 10, 20, 40 or 60 mL of water at room temperature for 4 hours without stirring. The tablet fully eroded after 4 hours. Hourly stirring or placing the tablet in warm water did not impact the dissolution time. This dosing strategy is only recommended for full doses in 100 mg increments. In vivo, the performance and bioavailability of the suspension is expected to be the same as ritonavir powder for oral suspension. Of note, ritonavir powder for oral suspension demonstrated bioequivalence to the discontinued oral solution when administered after a moderate-fat meal (Data on file, AbbVie Canada, Feb 20, 2017 and April 17, 2017). Liquid is unpalatable, bad aftertaste. Tips: - Mix oral solution with milk/chocolate milk or pudding - Give after popsicle/frozen juice to dull taste buds - Give after grape jelly, maple syrup, or peanut butter which coats mouth -Give strong flavour after dose: syrup, cheese, chewing gum Oral powder (100 mg/packet):

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
saquinavir	no		Liquid not being formulated due to unpalatability	In an open-label, randomized, 4 period study in adults, the bioavailability of 1000 mg opened saquinavir capsules suspended in simple syrup, baby formula and jelly jam (plus ritonavir 100 mg oral solution) was approximately 10%, 60% and 40% higher, respectively, than 1000 mg unopened saquinavir capsules plus ritonavir. In terms of palatability, saquinavir suspended in simple syrup or jelly jam ranked higher than saquinavir suspended in baby food.(McKay et al. 8th Int Workshop Clin Pharm HIV Ther 2007, abstract 6.)	The entire packet should be mixed with soft food such as apple sauce or vanilla pudding, or mixed with liquid such as water, chocolate milk, or infant formula. All soft food or liquid should be consumed within 2 hours of preparation. The bitter taste may be decreased if taken with food. The powder should be used in 100 mg increments only. The oral powder can also be administered via feeding tube after being mixed with water. **Note: Only Invirase® 500 mg tablets are marketed in Canada & US.** Invirase® and Fortovase® capsules are discontinued (verified April 2019). Hard gel caps (Invirase®) may be opened and powder sprinkled on food, simple syrup or water (unpleasant taste). Take with food. 6 x 200 mg Fortovase® (soft-gel caps) whole caps mixed with 50 mL of whole milk or Advera nutritional supplement took 5-15 minutes to dissolve when heated to 40, 60 or 80°C.

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
				An extemporaneously prepared saquinavir suspension (60 mg/mL) from saquinavir (Fortovase®) soft gelatin capsules was stable at both 5 and 25 degrees C for at least 30 days.(Tan et al. J Clin Pharm Ther 2000;28:457-63.)	The mixture remained in solution for up to 1 hour at room temperature. If refrigerated for 24 hours, it turned into a gel, but reliquified after reheating to 30 degrees C. The drug was still stable at 24 hours. (data on file, Hoffmann-LaRoche)
stavudine (d4T)	No longer available in Canada via SAP; product discontinued in Canada (Sept 30, 2018) & US	1 mg/mL oral suspension; 200 ml bottle. Fruit-flavoured. Shake well (no longer available in Canada & US)	Stable 30 days in fridge.		Can open up stavudine capsules and give in small portion of food or 5-10 mL cool tap water.
tenofovir DF	yes (US only)	40 mg per 1 gram of oral powder formulation. Oral powder should be mixed in a container with 2 to 4 ounces (60 to 120 mL) of soft food not requiring chewing (e.g., applesauce, baby food, yogurt). Do not attempt to mix in a liquid as the powder may float on top even after stirring.	Administer immediately to avoid a bitter taste.		Crushed tablet dissolves in 100 mL water in 20 minutes; grape juice may also be used. Consume immediately. NB: crushed tablets have very disagreeable taste. May also try splitting tablets and inserting into empty gelatin capsules to mask taste.

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
tipranavir	yes (US only; not in Canada)	100 mg/mL oral solution; contains 116 IU/mL vitamin E.	Store oral solution and room temperature (25ºC). Use solution within 60 days of opening the bottle.		250 mg capsule. Avoid splitting or crushing capsule.
zalcitabine (ddC)	no	Investigational oral solution is no longer available.			
zidovudine (AZT)	yes	10 mg/mL oral syrup; 240 mL bottle. Strawberry-flavoured.	Store at room temperature.		May open capsules & give in small portion of food or 5-10 mL cool tap water.
DIRECT ACTING ANTIVIRALS (DAAs)					
daclatasvir	no				Manufacturer recommends not chewing or crushing the tablet as it has a very unpleasant taste. (Daklinza Summary of Product Characteristics, EU, Feb 19, 2019).
elbasvir/ grazoprevir (Zepatier®)	no			Sustained virologic response was achieved in a 63 year old non-cirrhotic, treatment-naive man who received 16 weeks of crushed elbasvir/grazoprevir administered through a percutaneous endoscopic gastrostomy (PEG) tube. (Yap et al. J Clin Pharm Ther	The Zepatier® tablet uses an enabled formulation (i.e. not a simple enteric coated formulation) and breakage of the tablet integrity should not affect its bioavailability in presence of increased pH. (Reau et al. Hepatol Commun 2017;1(8):757-764.) However,

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
				2018;43:730-2.)	no pharmacokinetic studies have been performed, and caution is advised.
glecaprevir/ pibrentasvir (Maviret®, Mavyret®)	no			Case report of a 41 year-old female (36 kg) with treatment naive chronic HCV, spina bifida and hydrocephalus who received crushed glecaprevir/pibrentasvir (GLE/PIB) x 8 weeks via PEG tube feeds and achieved a 12-week sustained virologic response post-treatment. Prior to PEG tube administration, GLE/PIB was crushed into powder and suspended with water. The regimen was well-tolerated (constipation reported). (Tanaka et al. Clin J Gastroenterol 2019;12:588-91.)	In healthy adults, cutting the tablet in half had minimal impact on glecaprevir and pibrentasvir exposures (≤15% difference) compared to swallowing the whole tablets. In contrast, grinding or crushing the tablets resulted in 27-61% lower exposures for glecaprevir and 21-83% higher exposures for pibrentasvir. (Oberoi et al. J Pharm Sci 2018;107:1724-30.) These data suggest that Maviret® tablets may be cut in half for people who have difficulty swallowing whole tablets. Solubility: Glecaprevir has a solubility of less than 0.1 to 0.3 mg/mL across a pH range of 2–7 at 37ºC and is practically insoluble in water, but is sparingly soluble in ethanol. (Maviret Product Monograph; St-Laurent, QC. Aug 16, 2017).
ledipasvir/ sofosbuvir (Harvoni®)	no			In a treatment-experienced patient with compensated cirrhosis, SVR12 was achieved after 24 weeks of treatment with crushed ledipasvir/sofosbuvir	

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
				administered via a percutaneous endoscopic gastrostomy (PEG) tube. (Jindracek et al. J Pharm Pract 2018;31:522-4.) A 19-year old woman with HCV genotype 1 and HIV coinfection achieved SVR12 after 12 weeks of crushed ledipasvir/sofosbuvir administered via gastrosomy button. Each ledipasvir/sofosbuvir tablet was crushed, mixed with 10 mL warm water, and administered via syringe. Additional warm water was used to obtain all the powder. (Fulco et al. AJHP 2017;74:1761-2.)	
sofosbuvir	no				Sofosbuvir tablets can be disintegrated in water, juice, or milk with minor stirring and pressure with a spoon. However, the stability of sofosbuvir in these liquids is unknown at this time. Furthermore, there are no studies evaluating the pharmacokinetic parameters of the disintegrated or crushed sofosbuvir tablet versus the whole tablet. In addition, a disintegrated or crushed sofosbuvir tablet may have an unpleasant taste. (Personal

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
					communication, Gilead Sciences Canada, December 2013).
sofosbuvir/ velpatasvir (Epclusa®)	no			A case report of a 70 year-old female with an oropharyngectomy and infected with HCV genotype 1b studied drug absorption and exposure of crushed sofosbuvir/velpatasvir 400/100 mg tablet daily taken with an acidic beverage and meal (to enhance absorption). TDM showed that the absorption of both was increased compared to historical controls and the steady-state trough concentration (Css) of velpatasvir at weeks 1 and 10 showed concentrations above historical reference with no accumulation. Although the trough Css of sofosbuvir was below the limit of quantification, this was anticipated/expected given the short half-life of the drug (30 minutes). The drugs were well tolerated with improved liver parameters. The HCV viral load became undetectable after 4 weeks and remained undetectable 12 weeks post-treatment. (Lalanne et al. Ther Drug Monit 2019 Dec 4 [Epub ahead of print].)	Manufacturer states that Epclusa® tablets are not enteric-coated and are not sustained-release. Tablets can be disintegrated in water, juice, or milk with minor stirring and pressure with a spoon. A disintegrated, crushed or split tablet may have an unpleasant taste and there are no studies evaluating the pharmacokinetics of these methods of administration. (Gilead Sciences Canada, Data on File, Nov 11, 2018).

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
sofosbuvir/ velpatasvir/ voxilaprevir (Vosevi®)	no			A case report of a 62 year-old female with dysphagia and hepatitis C genotype 4 infection (non-cirrhotic, treatment naïve) was treated with sofosbuvir/velpatasvir given mixed with soft foods (e.g. apple sauce). The patient was adherent to therapy and it was well tolerated. The viral load was undetectable at weeks 4 and 12 of therapy and a 12 week sustained virologic response was achieved. Liver transaminases normalized during treatment. Mogul et al. Am J Health Syst Pharm 2020 Jan 13 [Epub ahead of print.]	Manufacturer states that Vosevi® tablets are not enteric-coated and are not sustained-release. Tablets can be disintegrated in water, juice, or milk with minor stirring and pressure with a spoon. A disintegrated, crushed or split tablet may have an unpleasant taste and there are no studies evaluating the pharmacokinetics of these methods of administration. (Gilead Sciences Canada, Data on File, Nov 19, 2018).

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
OTHER:					
acyclovir	yes	200 mg/5 mL;125 mL bottle. Banana-flavoured suspension.	Store between 15-25 ºC		
azithromycin	yes	pediatric oral powder/suspension 100 mg/5 mL (300 mg bottle) OR 200 mg/5 mL (600 & 900 mg bottles). Cherry-flavoured suspension.	Store reconstituted suspension between 5-30ºC. Dispose unused suspension after 10 days.		May also open capsules and mix with water (ingest immediately on empty stomach, follow with full glass of water).
clarithromycin	yes	125 mg/5 mL (55, 105, 150 mL bottles) OR 250 mg/5mL (105 mL bottles). Fruit-flavoured suspension. Shake well before use.	Store reconstituted liquid at room temperature.		
hydroxyurea	no				Can open up capsules and mix with water; take immediately. Some inert material (used as a vehicle in capsule) may not dissolve and may float on top. Do not allow powder to come in contact with skin and mucous

Drug	Oral Liquid Preparation			Case Reports/Clinical Compounding	Information on Crushing or Splitting Tablets
	Commercial Oral Liquid Available?	Formulation	Stability		
rifabutin	no				membranes. Avoid inhalation of powder when opening capsules. Can open capsules (experience in pediatrics: OK to mix with applesauce, syrup, cherry syrup); drug not soluble in water
Trimethoprim/ sulfamethox-azole (TMP/SMX)	yes	pediatric suspension TMP/SMX 40 mg/200 mg per 5 mL (= ½ SS tablet); 100 & 400 mL bottles Contains sorbitol. Example : Teva-Trimel suspension contains sorbitol 4 g/5 mL- (Personal communication, Teva Canada, April 2017)	Store at room temperature. Shake well before use.		

Key: SAP= Special Access Program, Health Protection Branch, Ottawa, Canada- phone: 613-941-2108; fax: 613-941-3194;
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